The Biology of Stress

Exploring the causes, types, and effects...

Mini Med School

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a place of mind THE UNIVERSITY OF BRITISH COLUMBIA

Faculty of Medicine







Territorial Acknowledgement

We acknowledge with respect the Lekwungen peoples on whose traditional territory the university stands and the Songhees, Esquimalt and <u>WSÁNEĆ</u> peoples whose historical relationships with the land continue to this day.

Introductions and Disclosures

• About the optional survey

• This talk will be recorded

Introductions and Disclosures

• I am a second year medical student

• This talk is intended for your entertainment and education, and is not meant to replace advice from your physician or another health care professional

• All pictures used are free stock photos or photos available under a Creative Commons license unless otherwise noted

• Thank you for joining us!

Agenda

- What is stress?
- Epidemiology of stress
- Types of stress
- Stress response pathway
- Effects of stress acute
- Intermission / Q&A
- Effects of stress chronic
- When to seek help
- Preview: stress management
- Suggested Resources
- Q&A

What is stress?

- A feeling of emotional or physical tension
 - Physiologic stressors
 - Put a strain on our body (e.g., injury, extreme temperatures, etc.)
 - Psychological stressors
 - Anything we interpret as negative or threatening (e.g., events, situations, people, comments, etc.)



What is stress?

- Common elements to all stressful situations...NUTS
 - Novelty
 - Unpredictability
 - Threat to the ego
 - Sense of control



https://humanstress.ca/stress/understand-your-stress/sources-of-stress/

- Statistics Canada survey 2020: Perceived life stress
 - 20.4% of people reported "most days are quite or extremely stressful"
 - Highest among those aged 35-49 (27.5%)
 - Lowest among those aged 65+ (10.3%)



Poll

What is the most common source of stress in people's day-today lives?

- Financial concerns
- Family
- Work
- Time pressures/not enough time

- Statistics Canada survey 2020: Main source of stress in day-to-day life
 - 1. Work
 - 2. Financial concerns
 - 3. Family
 - 4. Time pressures / not enough time





stress in America 2020: A NATIONAL MENTAL HEAlth crisis COVID-19 Is a Significant Stressor for Most Americans



Stress in America 2020 study: COVID-19

Nearly 8 in 10 (78%) say the coronavirus pandemic is a significant source of stress in their life

Nearly 7 in 10 (67%) say they have experienced increased stress over the course of the pandemic

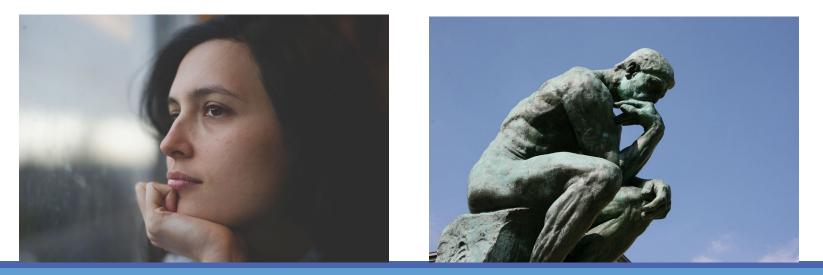
- Stress in America 2021 study: COVID-19 and decision making
 - 63% of adults agreed that uncertainty about what the next few months will be like causes them stress



- Stress in America 2021 study: COVID-19 and decision making
 - 49% of adults said the pandemic makes planning for the future feel impossible
 - Younger adults, especially millennials feel this way



- Stress in America 2021 study: COVID-19 and decision making
 - 61% of adults agreed the pandemic has made them re-think how they were living their life





Poll

True or false?

• All stress is bad

Types of stress

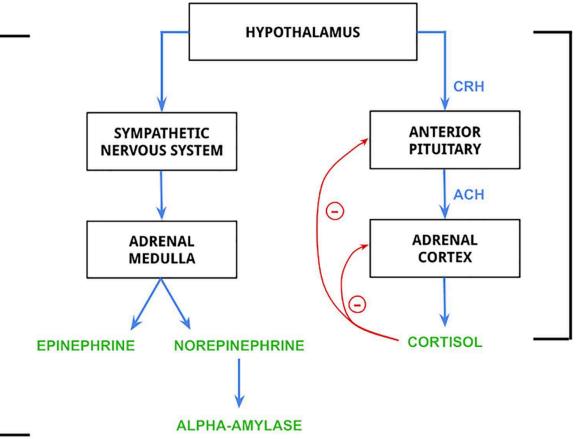
- Eustress (positive stress)
 - E.g., before performing in front of people
- Distress (negative stress)
 - Acute/episodic
 - E.g., Loss of a loved one
 - Chronic/toxic
 - E.g,. abuse

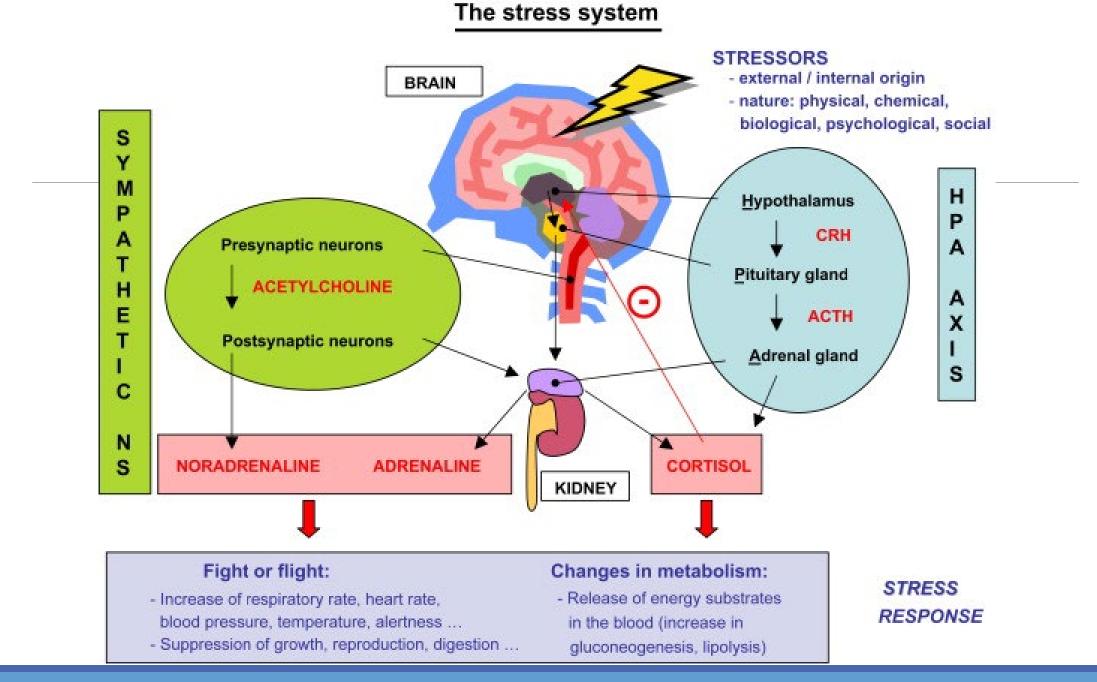




SYMPATHETIC ADRENAL MEDULLARY AXIS

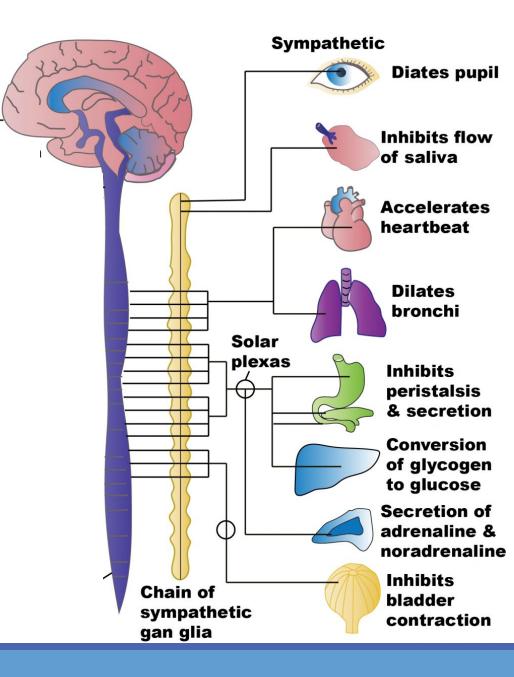
- The stress response is mediated by two systems:
 - Fast response: Sympathetic Nervous System (SNS)
 - Epinephrine (Adrenaline)
 - Slow response: Hypothalamus-Pituitary-Adrenal (HPA) Axis
 - Cortisol



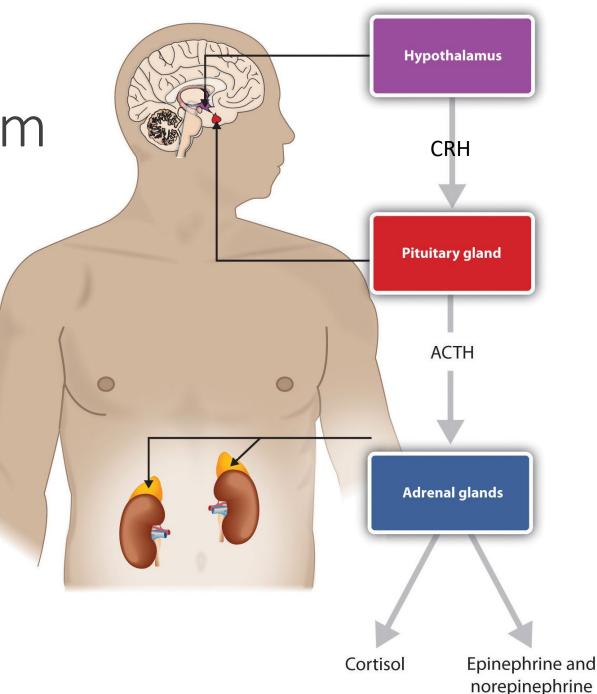


T. De Vriendt, L.A. Moreno, S. De Henauw, Chronic stress and obesity in adolescents: Scientific evidence and methodological issues for epidemiological research, Nutrition, Metabolism and Cardiovascular Diseases, Volume 19, Issue 7, 2009, Pages 511-519, ISSN 0939-4753, https://doi.org/10.1016/j.numecd.2009.02.009.

- Sympathetic Nervous System
 - Increases heart rate
 - Increases blood pressure
 - Enhances arousal
 - Reduces gut motility

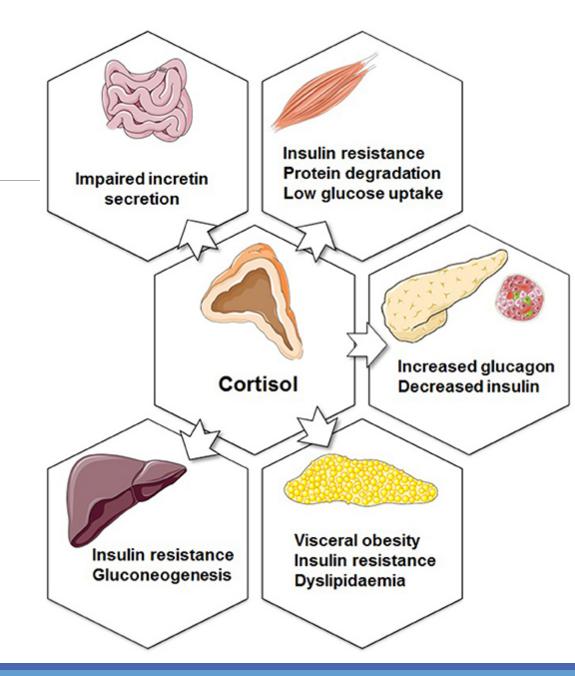


- HPA Axis: Cortisol
 - Promotes release of epinephrine
 - Suppresses immune response
 - Mobilization of energy stores



• HPA Axis: Cortisol

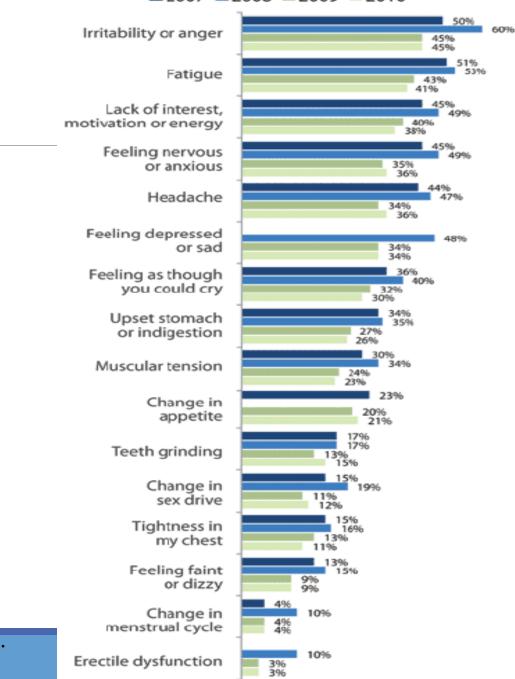
- Increases blood glucose
 - Promotes release of glucose from liver
 - Counteracts insulin
- Promotes lipolysis (breakdown of fat)





- What are the experienced effects of these systems?
 - Irritability or anger
 - Fatigue/lack of motivation
 - Feeling anxious
 - Headache
 - Upset stomach/indigestion

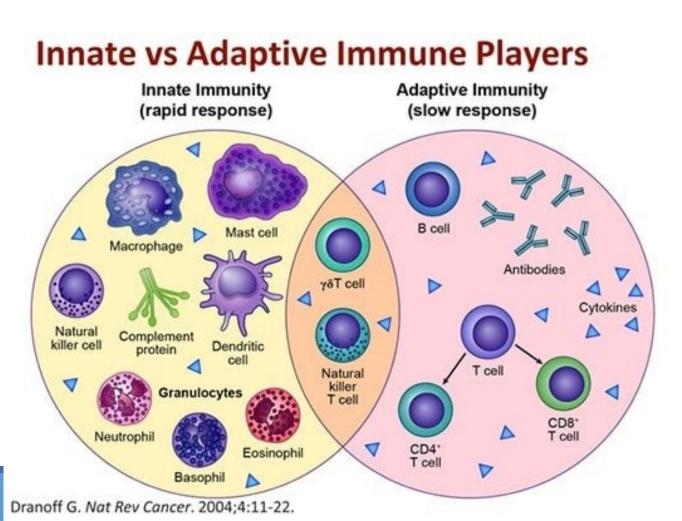
Clay, R. A. (2011, January). Stressed in America. *Monitor on Psychology*, 42(1). http://www.apa.org/monitor/2011/01/stressed-america



•These systems evolved to be adaptive/beneficial in times of stress

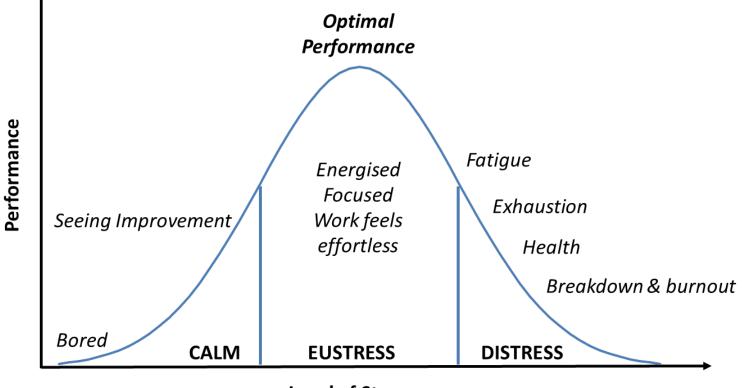
- In the short-term, they are!
- Short-term stress can have positive impacts on functioning
 - Upregulation of part of the immune system
 - Improved performance and memory

- Immune system
 - Innate: general protection that we are born with
 - E.g., skin as a barrier; mucus
 - Adaptive: immunity that develops throughout our lives with exposure
 - E.g., vaccines



- Immune system
 - Upregulates innate immunity = quick response, energy efficient
 - Prepares body for fighting infections from cuts, bite wounds, injury, etc.
 - Suppresses adaptive immunity = slow response, energy consuming

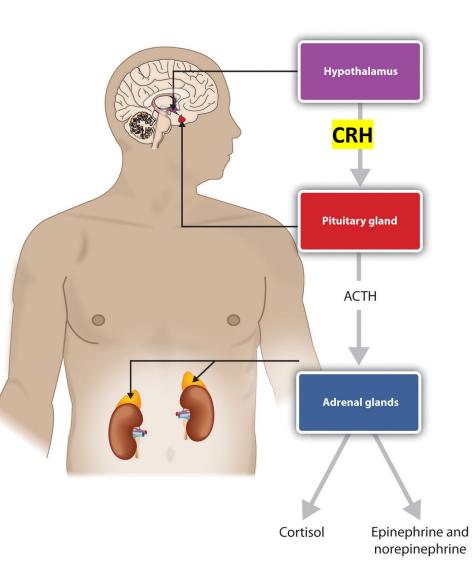
- Performance
 - Yerkes-Dawson Law
 - Eustress (short-term, positive stress) can optimize performance



Level of Stress

Memory

- Limited levels of stress induce rapid memory formation
- Potentially related to HPA axis signaling
 - CRH release from the hypothalamus acts not just on the pituitary, but other areas of the brain that are implicated in learning and memory



Intermission

• 10-minute break





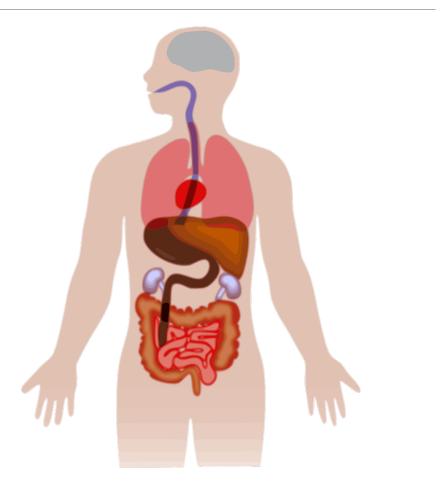


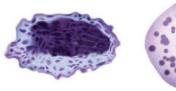
Poll

Which of the following body systems is/are impacted by chronic stress?

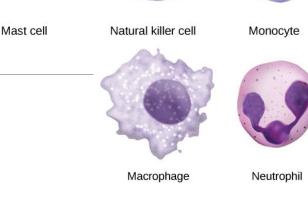
- Cardiovascular system
- Gastrointestinal system
- Immune system
- Nervous system
- All of the above

- Immune system
- Memory/cognition
- Cardiovascular system
- Diabetes
- Obesity
- Gastrointestinal system
- Mental health









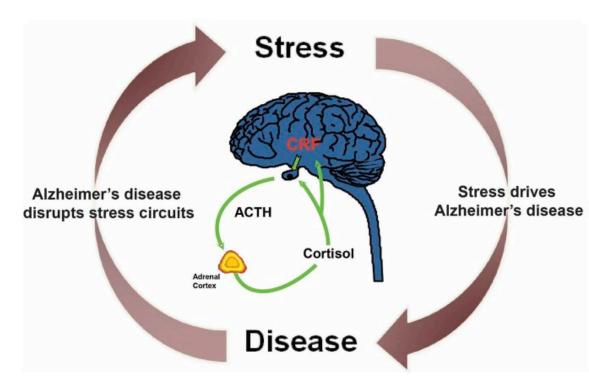


- In contrast to some beneficial effects during acute stress, chronic stress can lead to global immunosuppression and inflammation
 - Increasing stressor duration = shift from adaptive \rightarrow detrimental

• Inflammation + suppression of important immune cells (T cells and Natural Killer cells) may increase susceptibility to some cancers

Memory/cognition:

- Chronic psychologic stress is a risk factor for Alzheimer's Disease
 - Potentially HPA axis-mediated



Justice NJ. The relationship between stress and Alzheimer's disease. *Neurobiol Stress*. 2018;8:127-133. Published 2018 Apr 21. doi:10.1016/j.ynstr.2018.04.002

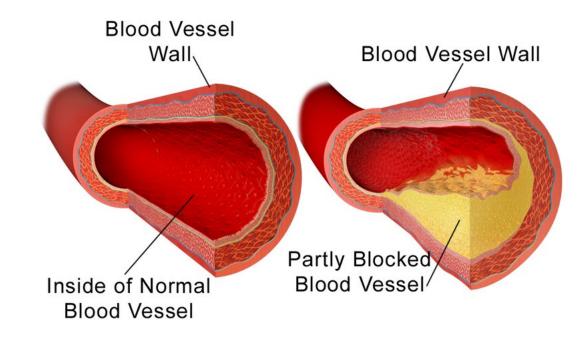
Cardiovascular system:

- Epinephrine (adrenaline) causes heart rate and blood pressure to increase
 - Hypertension damages blood vessel walls
 → atherosclerosis, aneurysm
 - Severe hypertension makes the heart work harder → heart walls can thicken/dilate → heart failure



Cardiovascular system:

 Cortisol and chronic low-grade inflammation contributes to development of atherosclerosis → coronary artery disease



Cardiovascular system:

• Stress can lead to unhealthy coping mechanisms (poor diet, smoking, etc.) that can further contribute to heart disease





Effects of stress

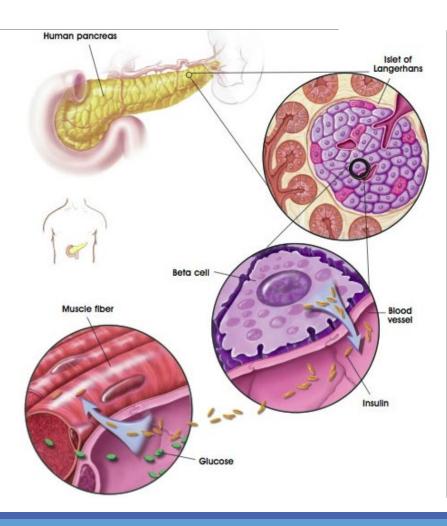
Cardiovascular system:

- Extreme stress can lead to "Broken Heart Syndrome" (Takotsubo cardiomyopathy)
 - Exact mechanism not known, but thought that high levels of stress hormones cause changes in heart muscle cells → weak heart walls and poor contraction of heart → 20% of people experience heart failure
 - Most people recover with no long-term damage



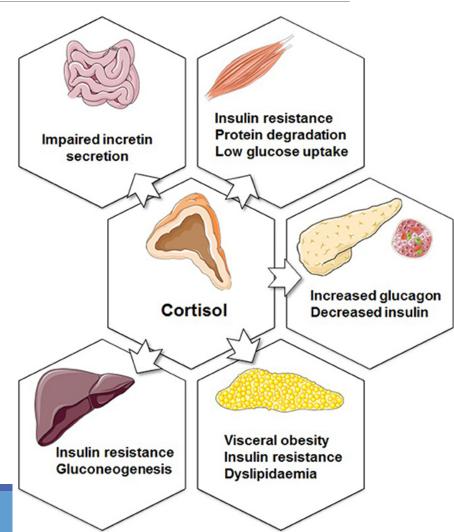
Diabetes:

 Moderate/high levels of stress associated with 2.3-fold increase in the odds of diabetes three years later



Diabetes:

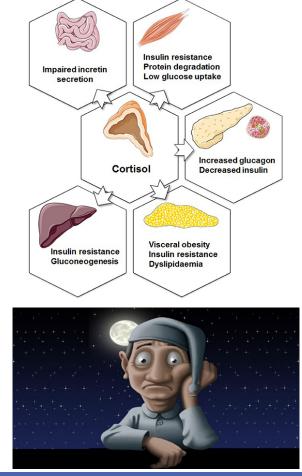
- High levels of cortisol increase blood glucose and may prevent insulin-producing cells from working properly → Type 2 Diabetes
 - Cushing disease (too much cortisol produced)
 → Type 2 Diabetes

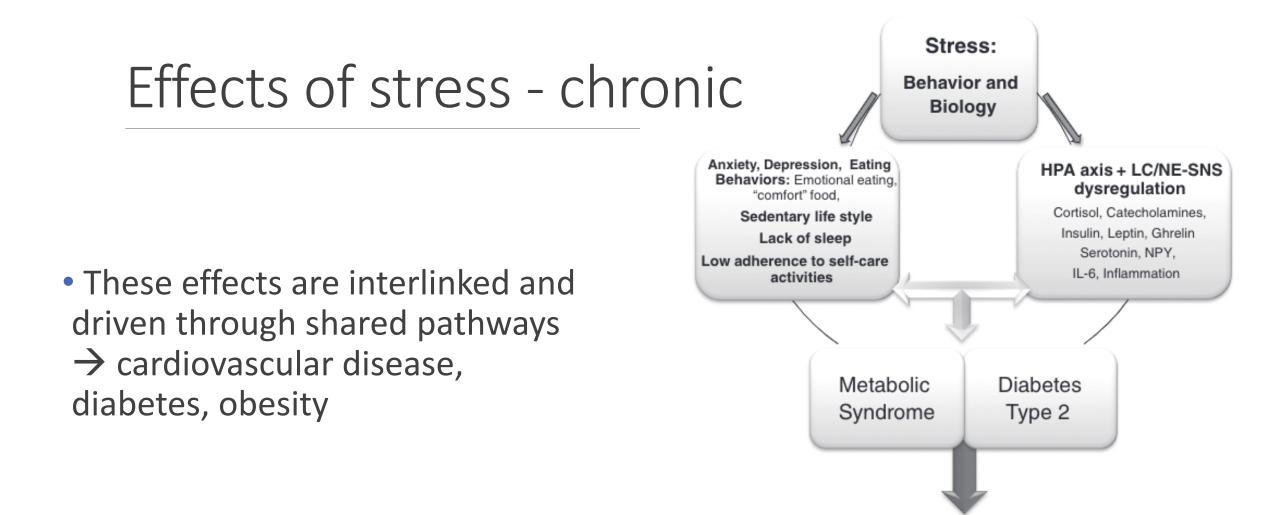


Obesity:

- Behavioural and cellular mechanisms promote obesity and metabolic issues in chronically stressed people
 - Stress \rightarrow unhealthy lifestyles/coping
 - Stress \rightarrow cortisol, epinephrine, inflammation \rightarrow central obesity, insulin resistance
 - Stress \rightarrow poor sleep \rightarrow obesity







Panagiota Pervanidou, George P. Chrousos, Metabolic consequences of stress during childhood and adolescence, Metabolism, Volume 61, Issue 5, 2012, Pages 611-619, ISSN 0026-0495, https://doi.org/10.1016/j.metabol.2011.10.005.

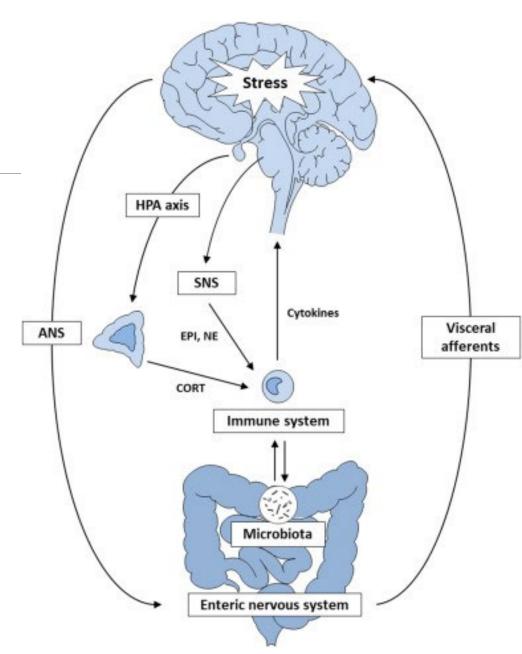
Fig. 2 – Biologic and behavioral pathways linking stress to obesity and the metabolic syndrome.

Cardiovascular

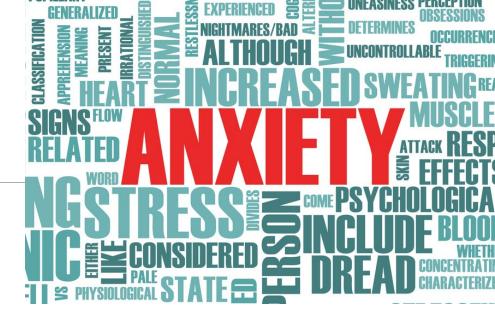
Disease, Clinical Events

Gastrointestinal system:

- Gut-brain axis
 - Interconnected bidirectional pathways between the brain and gut that are modulated by stress
 - Gut motility, immune function/microbiota
 - Functional gastrointestinal disorders (e.g., Irritable Bowel Syndrome) strongly linked to stress



Mental Health



- Anxiety
 - Stress is the response to a situation (external), anxiety is a reaction to stress (internal)
 - Prolonged exposure to stress hormones can contribute to the development of anxiety

Effects of stress

Mental Health:

- Post-Traumatic Stress Disorder (PTSD)
 - Intrusive thoughts, nightmares/flashbacks of trauma, avoidance, hypervigilance, sleep disturbances
 - Occurs in 5-10% of people exposed to traumatic events
 - Underlying cause unclear but thought to relate to dysregulation of HPA axis



Mental Health:



- Depression
 - Strong links between chronic/toxic stress and depression
 - Behavioural and biological pathways involved
 - Stress \rightarrow sedentary lifestyle, lack of sleep, poor self-care, relationship challenges
 - Stress \rightarrow HPA axis dysregulation
 - = increased risk of depression



Poll

True or False?

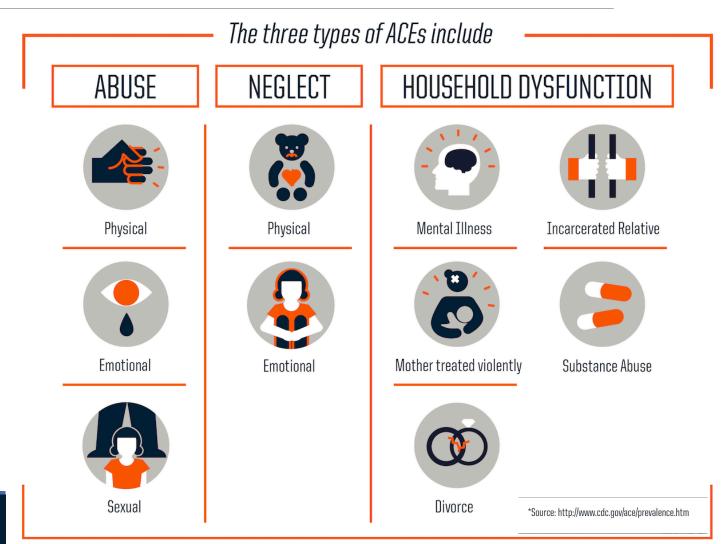
 Stressful/traumatic experiences in childhood can lead to longterm health consequences

Robert Wood Johnson Foundation

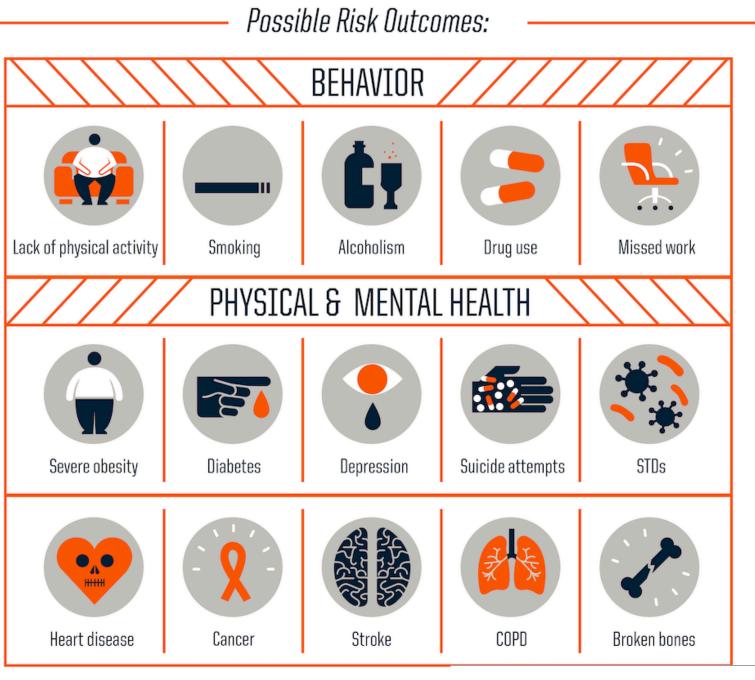
- Adverse Childhood
 Experiences (ACEs)
 - Abuse
 - Neglect

rwjf.org/aces

- Household dysfunction
- Higher number of ACES = higher risk of health problems*



- Adverse Childhood
 Experiences (ACEs)
 - Lead to toxic stress
 - Increase risk of negative behaviours and chronic health conditions



*Source: http://www.cdc.gov/ace/prevalence.htm



rwjf.org/aces



When to seek help

- Feeling overwhelmed
- Feeling worried about your level of stress and/or its impacts
- Using drugs or alcohol to cope
- Thoughts of hurting yourself

• Reach out to your primary care provider, mental health provider (counsellor, therapist, etc.), or crisis line

Preview: stress management

- Relaxation activities meditation, yoga, breathing exercises
- Exercise, eating well, sleep
- Positive attitude and gratitude
- Acceptance/reframing
- Social connection and talking



Suggested Resources

- Centre for Studies on Human Stress: https://humanstress.ca/stress/
- Canadian Mental Health Association: https://cmha.ca/
- Crisis Line Association of BC: https://www.crisislines.bc.ca/
 - 310Mental Health Support: **310-6789** (no area code needed)
- HealthLinkBC: Stress Management https://www.healthlinkbc.ca/health-topics/stress-management
- Telephone 8-1-1 (7-1-1 for the hearing impaired)
 - Speak to a Registered Nurse, Registered Dietician, exercise professional, or Pharmacist
- Dr. Gair book recommendation: Burnout, the Secret to Unlocking the Stress Cycle -Nagoski



• Questions or comments about the talk?

Closing Remarks

• Thank you to Dr. Jane Gair, our supervisor for this activity, and to my classmates, Julia De Pieri and Drew Smith, who will be delivering several of the talks in this series

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• Thank you for attending today!