



OBESITY: A Rising Threat to Global Health

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Disclosure Statement

Speaker:

Dr. Fatima Cody Stanford

- Has no relevant financial relationships to disclose
- Will not be discussing unlabeled/unapproved use of drugs or products



Weill Cornell Medicine-Qatar

Points of Discussion

- ▶ What is obesity and what is its prevalence in the US and throughout the world?
- ▶ How did obesity become a threat to global health?
- ▶ Do I play a role in weight bias? If so, how do I make a change?
- ▶ What strategies might we take to address the obesity epidemic?

Obesity defined- the role of Body Mass Index (BMI)

Metric measurements:

$$\frac{\text{Weight (kg)}}{\text{Height (m)}^2}$$

English measurements:

$$\frac{\text{Weight (lb)} \times 703}{\text{Height (in)}^2}$$

How is Obesity defined in Children and Adolescents?

Weight Status Category	BMI Percentile Range
Underweight	< 5 th Percentile
Healthy Weight	5 th Percentile - <85 th Percentile
Overweight	85 th Percentile- <95 th Percentile
Obesity	≥ 95 th Percentile

How is Obesity defined in Adults?

Weight Status Category	Body Mass Index (BMI)
Underweight	< 18.5
Normal Weight	18.5-24.9
Overweight	25- 29.9
Class I Obesity	30-34.9
Class II Obesity	35-39.9
Class III Obesity	≥40

Body Mass Index Calculation

Metric measurements:

$$\frac{\text{Weight (kg)}}{\text{Height (m)}^2}$$

English measurements:

$$\frac{\text{Weight (lb)} \times 703}{\text{Height (in)}^2}$$

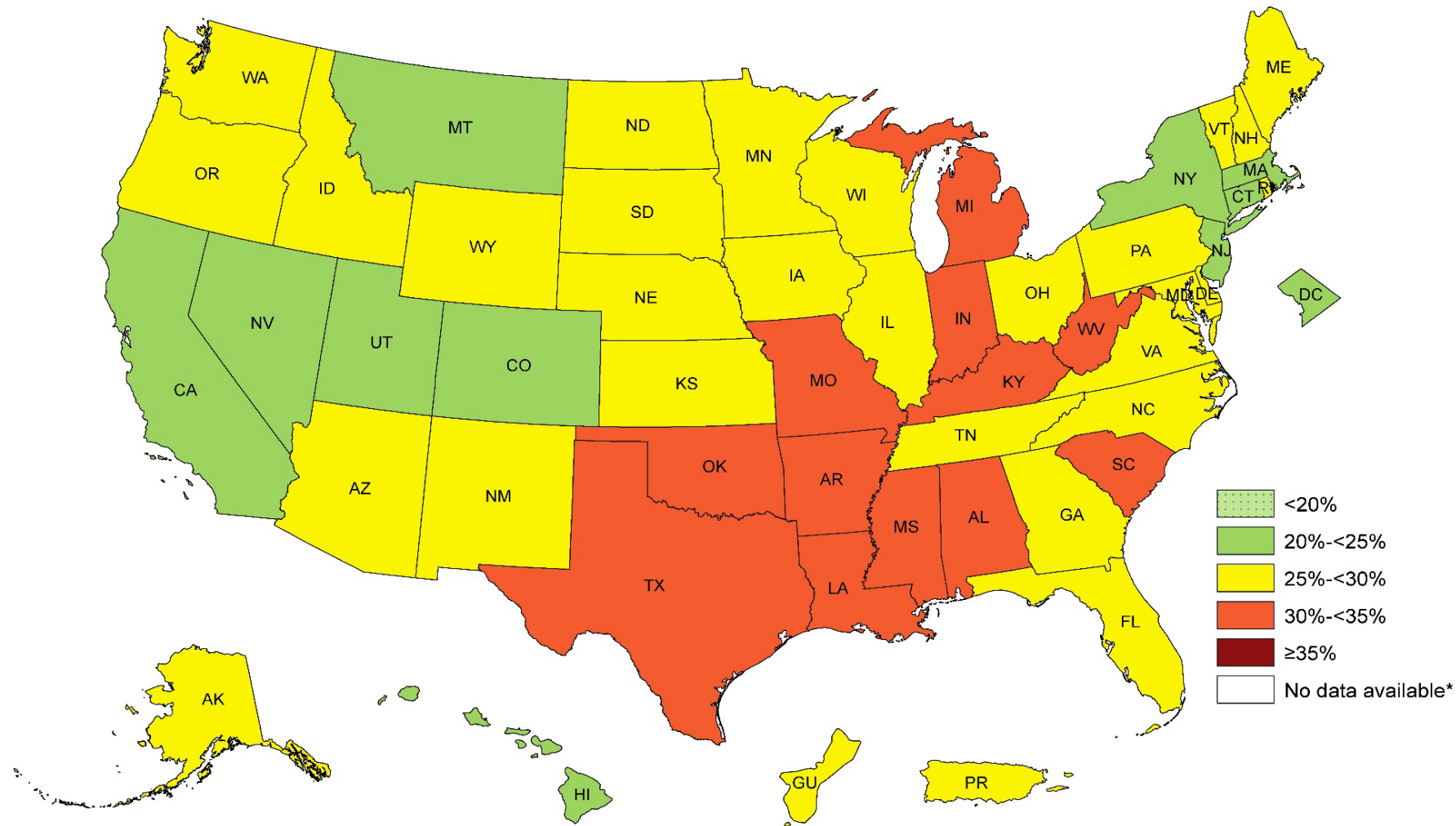
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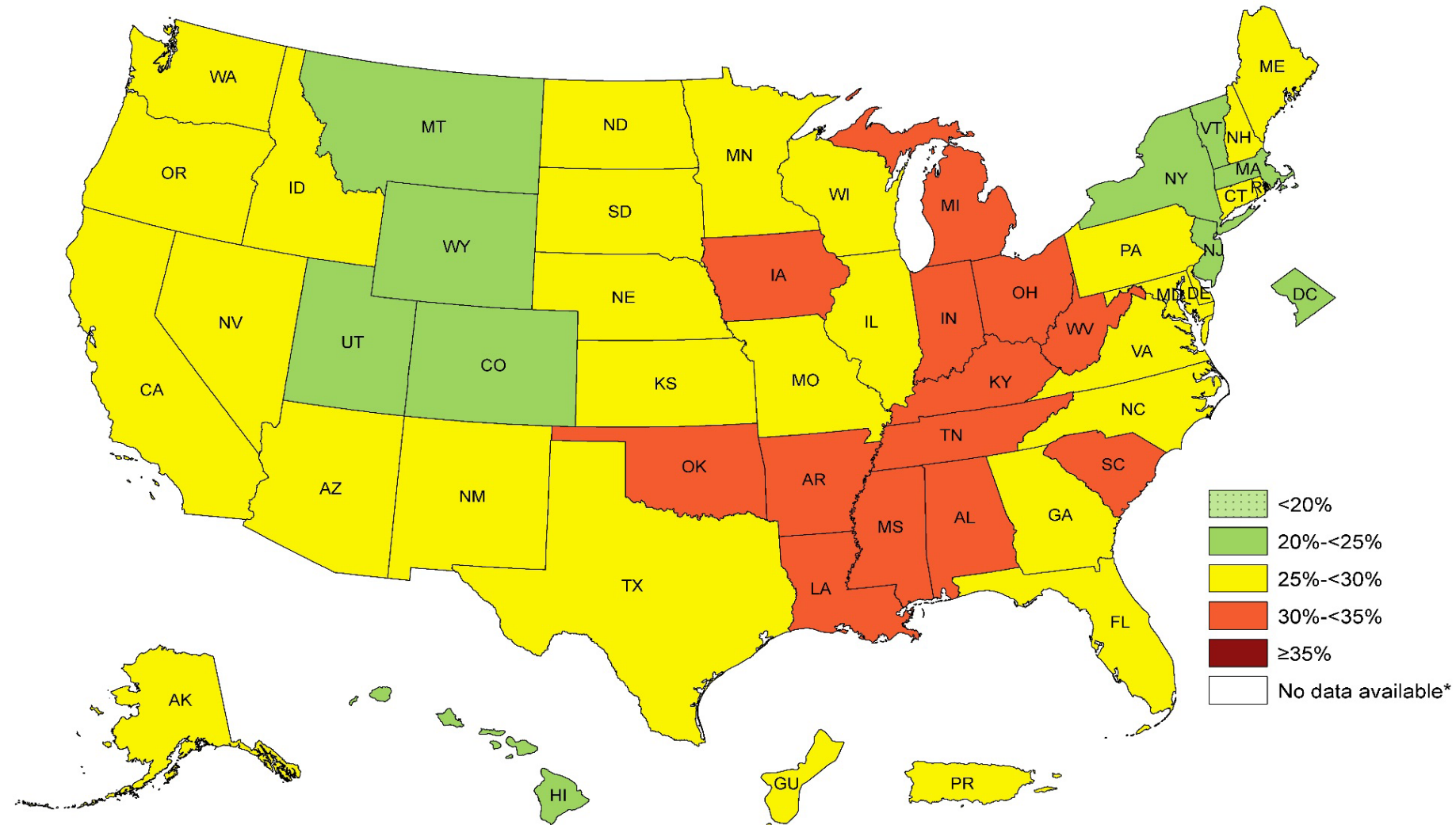
Prevalence[¶] of Self-Reported Obesity Among U.S. Adults by State and Territory, BRFSS, 2011



[¶] Prevalence estimates reflect BRFSS methodological changes started in 2011. These estimates should not be compared to prevalence estimates before 2011.

*Sample size <50 or the relative standard error (dividing the standard error by the prevalence) ≥ 30%.

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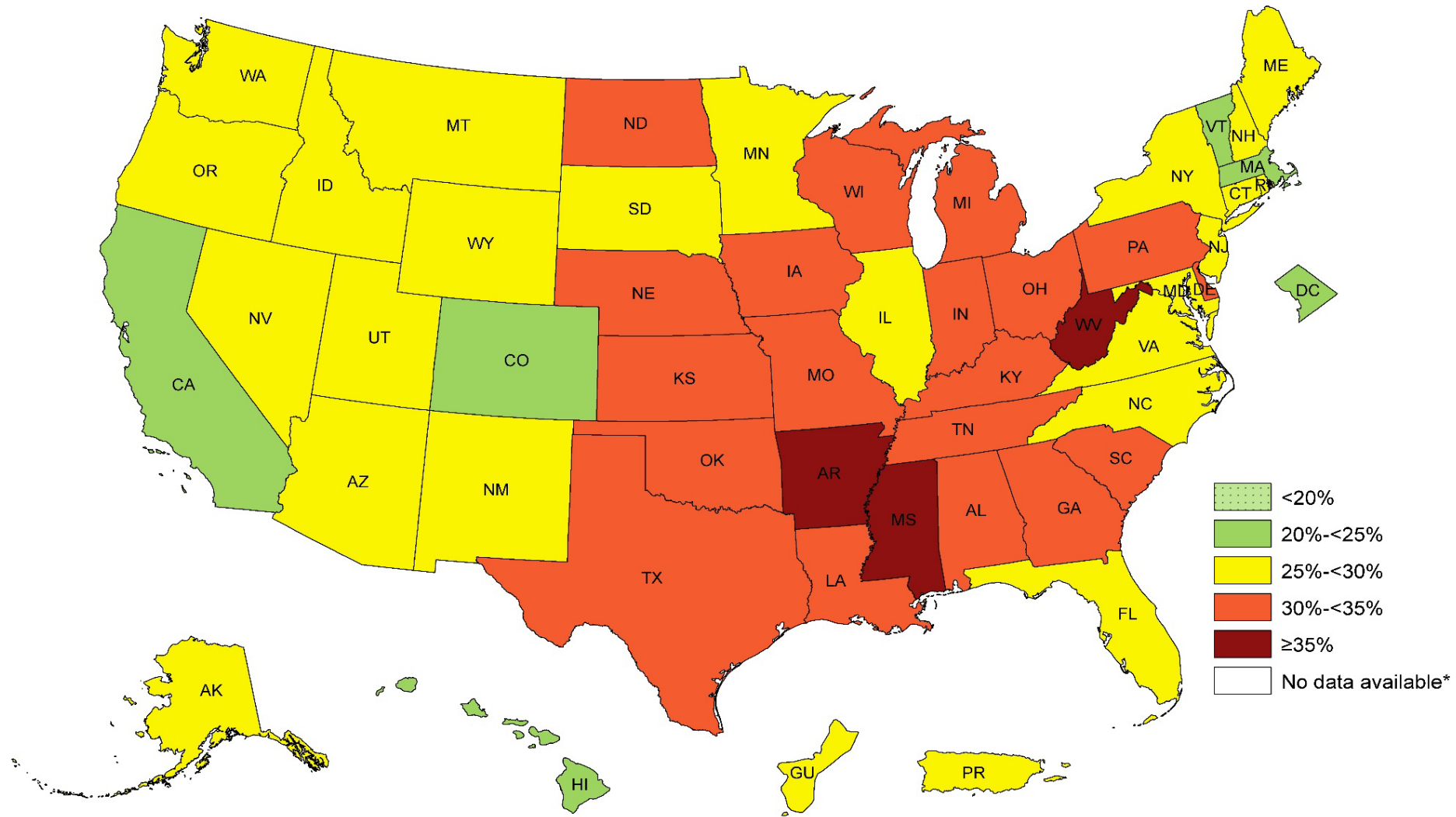
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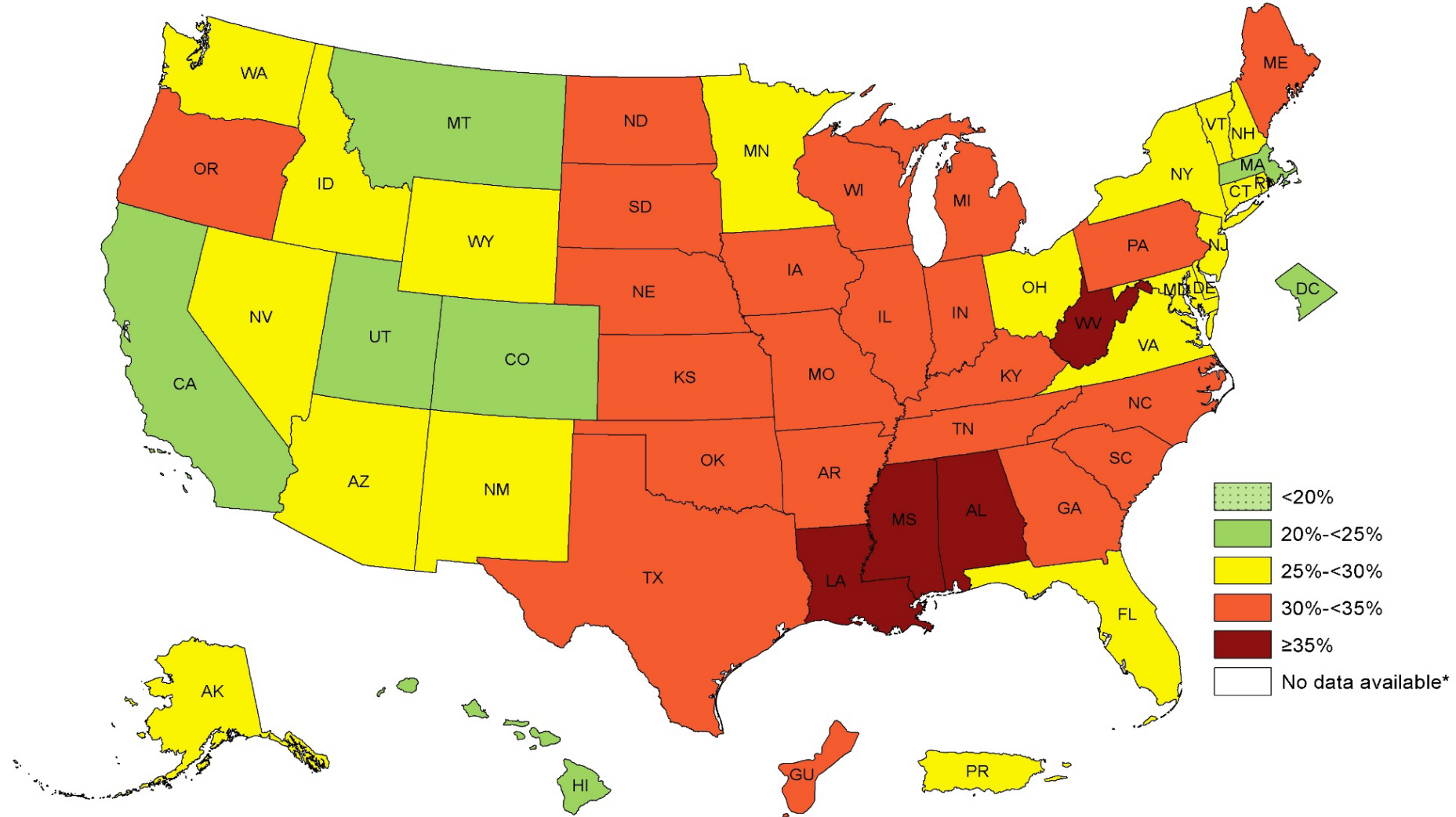
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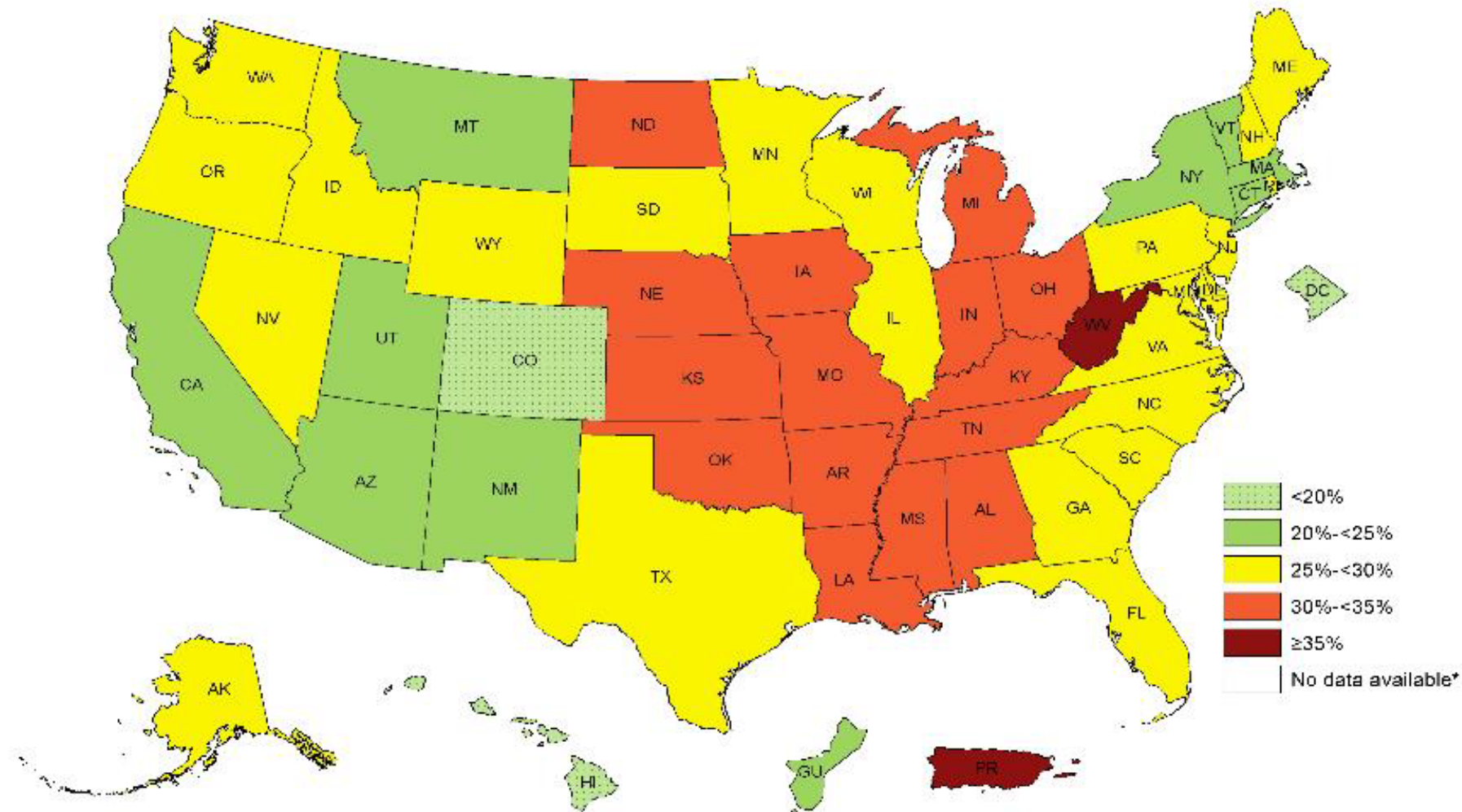
Prevalence[¶] of Self-Reported Obesity Among U.S. Adults by State and Territory, BRFSS, 2015



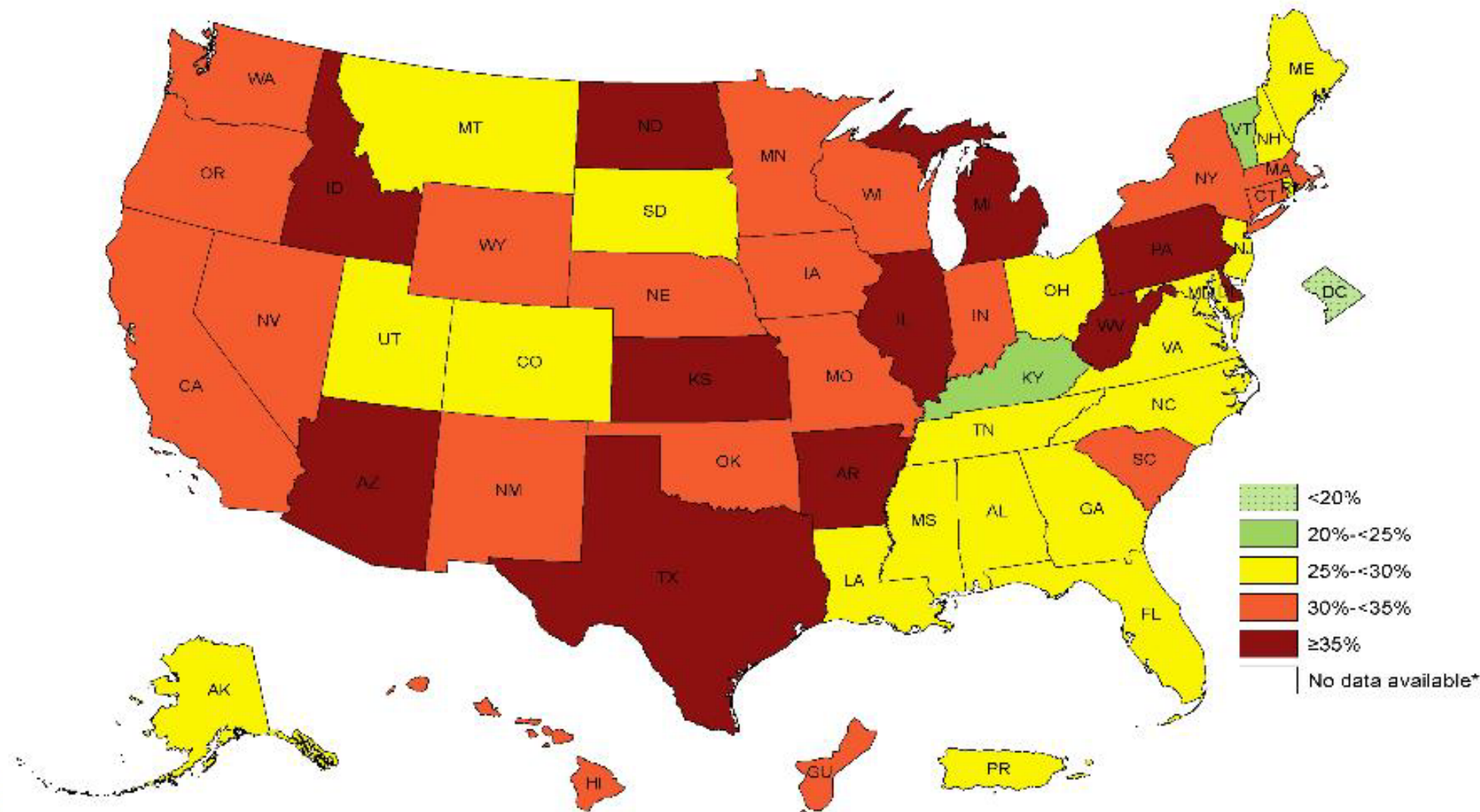
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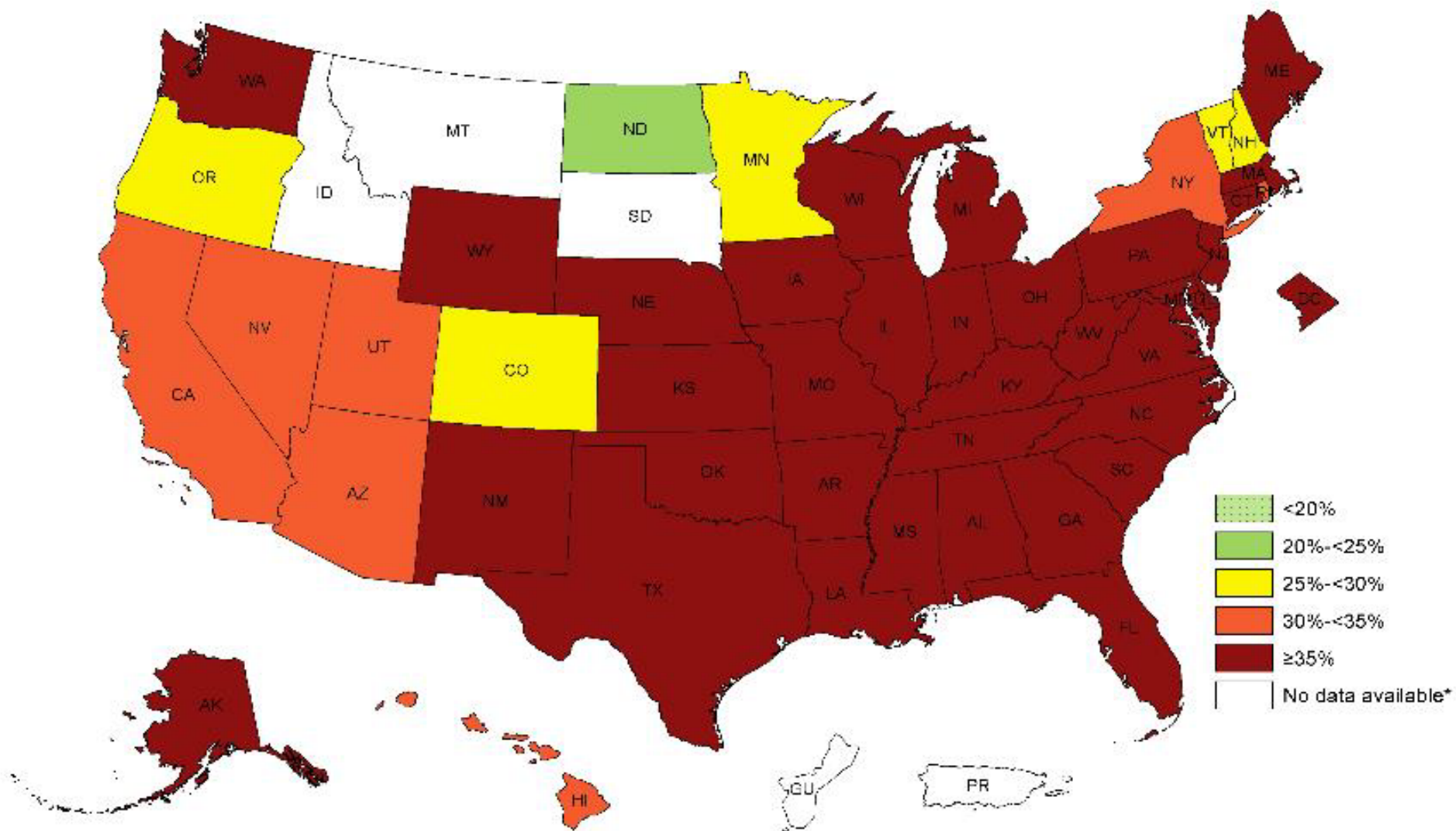
Prevalence of Self-Reported Obesity Among Non-Hispanic White Adults, by State and Territory, BRFSS, 2013-2015



Prevalence of Self-Reported Obesity Among Hispanic Adults, by State and Territory, BRFSS, 2013-2015



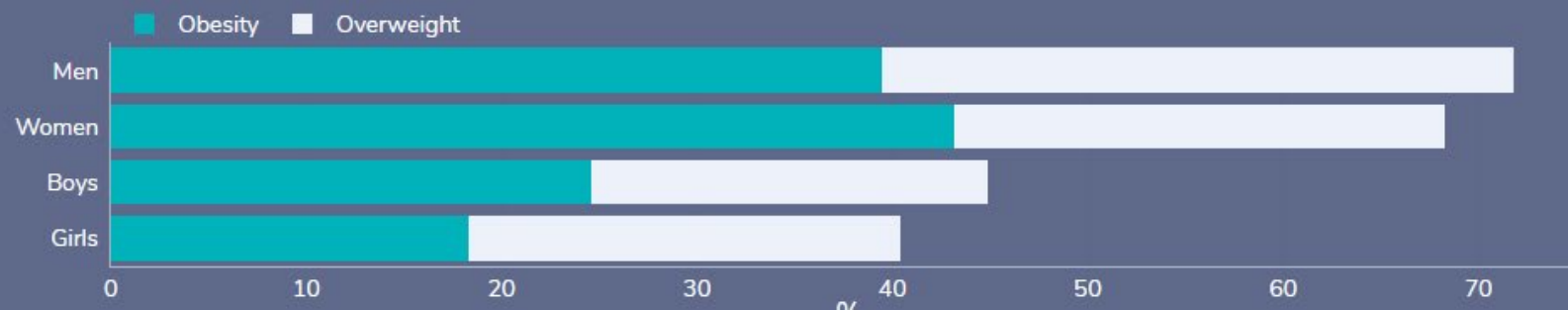
Prevalence of Self-Reported Obesity Among Non-Hispanic Black Adults, by State and Territory, BRFSS, 2013-2015



Overweight and Obesity in Qatar

Qatar

Overview | Obesity prevalence | Population breakdowns | Drivers | Comorbidities | Health systems | Actions



National obesity risk *

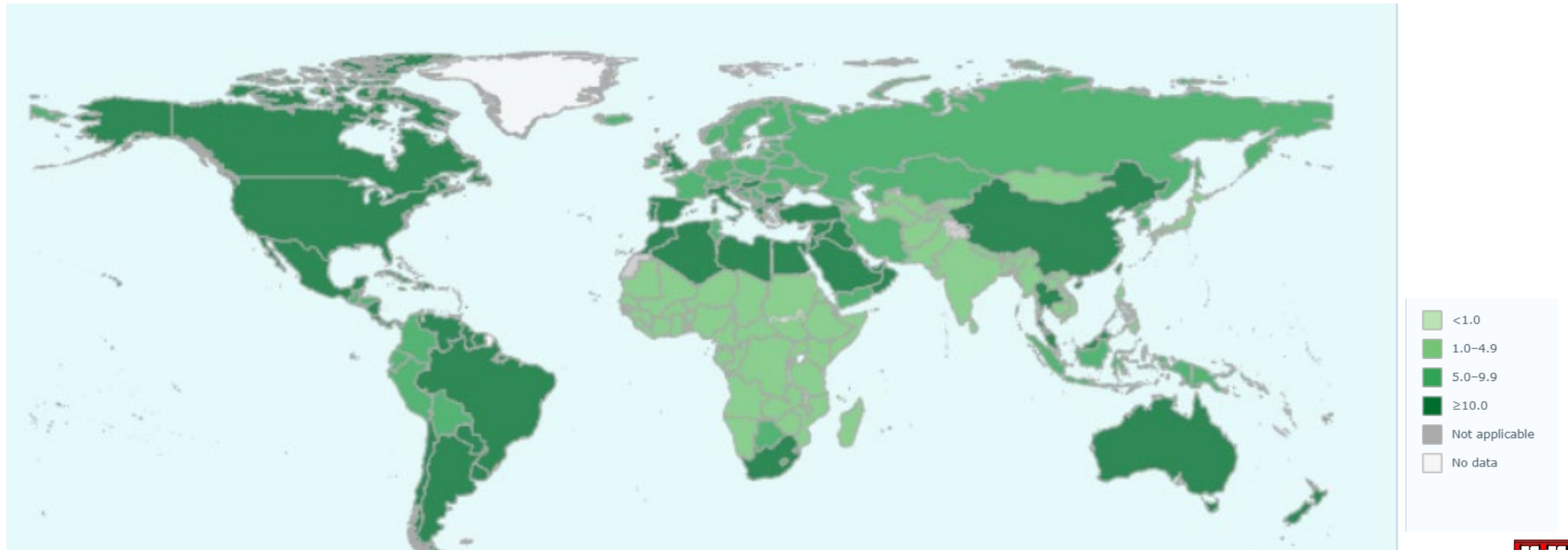
7.5/10

Childhood obesity risk *

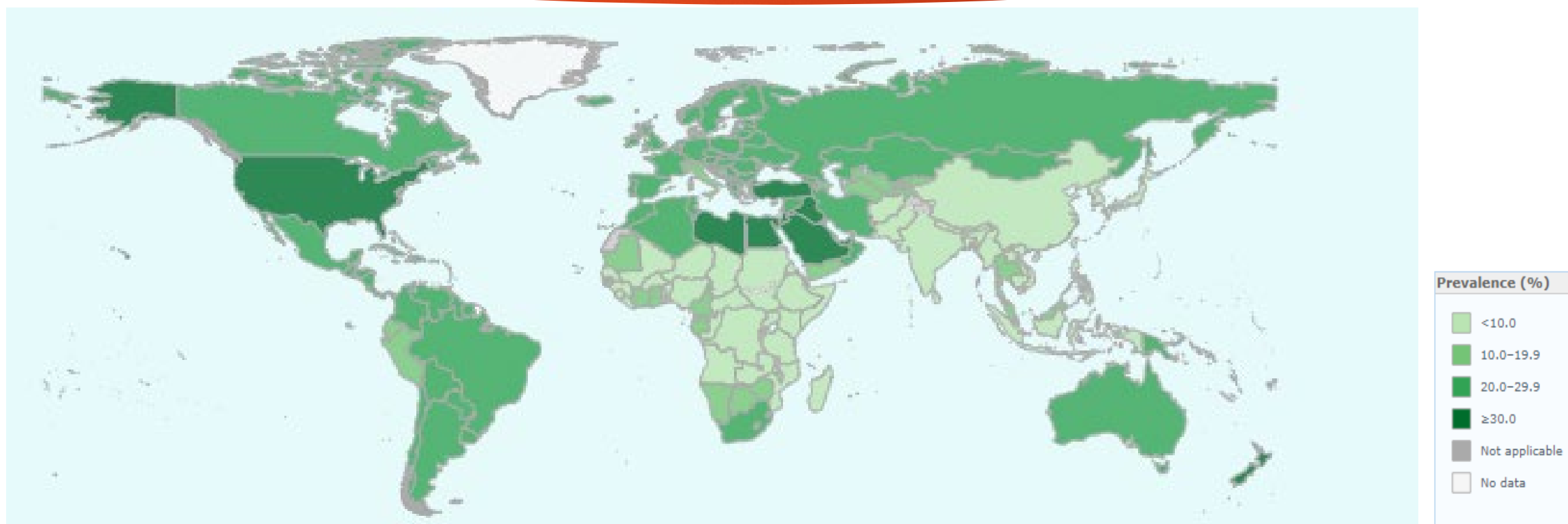
8.5/11

* Based on estimated data. For more information see Publications

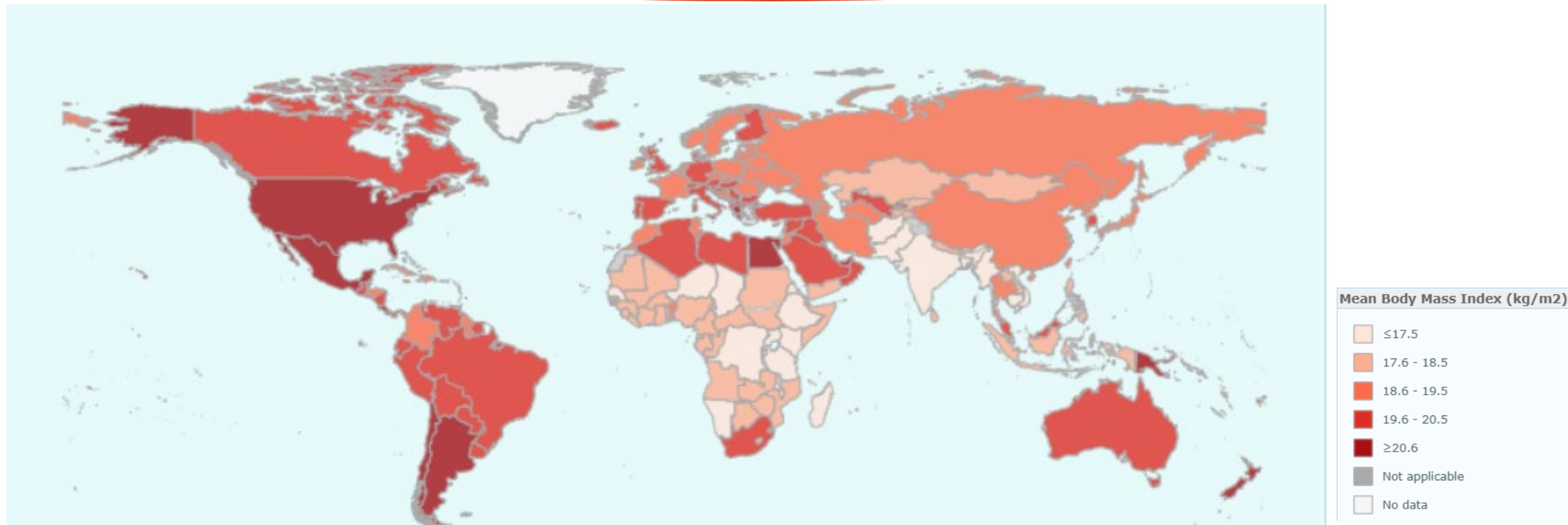
Prevalence of Obesity Among Children and Adolescents, Both Sexes, 2016



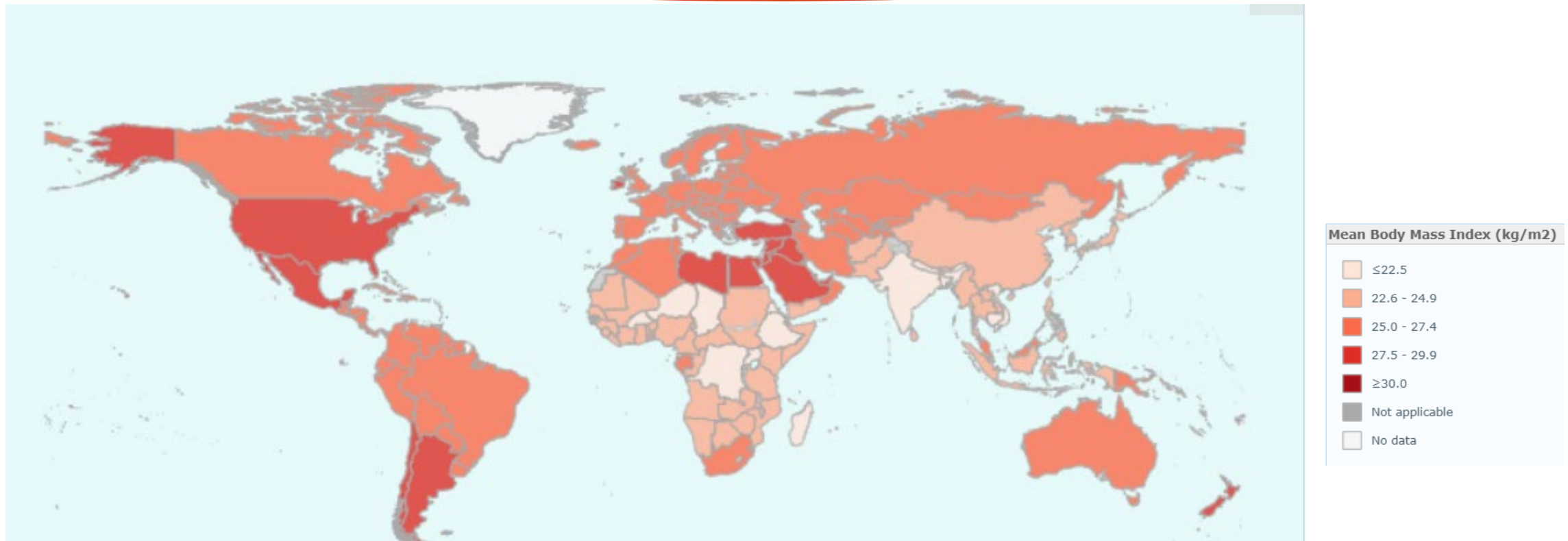
Prevalence of Obesity Among Adults, Both Sexes, 2016



Prevalence of Mean Body Mass Index Among Children and Adolescents, Both Sexes, 2016

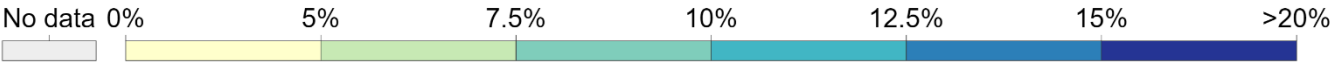
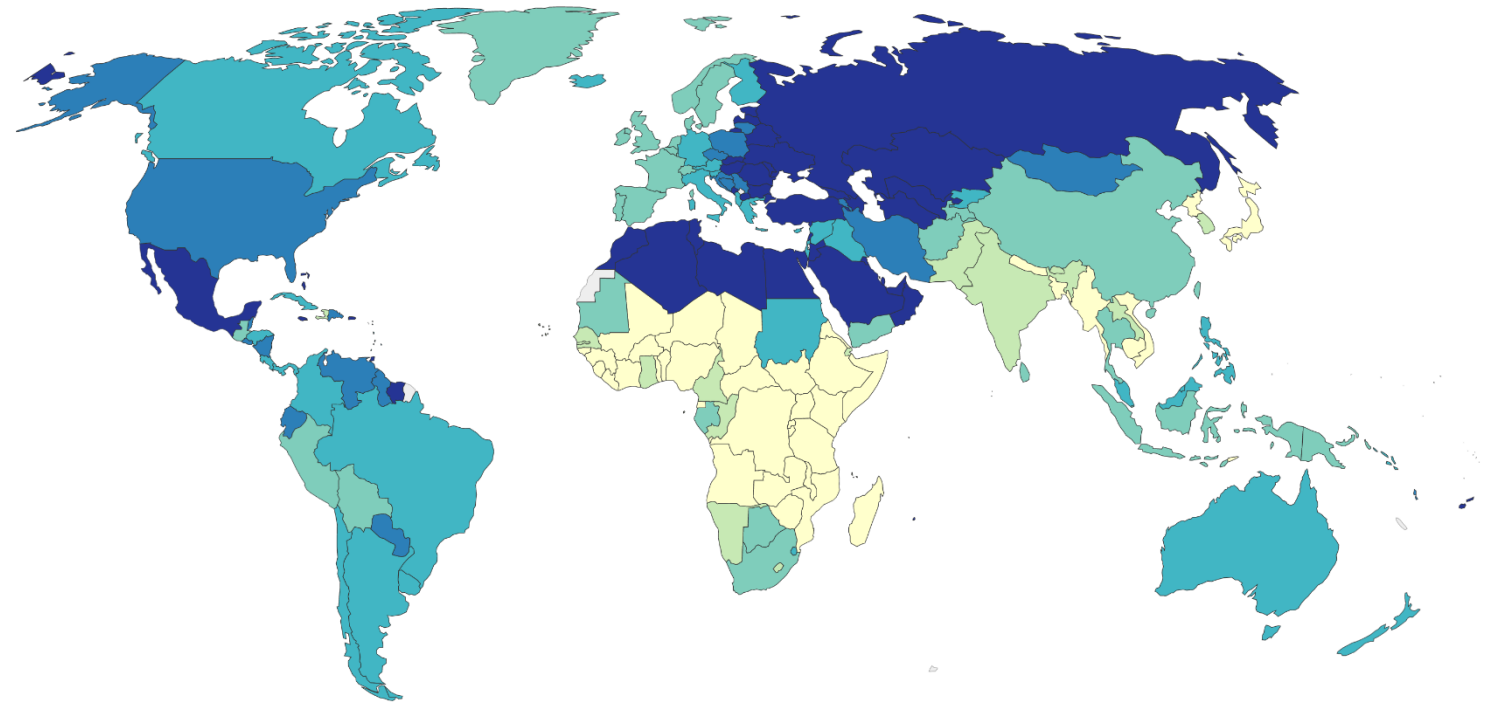


Prevalence of Mean Body Mass Index Among Adults, Both Sexes, 2016



Share of deaths attributed to obesity, 2017

Obesity is defined as having a body-mass index (BMI) equal to or greater than 30. BMI is a person's weight in kilograms divided by his or her height in metres squared.

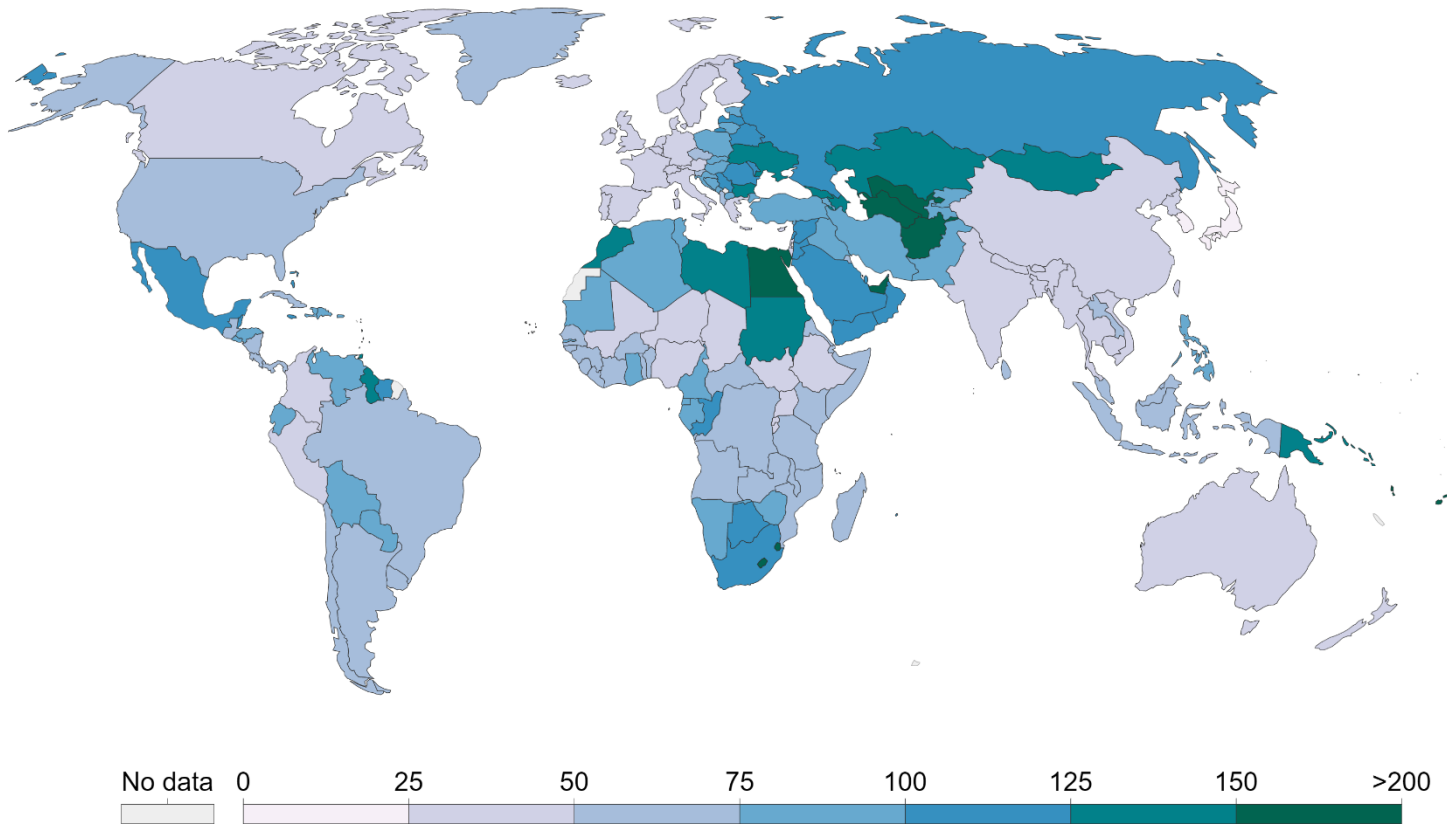


Share of Deaths Attributable to Obesity in 2017

Death rate from obesity, 2017

Premature deaths attributed to obesity per 100,000 individuals. Obesity is defined as having a body-mass index (BMI) equal to or greater than 30. BMI is a person's weight in kilograms divided by his or her height in metres squared.

Our World
in Data



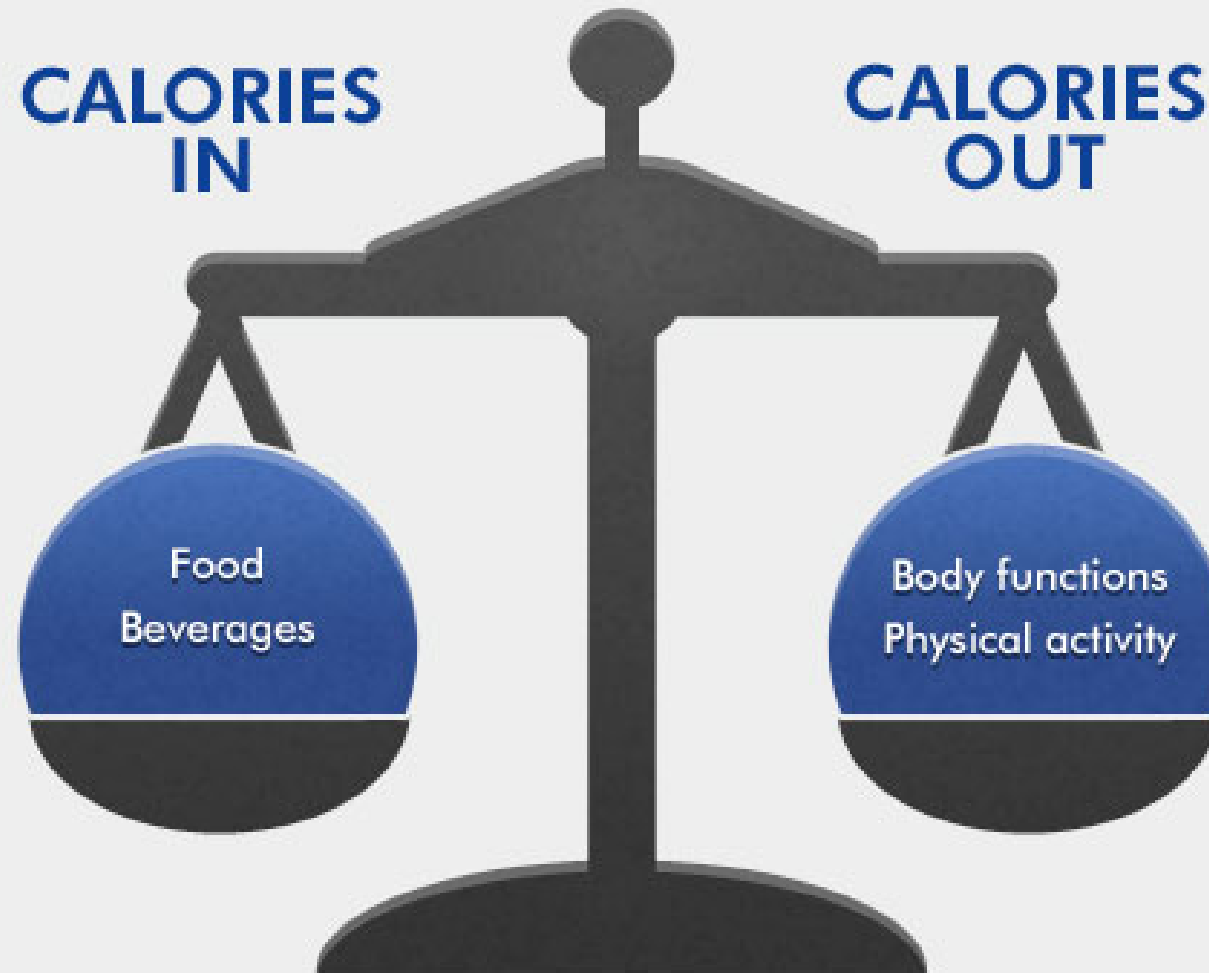
Source: IHME, Global Burden of Disease

Note: To allow comparisons between countries and over time this metric is age-standardized.

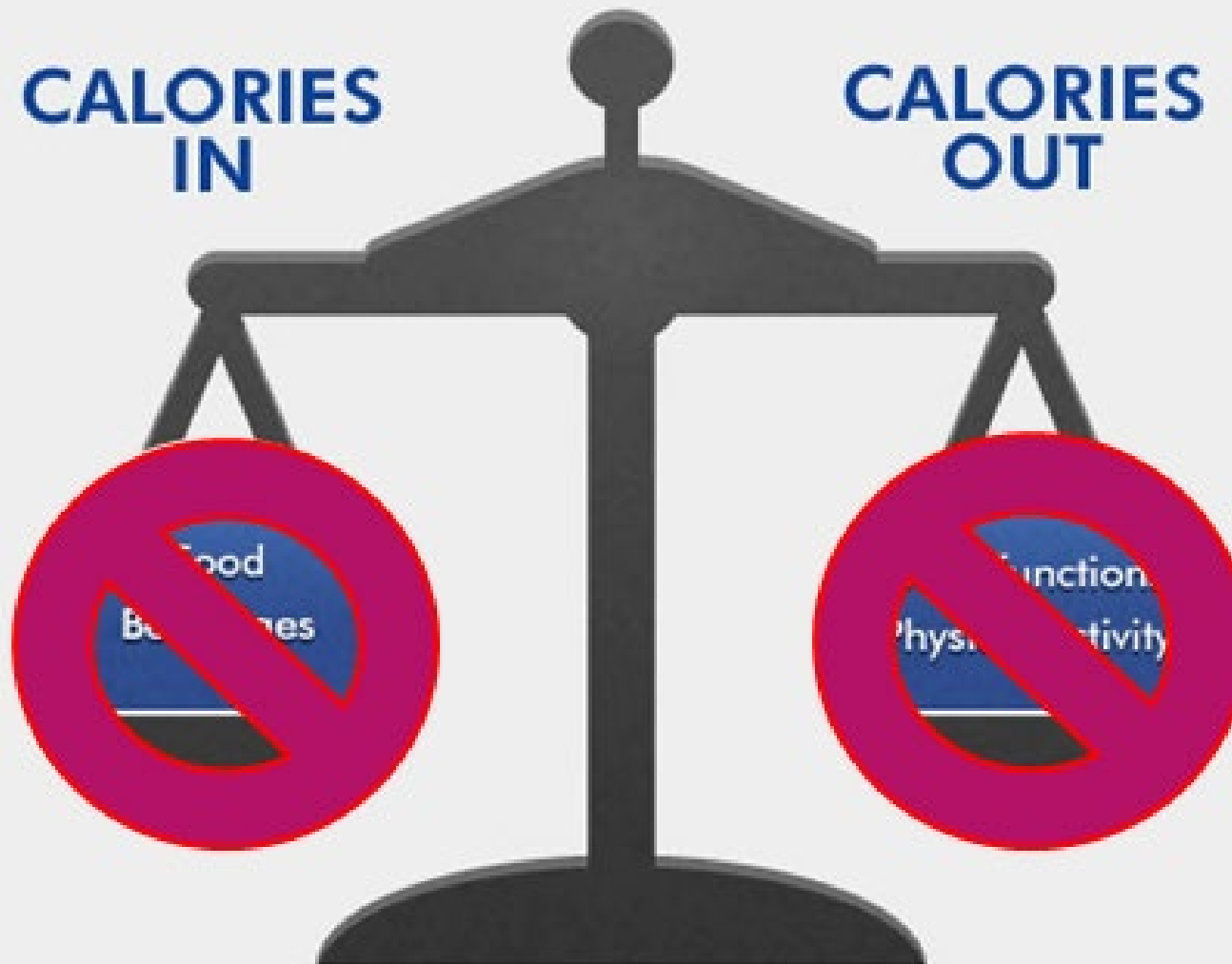
OurWorldInData.org/obesity • CC BY

Death Rate from Obesity, 2017

Energy Balance (simple)



Energy Balance (simple)



Obesity: A Multi-factorial Disorder



Genetics



Environment



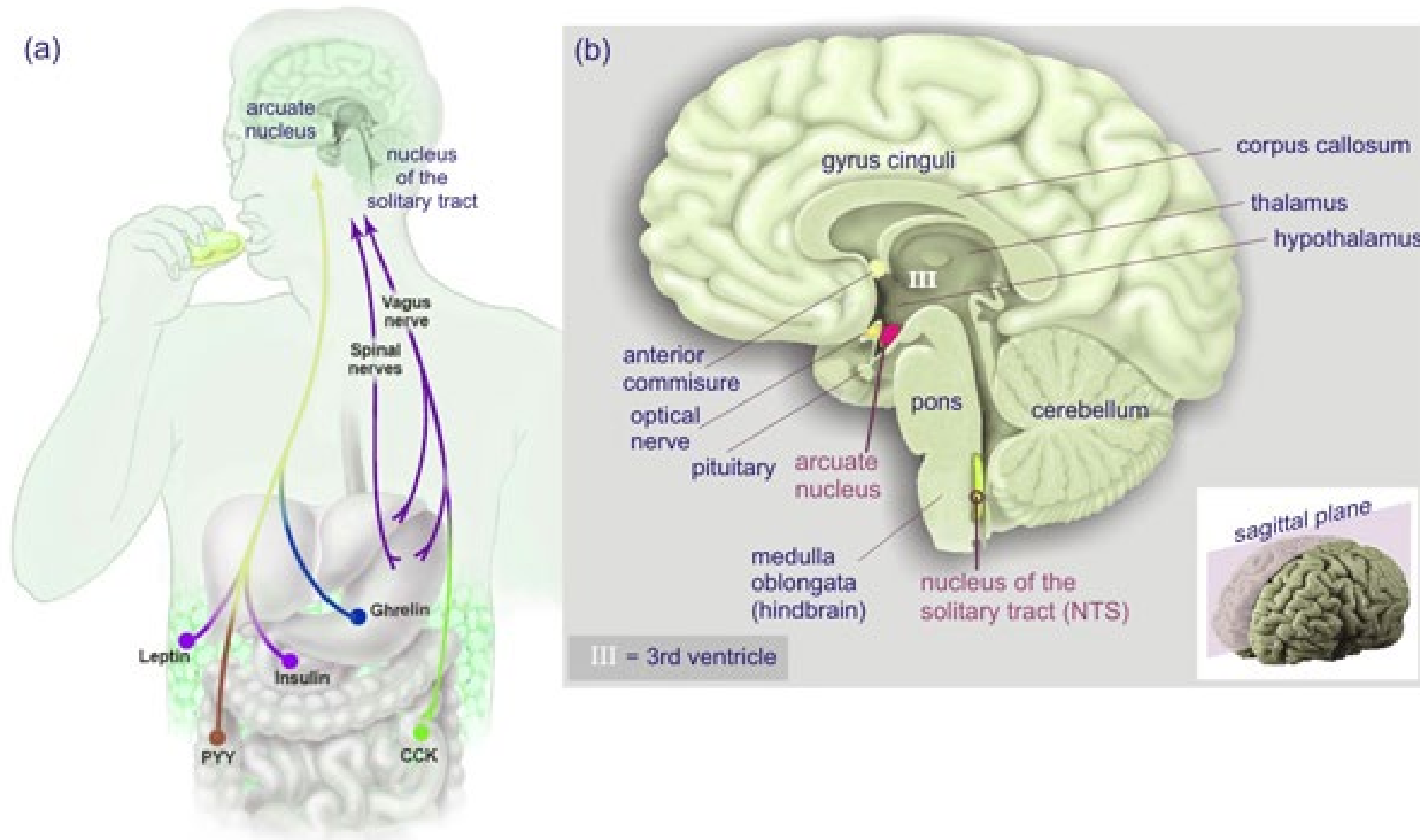
Development



Behavior



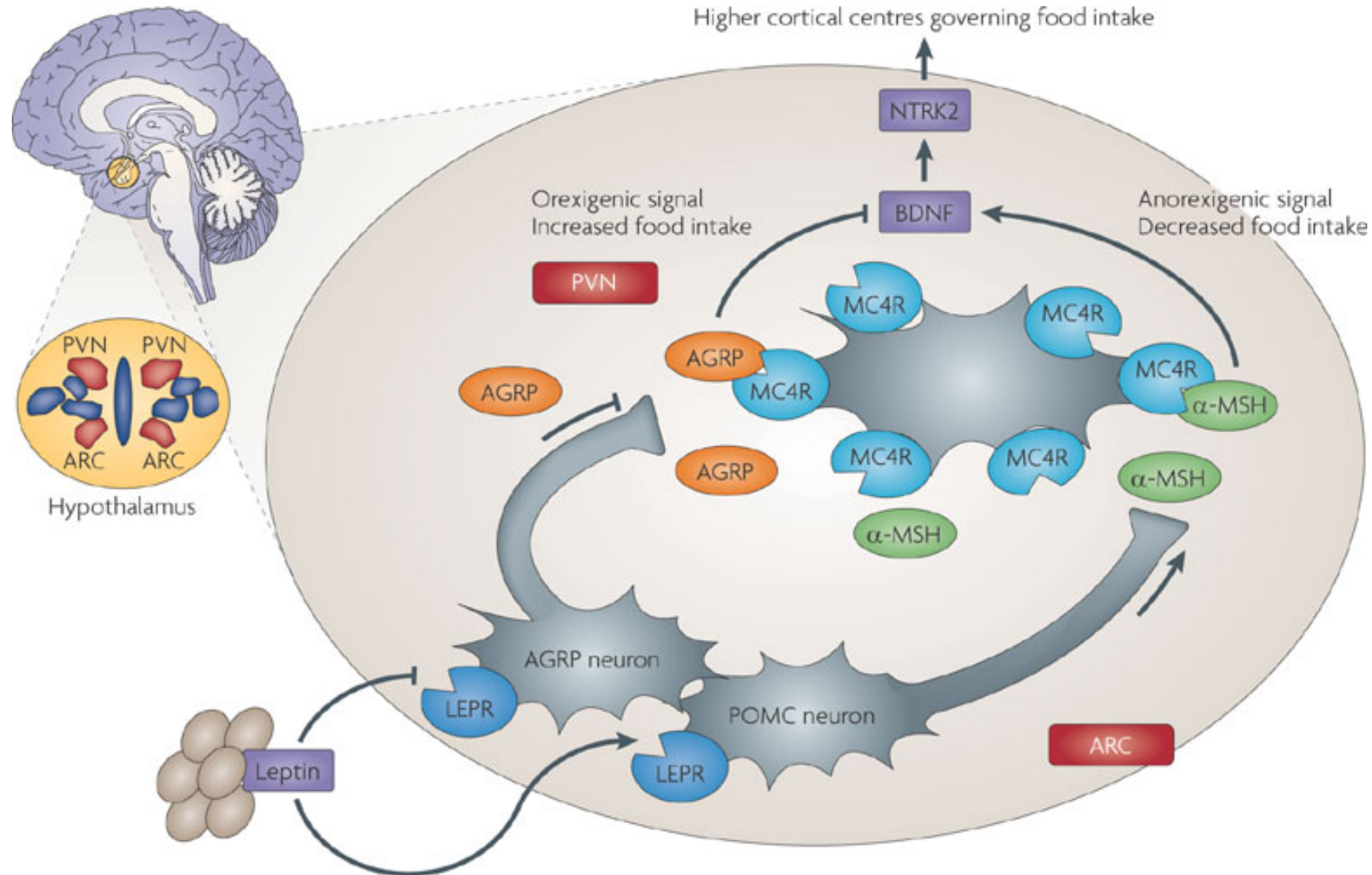
Regulation of Food Intake



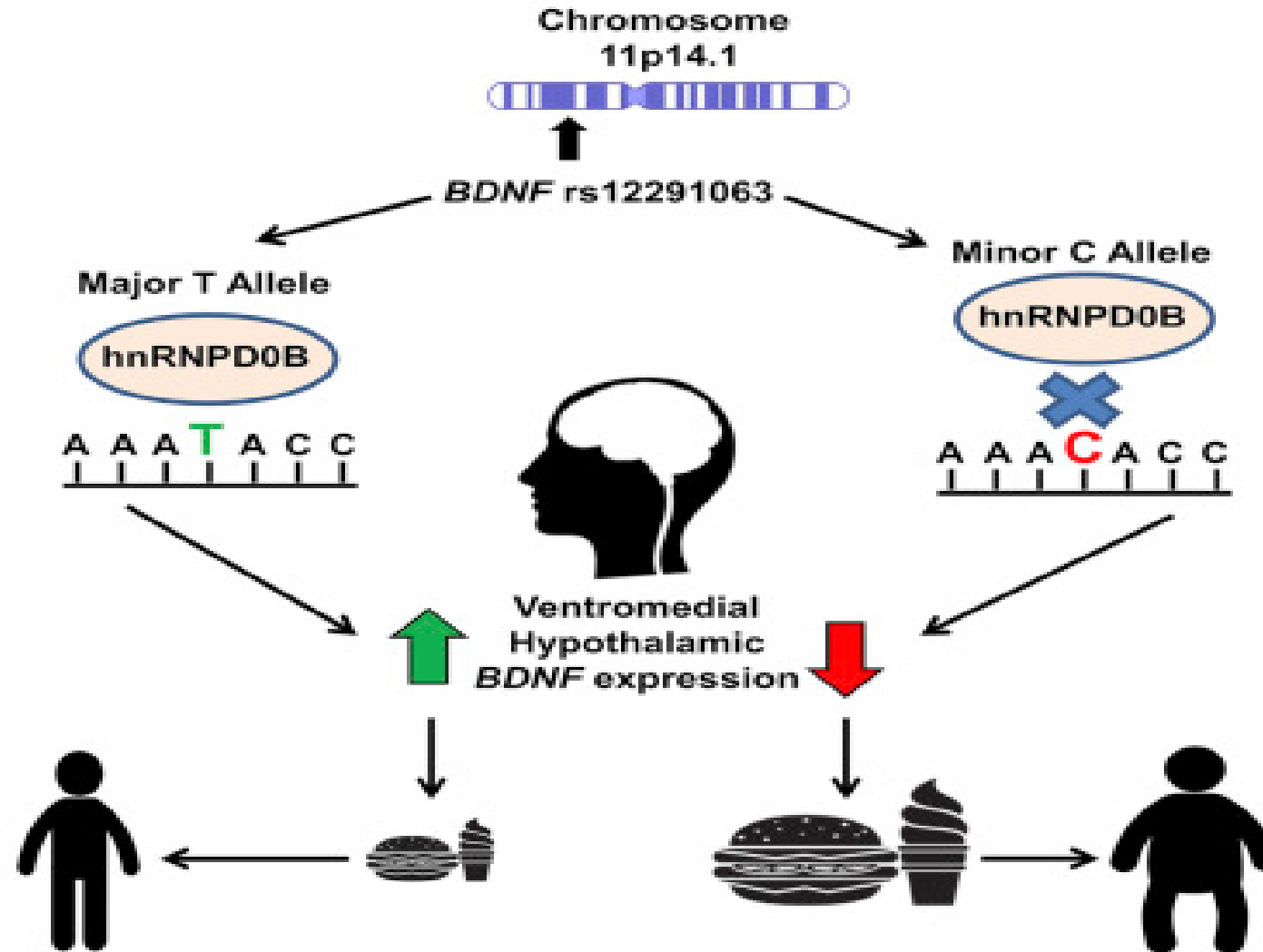
Regulation of Food Intake

Substance	Production Site	Effect (Relevant for Feeding)
Ghrelin (grow)	<ul style="list-style-type: none"> •Stomach (fundus region, entero-endocrine cells) •Neurons in the hypothalamus 	Appetite (orexigenic)
Anandamide (endocannabinoids, ananda; bliss, delight + amide)	Small intestine	Appetite (orexigenic)
Insulin (insula; island or islet)	Pancreas (β -cells in islets of Langerhans)	<ul style="list-style-type: none"> •Satiety (anorexigenic) •glycogen and lipid storage
Leptin (leptos, thin)	<ul style="list-style-type: none"> •Adipocytes (long term) •Stomach (short term) 	Satiety (anorexigenic)
CCK (cholecystokinin, "move the bile-sac")	Small intestine	<ul style="list-style-type: none"> •Early satiety (anorexigenic) •release of digestive enzymes from exocrine pancreas, bile from the gallbladder and H^+ from parietal cells in stomach
PYY (peptide tyrosine tyrosine)	<ul style="list-style-type: none"> •Ileum •colon 	Satiety (anorexigenic)

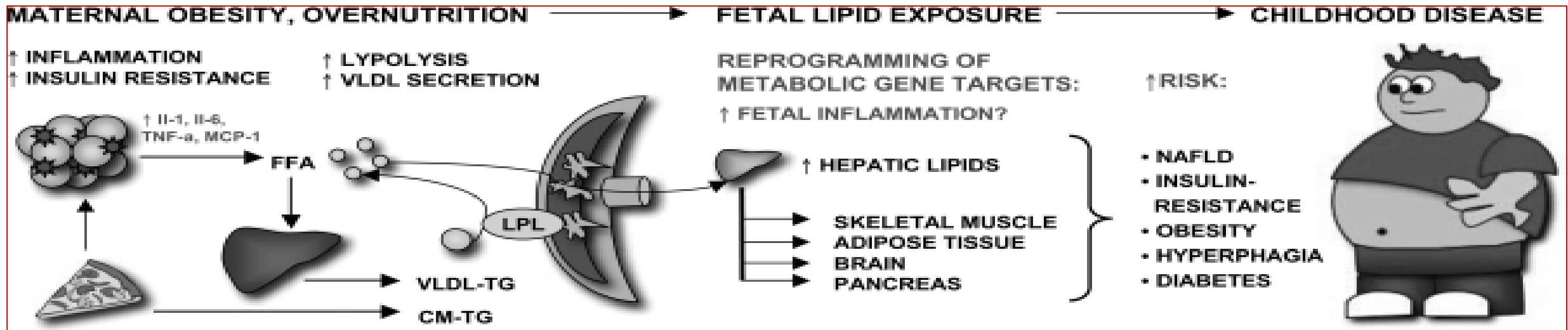
Regulation of Food Intake



BDNF Regulation and Obesity



Development- Fetal Programming



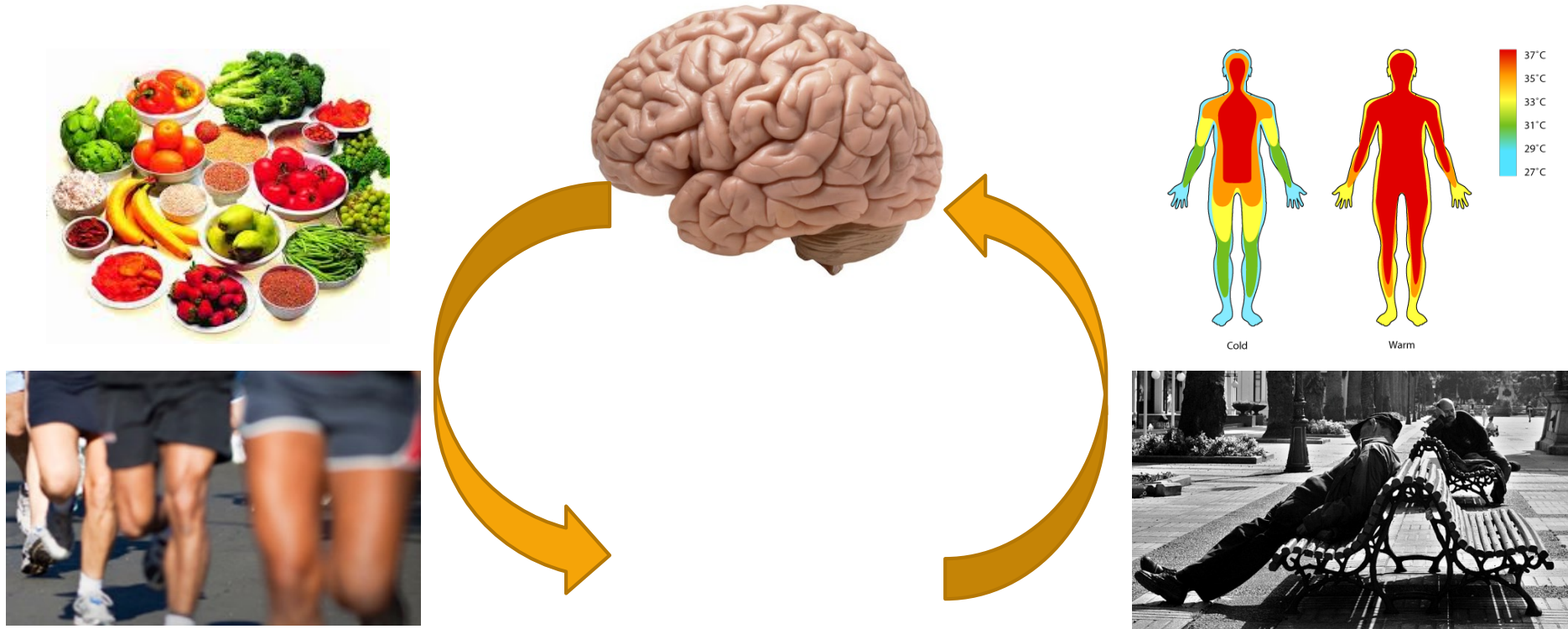
Anthropometric and Plasma Profile of Children Born BMS vs. AMS

	BMS (n=54)	AMS (n=57)	P-value
Male/Female	26/28	24/33	NS
Birth weight (kg)	3.3± 0.1	2.9 ± 0.1	0.003
Macrosomia	8 (14.8%)	1 (1.8%)	0.03
Age	16.6 ± 0.6 (6.8-25.5)	10.7 ± 0.5 (2.4-20.8)	<0.0001
BMI %	79.4 ± 3.4	68.4± 4.0	0.04
Weight (kg)	74.5 ± 4.5	46.5 ± 3.5	<0.0001
Body Fat %	29.7 ± 2.3	24.1 ± 2.1	0.08
Insulin (μU/ml)	21.1 ± 1.8	15.1 ± 1.5	0.01
Glucose (mmol/L)	5.08 ± 0.06	4.86 ± 0.05	0.008

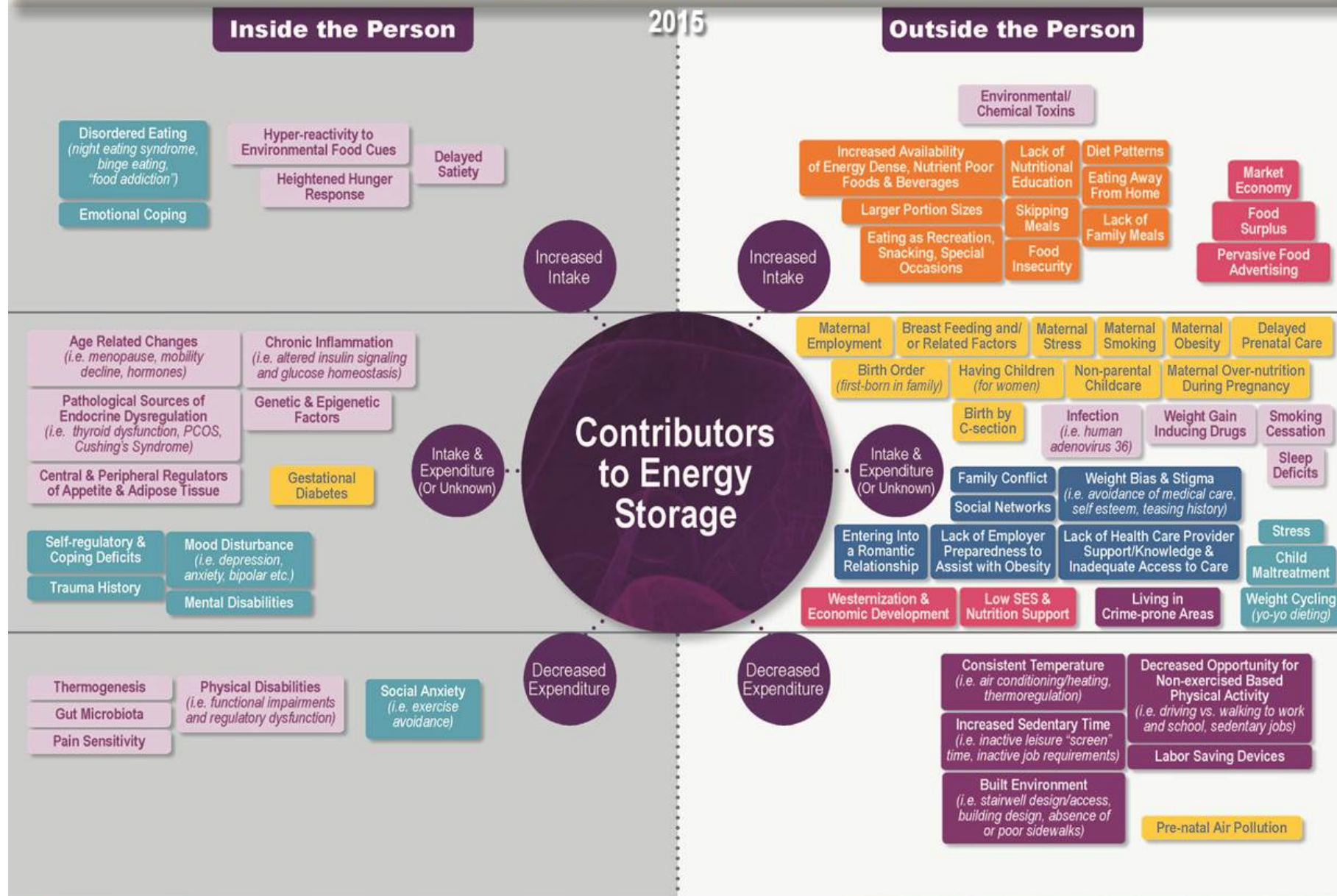
Children Born AMS vs. BMS

- ▶ 3-fold↓ prevalence of severe obesity (11 vs. 35%, $P = 0.004$)
- ▶ ↑insulin sensitivity (homeostasis model assessment of insulin resistance index 3.4 ± 0.3 vs. 4.8 ± 0.5 , $P = 0.02$)
- ▶ Improved lipid profile
 - ▶ cholesterol/HDL cholesterol 2.96 ± 0.11 vs 3.40 ± 0.18 , $P = 0.03$
 - ▶ HDL cholesterol 1.50 ± 0.05 vs. 1.35 ± 0.05 mmol/liter, $P = 0.04$)
- ▶ ↓CRP (0.88 ± 0.17 vs. 2.00 ± 0.34 $\mu\text{g/ml}$, $P = 0.004$)
- ▶ ↓Leptin (11.5 ± 1.5 vs. 19.7 ± 2.5 ng/ml, $P = 0.005$)
- ▶ ↑ghrelin (1.28 ± 0.06 vs. 1.03 ± 0.06 ng/ml, $P = 0.005$)

Central Nervous System regulates weight

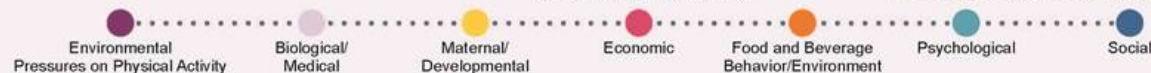


POTENTIAL* CONTRIBUTORS TO OBESITY

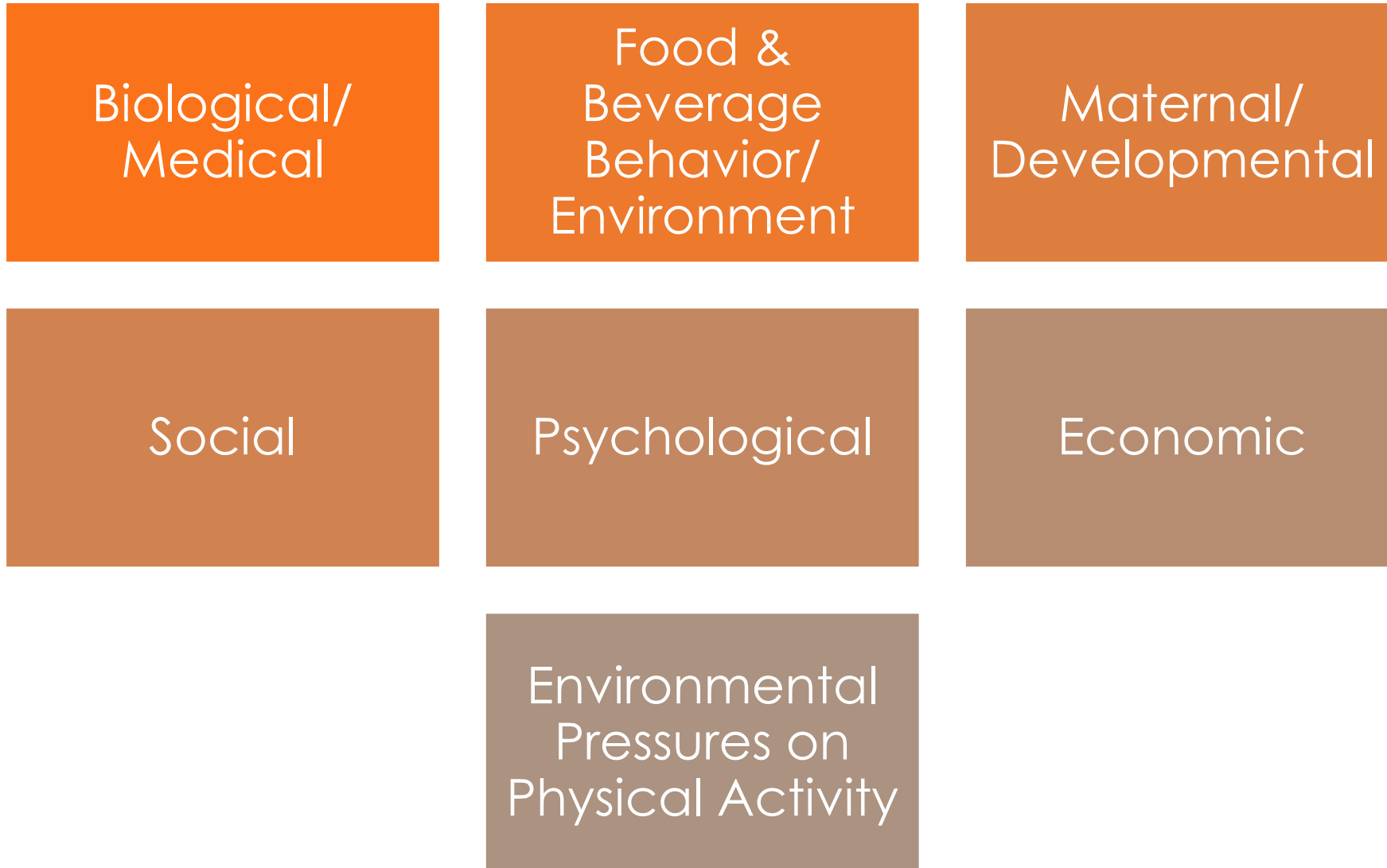


Contributor/Influencer

* Potential contributors indicate anything that has been put forth in the research literature as a question of investigation and is not intended to be a verification of whether or not, or the extent to which, each may or may not contribute.



Contributors/ Influencers to Obesity



Contributors to Obesity- **Inside** the Person

↑ Intake

Hyper-reactivity to
Environmental Food
Cues

Delayed Satiety

Disordered Eating

↓ Expenditure

Gut Microbiota

Thermogenesis

Physical Disabilities

↑ Intake/ ↓ Expenditure

Genetic and
Epigenetic Factors

Age Related
Changes

Mood Disturbances

Contributors to Obesity- **Outside** the Person

↑ Intake

Environmental/
Chemical Toxins

Pervasive
Food advertising

Large Portion Sizes

↓ Expenditure

Built Environment

Sedentary Time

Labor Saving
Devices

↑ Intake/
↓ Expenditure

Stress

Weight Cycling

Maternal/Paternal
Obesity

Toddler's Bias and Maternal Anti-fat Prejudice

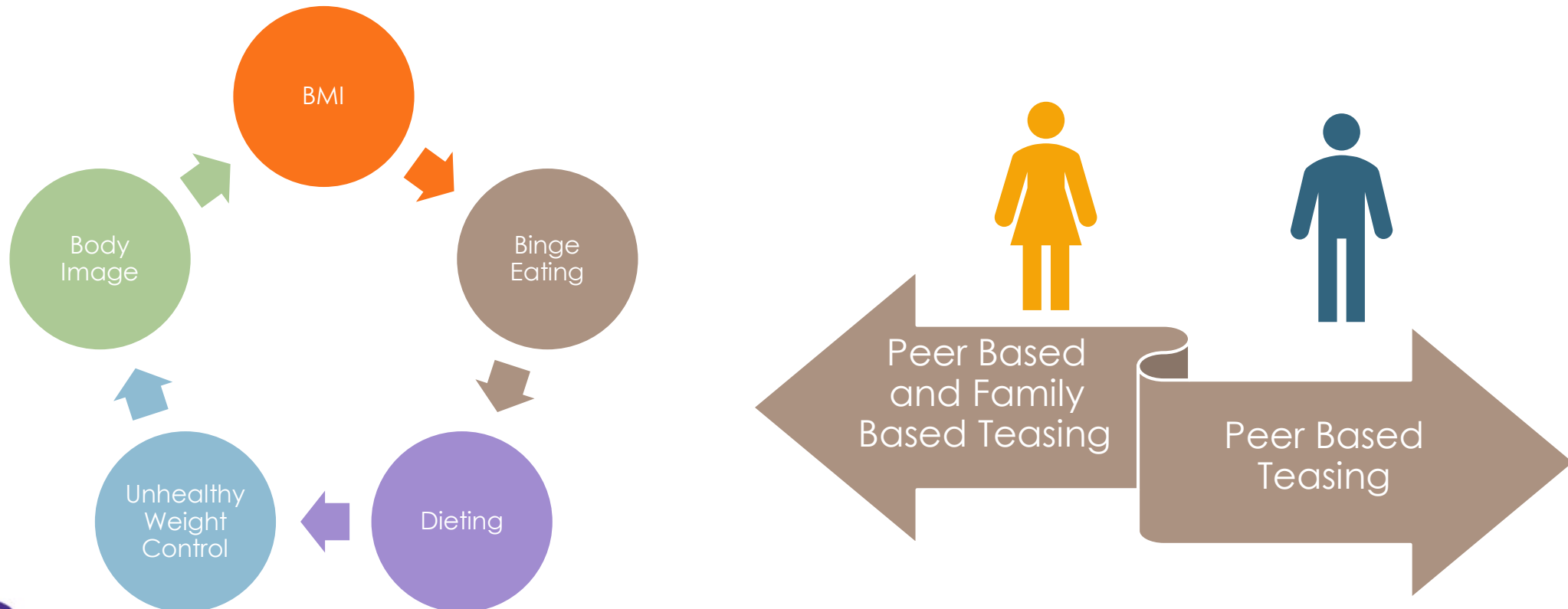


- Mother-child dyads (N=70)
- Older infants (M=11 months) had a bias for looking at the figures with obesity
- Older toddlers (M=32 months) instead preferred looking at the average-sized figures
- Older toddlers' preferential looking was correlated significantly with maternal anti-fat attitudes



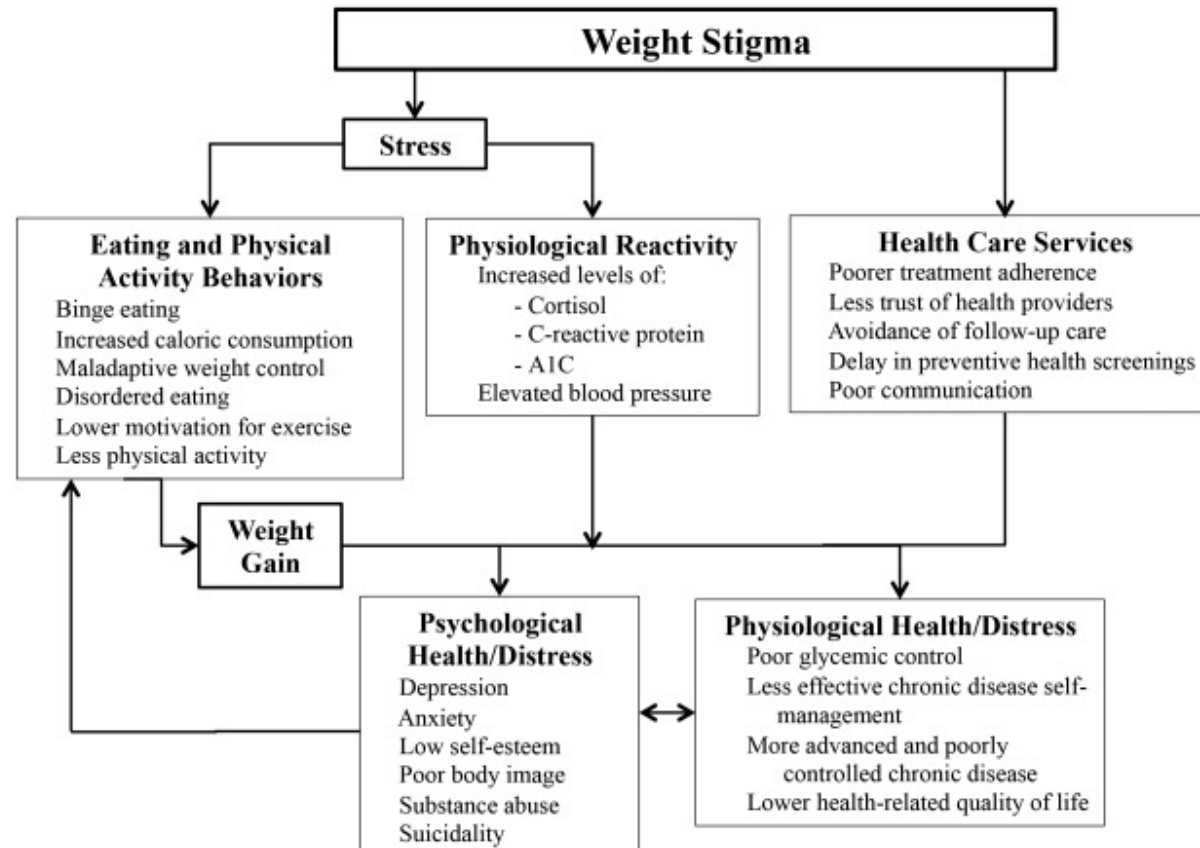
Weight Teasing in Adolescence and Weight Related Outcomes in Adulthood

Weight-based teasing in adolescence predicted higher BMI and obesity 15 years later



Prev Med. 2017 Jul;100:173-179.

Overcoming Weight Stigma in the Treatment of Obesity



Guidelines for Selecting Obesity Treatment

BMI Category

Treatment	25-26.9	27-29.9	30-34.9	35-39.9	≥40
Diet, PA, & Behavioral Therapy	With co-morbidities	With co-morbidities	+	+	+
Pharmacotherapy		With co-morbidities	+	+	+
Weight Loss Surgery			With co-morbidities		

•Prevention of weight gain with lifestyle therapy is indicated in any patient with a **BMI ≥ 25 kg/m²**, even without co-morbidities, while weight loss is not necessarily recommended for those with a BMI of 25–29.9 kg/m² or a high waist circumference, unless they **have two or more co-morbidities**.

•Consider pharmacotherapy only if a patient has not **lost 1 pound per week** after **6 months** of combined lifestyle therapy.

The + represents the use of indicated treatment regardless of co-morbidities.

Common Weight Promoting Medications

▶ **Anti-psychotics**

- ▶ Risperidone
- ▶ Lithium
- ▶ Quetiapine
- ▶ Aripiprazole
- ▶ Olanzapine
- ▶ Valproic Acid

▶ **Anti-depressants**

- ▶ Citalopram
- ▶ Duloxetine
- ▶ Venlafaxine

● **Sleep Agents**

- Zolpidem
- Eszopiclone
- Trazadone
- Zaleplon

● **Neuropathic Agents**

- Gabapentin
- Pregablin

● **β-Blockers**

● **Steroids**

● **Insulin**

● **Hypoglycemic Agents**

Treatment Strategy for Weight Promoting Medications

- ▶ Investigate whether medications are a likely source of weight gain in patients.
- ▶ If a weight promoting drug may be discontinued, discontinue the agent.
- ▶ If discontinuation of a weight promoting medication is not feasible, consider the use of anti-obesity pharmacotherapy for weight loss in conjunction with appropriate lifestyle changes.

Anti-obesity pharmacotherapy agents

- ▶ Most agents may be characterized into **3 primary groups**
 - 1) Centrally acting medications that impair dietary intake
 - 2) Medications that act peripherally to impair dietary absorption
 - 3) Medications that increase energy expenditure

FDA Approved Anti-obesity pharmacotherapy agents

Drug class/name
CNS Stimulants/ Anorexiant: Phentermine *Phentermine/topiramate Diethylpropion Phendimetrazine Benzphetamine
Anti-Depressants/ Dopamine Reuptake Inhibitors/ Opioid Antagonists: *Bupropion/ Naltrexone
Gastrointestinal Agents/Other: *Orlistat *GLP-1 agonists (liraglutide)

Other Anti-obesity pharmacotherapy agents

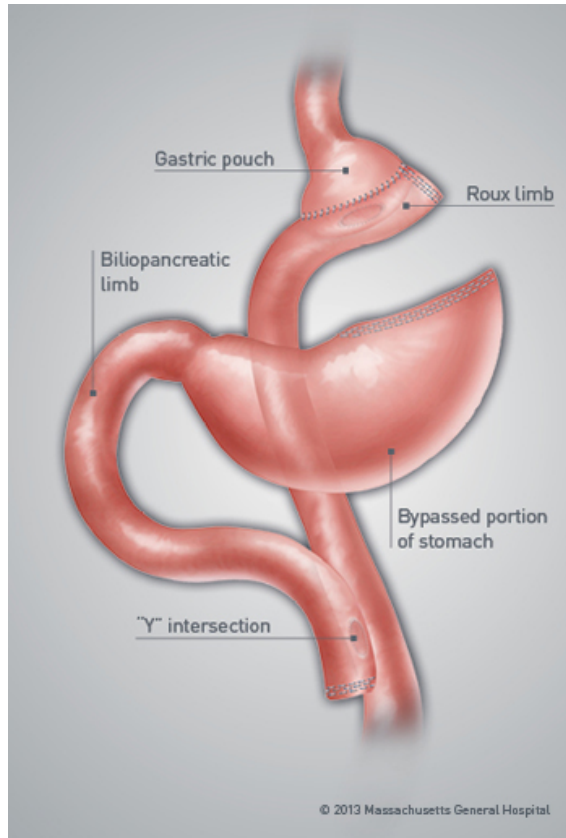
Drug class/name
Topiramate
Zonisamide
Bupropion
Metformin
Amylin agonist (pramlinitide)
SGLT2 Inhibitors (canagliflozin, dapagliflozin)

Criteria for Weight Loss Surgery

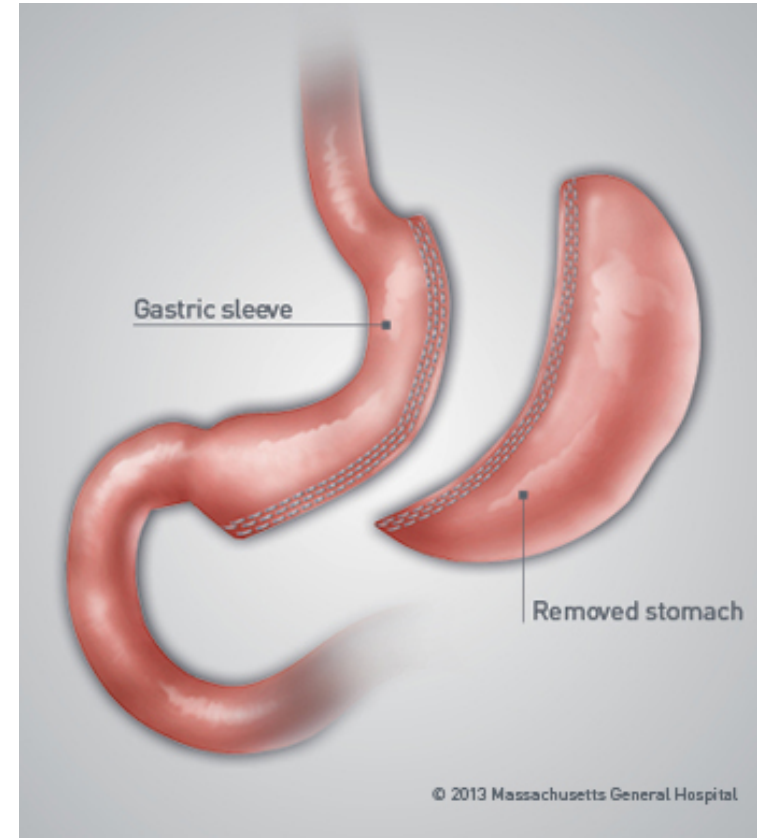
- ▶ Body Mass Index (BMI) ≥ 40 OR
- ▶ BMI of 35-39.9 + 1 serious co-morbidity
 - ▶ Type 2 Diabetes Mellitus
 - ▶ Coronary Artery Disease
 - ▶ Obstructive Sleep Apnea
- ▶ Prior Unsuccessful Weight Loss Attempts
- ▶ Acceptable operative risks
- ▶ Ability to participate in treatment and long term follow-up
- ▶ An understanding of the operation and the lifestyle changes needed to sustain long term weight loss

Most Common Weight Loss Surgeries in the US

Roux-en-Y Gastric Bypass



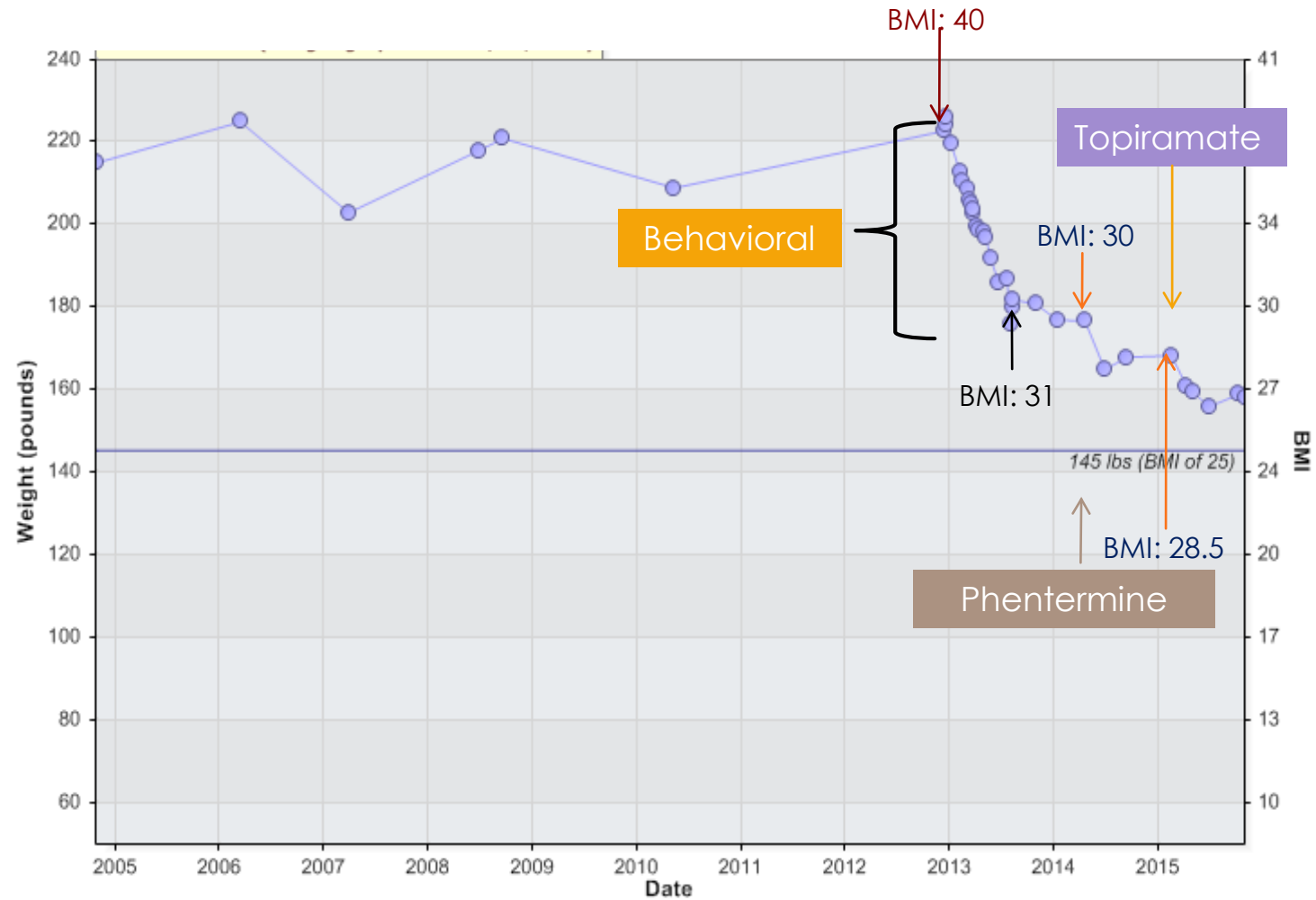
Vertical Sleeve Gastrectomy



Case #1

- ▶ 54-year-old woman
- ▶ Past medical history:
 - ▶ Untreated hypertension
 - ▶ Migraine headaches
 - ▶ GERD
 - ▶ Irritable Bowel Syndrome
 - ▶ Metabolic syndrome
- ▶ Retained 20 lbs with each of her 2 pregnancies
- ▶ Tried many commercial programs which lead to 20 lbs of unsustainable weight loss with each attempt
- ▶ Most significant weight loss with the use of phen-fen in the 1990's (~50 lbs over 6 months)
- ▶ Interested in weight loss medications + behavioral Tx

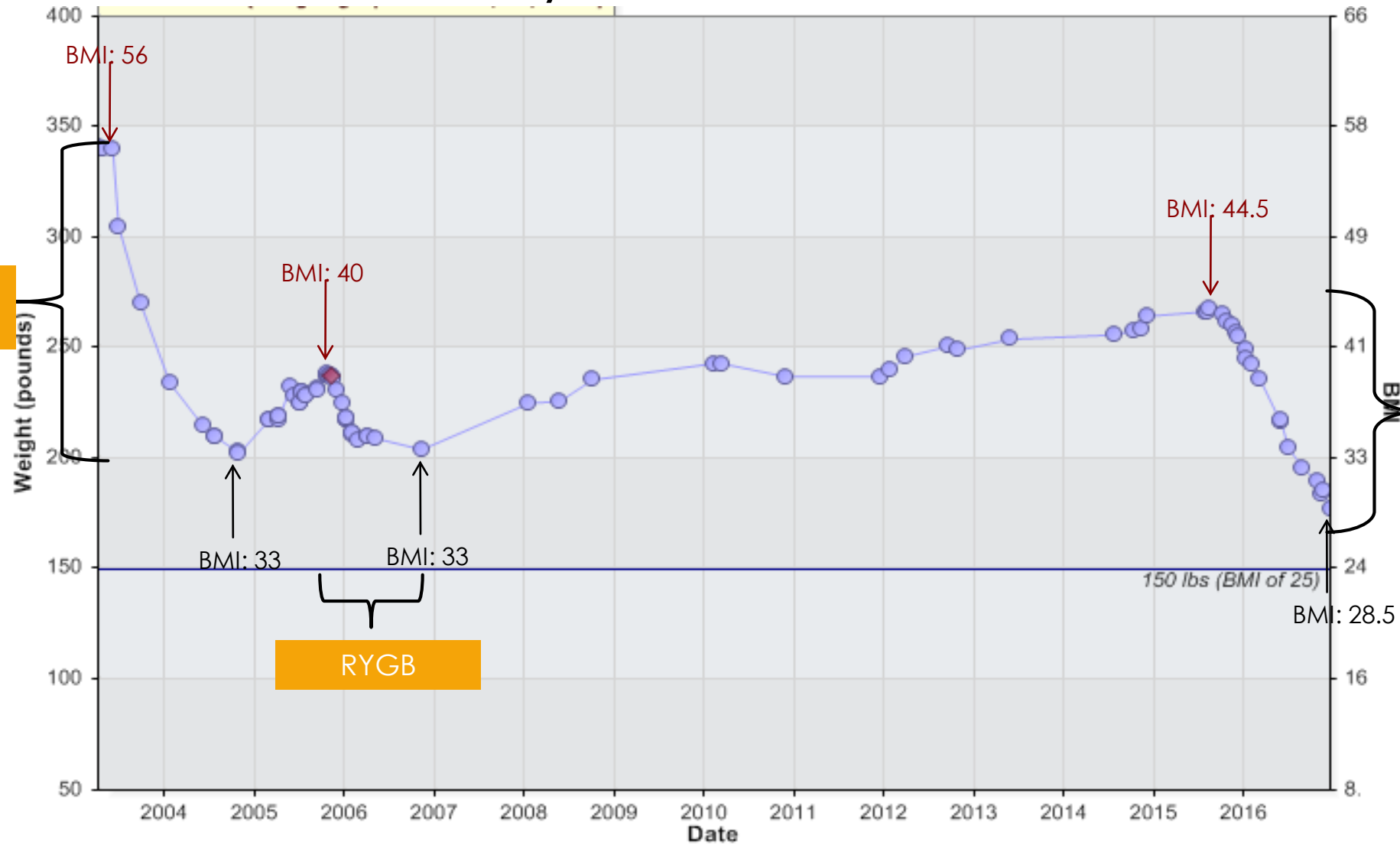
54-year-old woman



Case #2

- ▶ 57-year-old woman
- ▶ Past medical history:
 - ▶ Dyslipidemia
 - ▶ Breast Cancer
 - ▶ Hypertension
 - ▶ Depression
 - ▶ Pernicious Anemia
- ▶ Diet:
 - ▶ Breakfast: Brown Rice, Cashews, Goat Cheese
 - ▶ Snack: Denies
 - ▶ Lunch: Fish; may be on a sandwich with vegetables
 - ▶ Snack: Cheese and Crackers, Cashew nuts, Protein Bars
 - ▶ Dinner: Salad (Spinach) with cucumbers, tomatoes, goat cheese, peppers, vinaigrette
 - ▶ Snack: Cheese and Crackers, Cashew nuts, Protein Bars
- ▶ Activity: Exercise class (cardio interval circuit- 3 times per week, 1 hour); 2 videos 1/2 hour (low impact cardio); Yoga at night
- ▶ Sleep: 8 hours per night of restful sleep

57-year-old woman



Phentermine/
Topiramate

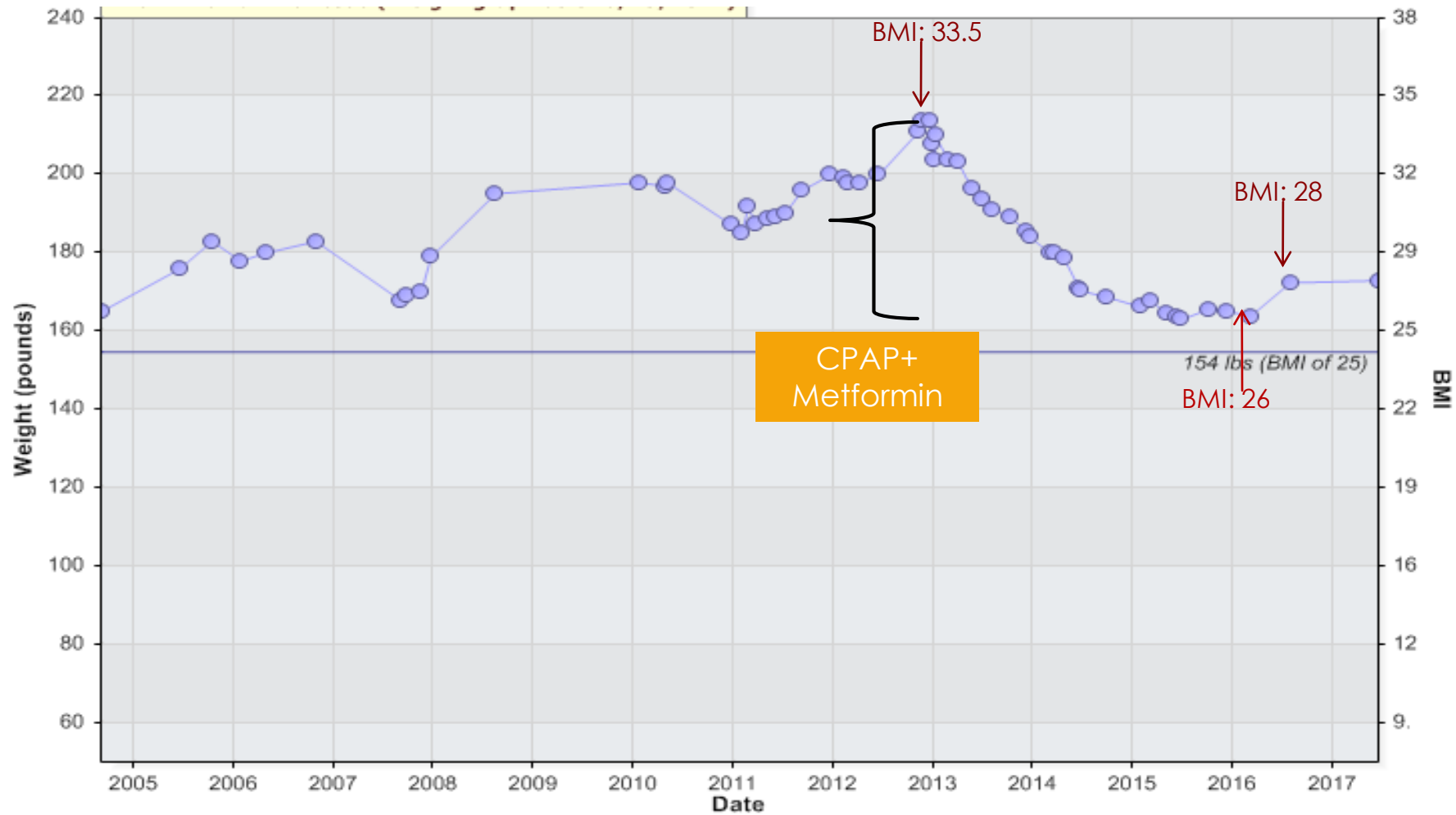
Phentermine
+Topiramate

RYGB

Case #3

- ▶ 46-year-old woman
- ▶ Past medical history:
 - ▶ Hypertension
 - ▶ Anxiety/ Depression
 - ▶ Asthma
 - ▶ Fibromyalgia
 - ▶ Bipolar disorder
 - ▶ Gastroesophageal reflux disease
 - ▶ Metabolic syndrome X
- ▶ History of being on several weight promoting: Ziprasidone, Quetiapine, Duloxetine, Citalopram, Fluoxetine, Zolpidem, Trazodone, Atenolol, Pregablin, and Nortriptyline
- ▶ Postpartum weight retention of 20 pounds
- ▶ Poor sleep- daytime hypersomnolence, snoring, morning headache

46-year-old woman



Ann Pharmacother. 2010 Apr;44(4):668-79
Am J Psychiatry. 2013 Sep;170(9):947-52.

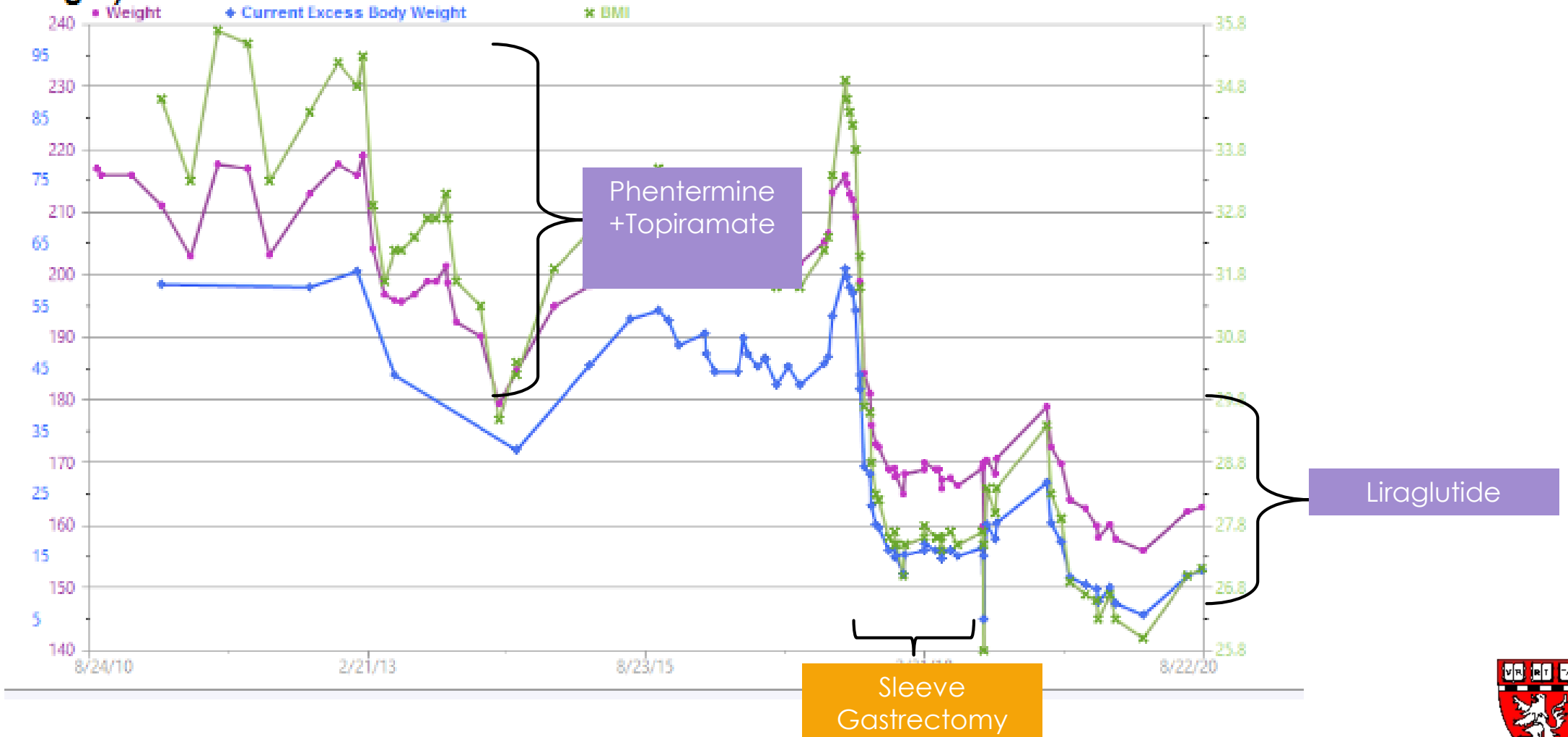
Case #4

- ▶ 64-year-old man
- ▶ Past medical history:
 - ▶ Type 2 Diabetes Mellitus
 - ▶ Male hypogonadism
 - ▶ Vitamin D deficiency
 - ▶ Obstructive sleep apnea on CPAP
 - ▶ Dyslipidemia
 - ▶ Hypertensive disorder
 - ▶ Steatosis of liver
- ▶ Hunger: Normal to high on topiramate; liraglutide
- ▶ Satiety: Fills Quickly
- ▶ Physical Activity: Peloton- 3-4 times per week (45 minutes)
- ▶ Sleep: 7 hours on CPAP
- ▶ Stress: Normal

64-year-old man

At this time, he has lost 92.2% of his excess body weight and 22.4% of his total body weight

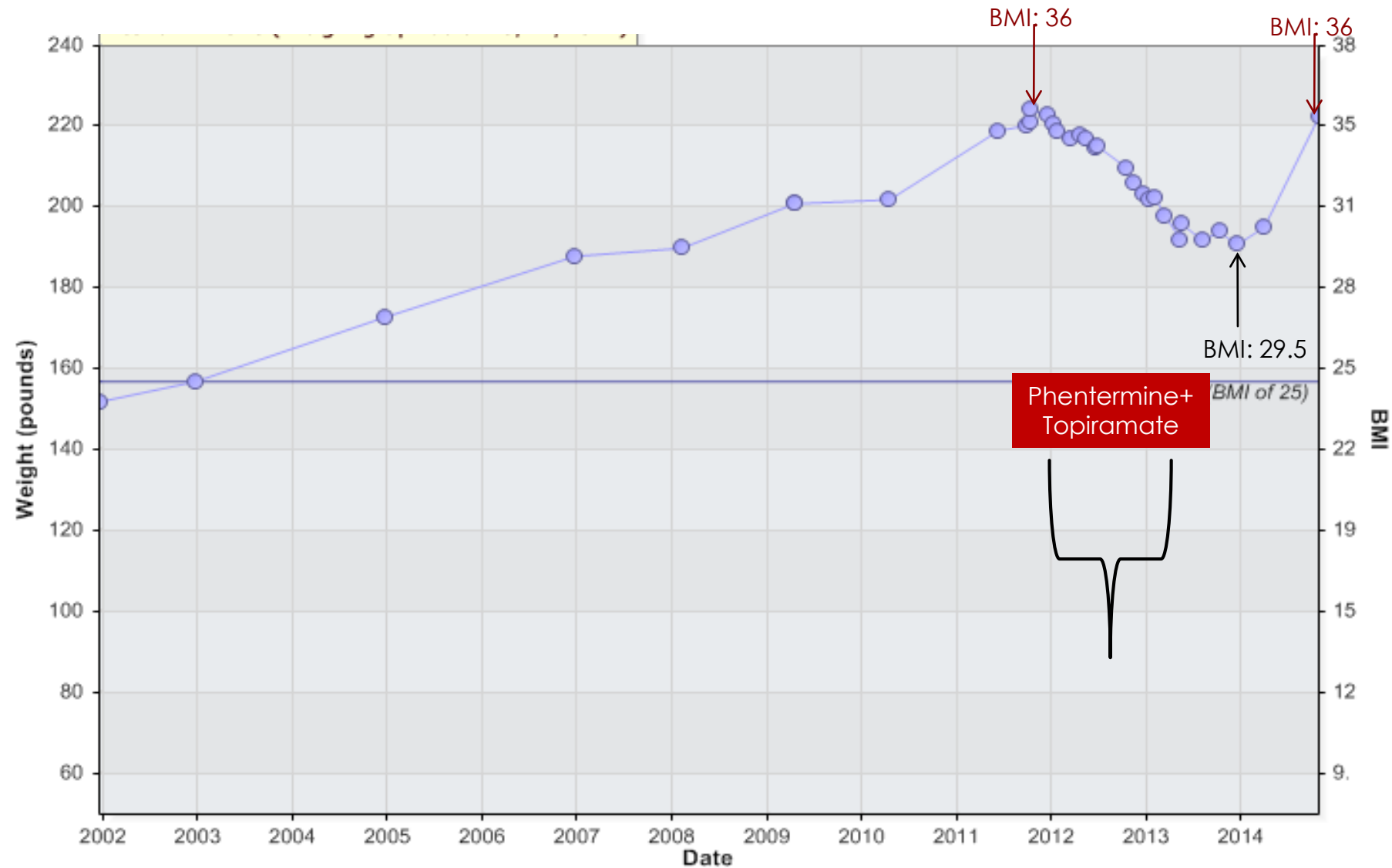
(NADIR post op: Lost 81.5% of his excess body weight and 20.4% of his total body weight)



Case #5

- ▶ 36-year-old woman
- ▶ Past medical history:
 - ▶ Hypothyroidism
 - ▶ Dysthymia
 - ▶ Allergic Rhinitis
 - ▶ Chronic Back Pain
 - ▶ Migraine Headaches
- ▶ Diet:
 - ▶ Breakfast: Oatmeal and Egg Whites
 - ▶ Lunch: Salad with Chicken
 - ▶ Snack: Fruit (Less Recently)
 - ▶ Dinner: Fruit, Chicken, Rice, Broccoli, Increase in Plant Based Protein
- ▶ Activity: Cardio at gym- 5 days per week (elliptical)-1 hour; Walks at lunch time (45 minutes); strength training with 3 times per week- 1 hour
- ▶ Sleep: 6-7 hours per night of restful sleep

36-year-old woman



Summary

- ▶ Obesity is a multifactorial disorder that is syndemic
- ▶ Conventional weight loss consisting of diet and exercise should be 1st line
- ▶ Pharmacotherapy and/or bariatric surgery should be considered as an option for appropriate patients
- ▶ Weight bias is seen as early as infancy
- ▶ Maternal and Paternal anti-fat bias influences children
- ▶ Weight stigma has a negative impact on the health and psychological health of patients who struggle with obesity
- ▶ Obesity is a chronic disease which requires lifelong treatment

Thank You For Your Time

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