

The Impact of Domestic Violence on Children's Brain Development

Peter S. Martin, MD, MPH

Director, Division of Community Psychiatry

Associate Director, Forensic Psychiatry Fellowship

Co-Director, GHHS Resident/Fellow Chapter

Assistant Professor of Psychiatry

University at Buffalo

Objectives

- Define the types of abuse
- Review background information on the effects of abuse, including domestic violence, to better understand short- and long-term consequences
- Describe the different brain regions that are directly, and indirectly, impacted by domestic violence
- Discuss how this knowledge can shape diagnostic and treatment recommendations

The challenges





Definitions

Definitions

- Child Maltreatment
 - Physical abuse
 - Physical neglect
 - Emotional abuse
 - Emotional neglect
 - Sexual abuse

Neglect and Abuse

- Neglect: parent/guardian has
 - Harmed the child
 - Failed to properly care for the child by providing inadequate education or medical care
 - Abandoned the child
- Abuse: parent/guardian injured the child
 - Physically
 - Sexually
 - Mentally
 - Emotionally

Background

(Not at all) Fun Facts about Abuse

- Annually
 - 702,000 children in US maltreated in 2009 (reported)¹
 - ~900,000 child victims of abuse/neglect (12/1,000 children)²
- CPS evaluations²: 30% substantiated → youngest children at highest risk
 - 60% neglect (2% medical neglect)
 - 16% physical
 - 8% psychological³

1. Klika, JB & Herrenkohl, TI. A review of developmental research on resilience in maltreated children.

2. Hayes, JP. Parenting Assessment in Abuse, Neglect and Permanent Wardship Cases.

3. Keyes KM et al. Childhood maltreatment and the structure of common psychiatric disorders.

(Not at all) Fun Facts about Abuse

- Costs: lifetime exceed \$210,000 per victim (law enforcement, child welfare, medical treatment/hospitalizations, loss of productivity)
 - Helps justify why financially you would want to prevent/fund prevention programs
 - Such as The Nurse Family Partnership, Positive Parenting Program (Triple P)

Background

- Children's brains primed to learn new experiences, but also makes them more susceptible to environmental stresses

Biological

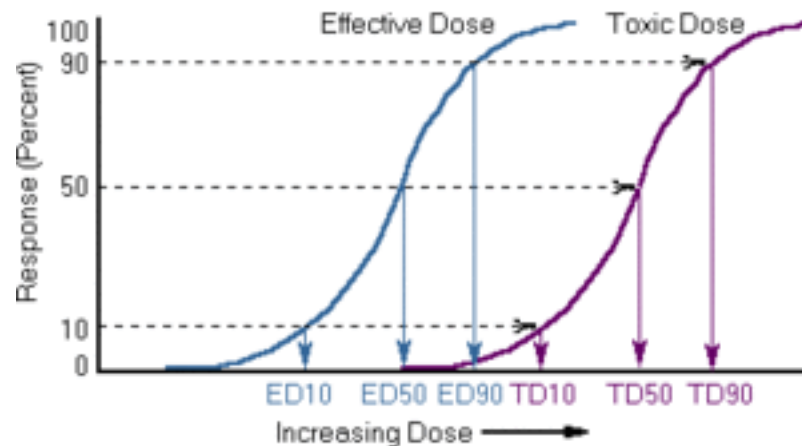
- Early Years Study 2: poor caregiver-infant interactions compromised formation of neural circuits and pathways¹
- Cortisol and catecholamines: ↑ with stress from abuse → destruction of brain cells, disrupts neuronal connections²

1. McCain NM, Mustard F, & Shanker S. Early Years Study 2. Putting science into action.

2. Odhayani, AA, Watson, WJ & Watson, L. Behavioral consequences of child abuse.

Background

- Clear dose–response relationship between severity of exposure and magnitude of the neurobiological findings



1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.
2. Dannlowski, U et al. Limbic scars: Long-term consequences of childhood maltreatment revealed by functional and structural magnetic resonance imaging.
3. Edmiston, EE et al. Corticostriatal-limbic gray matter morphology in adolescents with self-reported exposure to childhood maltreatment.
4. Teicher, MH, Anderson, CM, & Polcari, A. Childhood maltreatment is associated with reduced volume in the hippocampal subfields CA3, dentate gyrus, and subiculum.
5. Treadway, MT et al. Early adverse events, HPA activity and rostral anterior cingulate volume in MDD.

Background

- Survivors of childhood maltreatment have a higher prevalence of mental health/cognitive effects:¹⁻⁷
 - Depression
 - Anxiety
 - Substance abuse
 - Eating disorders
 - Suicidal symptomatology
 - Psychosis
 - Personality disorder
 - Diminished cognitive functioning
 - Poorer treatment response
- Maltreatment accounted for **45%** of the population attributable risk for childhood onset psychiatric disorders⁸

1. Ball, JS & Links, PS. Borderline personality disorder and childhood trauma: Evidence for a causal relationship.
2. de Bellis, MD et al. Neuropsychological findings in childhood neglect and their relationships to pediatric PTSD.
3. Bendall, S et al. Childhood trauma and psychotic disorders: A systematic, critical review of the evidence.
 4. Gould, F et al. The effects of child abuse and neglect on cognitive functioning in adulthood.
5. Nanni, V, Uher, R & Danese, A. Childhood maltreatment predicts unfavorable course of illness and treatment outcome in depression: A meta-analysis.
6. Norman, RE et al. The long-term health consequences of child physical abuse, emotional abuse, and neglect: A systematic review and meta-analysis.
7. Teicher, MH & Samson, JA. Childhood maltreatment and psychopathology: A case for ecophenotypic variants as clinically and neurobiologically distinct subtypes.
8. Green, JG et al. Childhood adversities and adult psychiatric disorders in the national comorbidity survey replication I: Associations with first onset of DSM-IV disorders.

Background

- Survivors of childhood maltreatment have increased prevalence of physical health disorders:
 - Inflammation¹
 - Metabolic syndrome²
 - Arthritis³
 - Ischemic heart disease⁴
 - Cancer⁵
 - Shortened telomeres associated with reduced life expectancy^{6,7}

1. Danese, A et al. Childhood maltreatment predicts adult inflammation in a life-course study.

2. Danese, A et al. Adverse childhood experiences and adult risk factors for age-related disease: Depression, inflammation, and clustering of metabolic risk markers.

3. Spitzer, C et al. Gender-specific association between childhood trauma and rheumatoid arthritis: A case-control study.

4. Dong, M et al. Insights into causal pathways for ischemic heart disease: Adverse childhood experiences study.

5. Brown, DW et al. Adverse childhood experiences are associated with the risk of lung cancer: A prospective cohort study.

6. Price, LH et al. Telomeres and early-life stress: An overview.

7. Brown, DW et al. Adverse childhood experiences and the risk of premature mortality.

The Brain

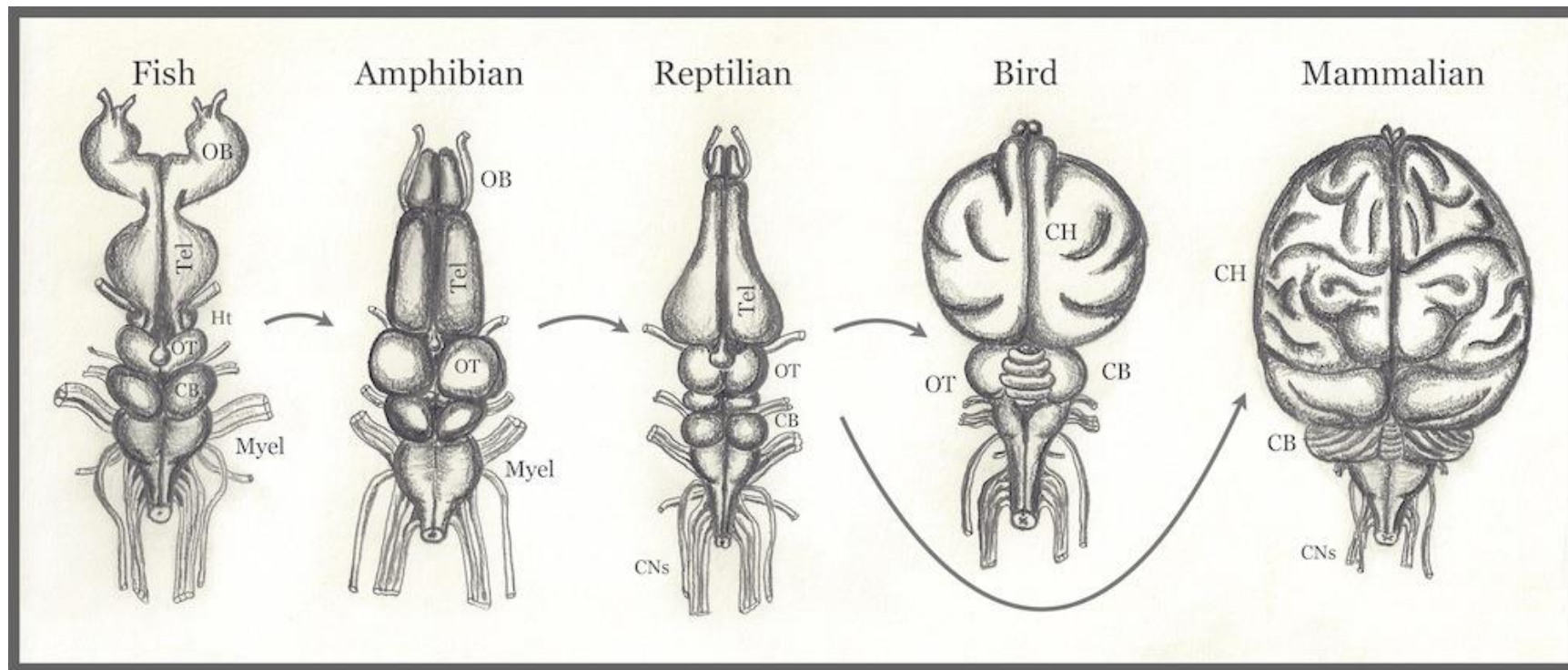
Background

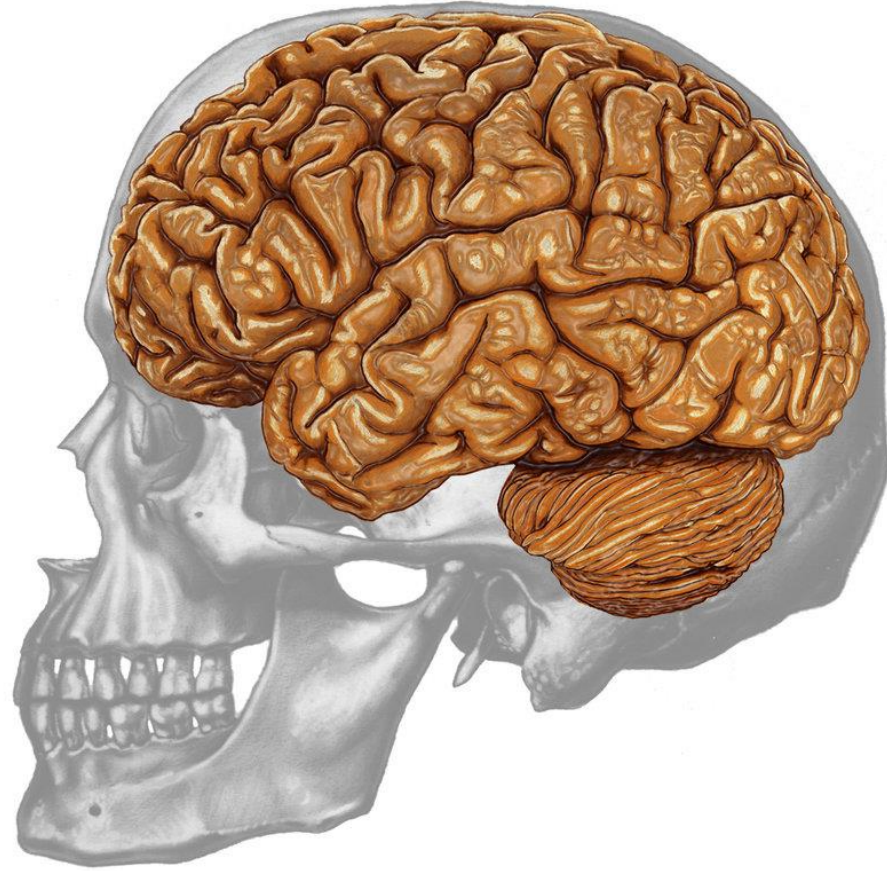
- Clearly able to show: link between maltreatment, development and alterations in structure and function of stress-susceptible brain regions
 - Substantial alterations in connectivity and network architecture
- Less clear: how these changes directly link to psychopathology

Background

- Chronic stress:
 - Potentially adaptive changes of brain → can lead to psychopathology when adaptive responses/experiences so much differ as to become maladaptive

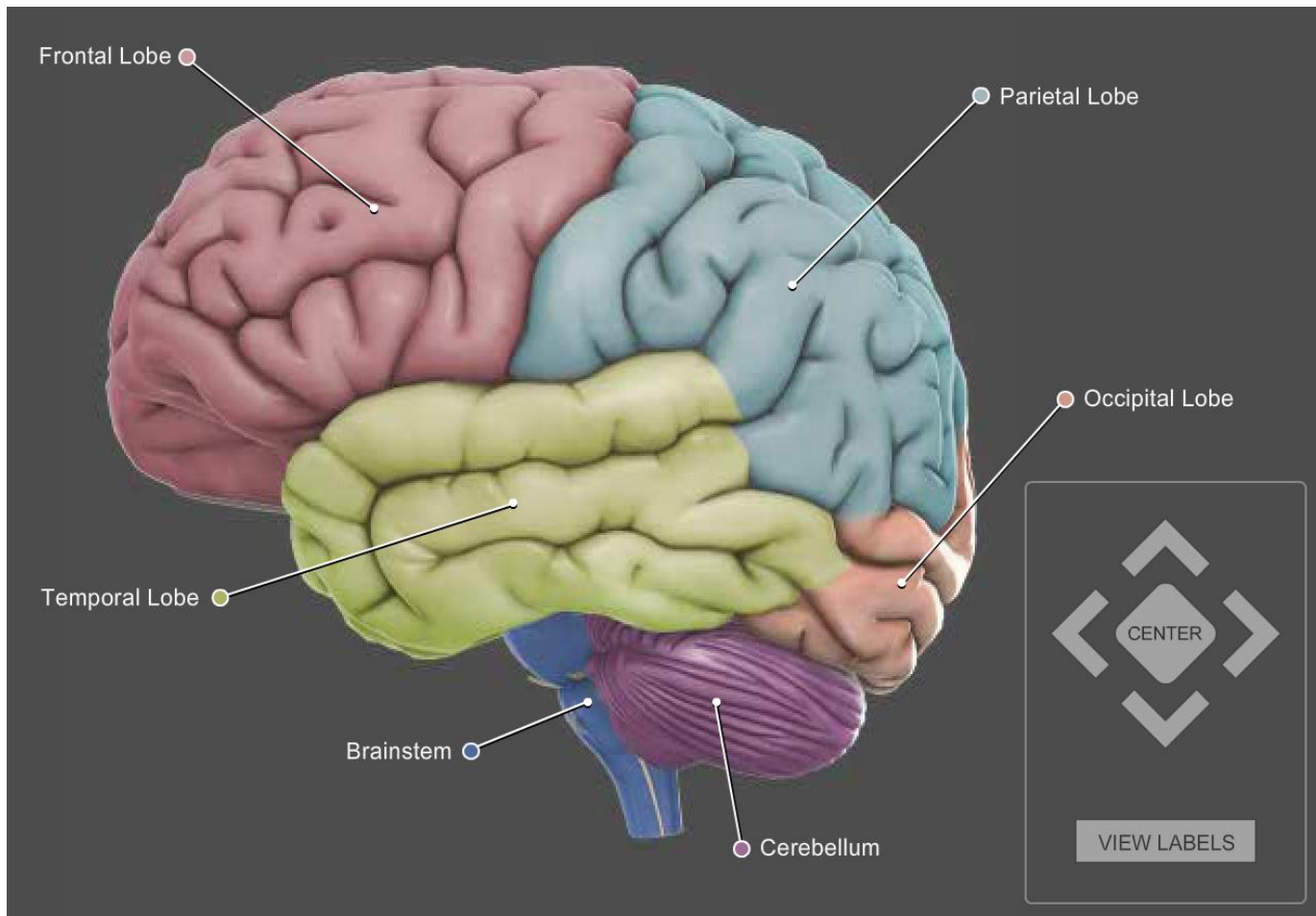
The Brain (101)



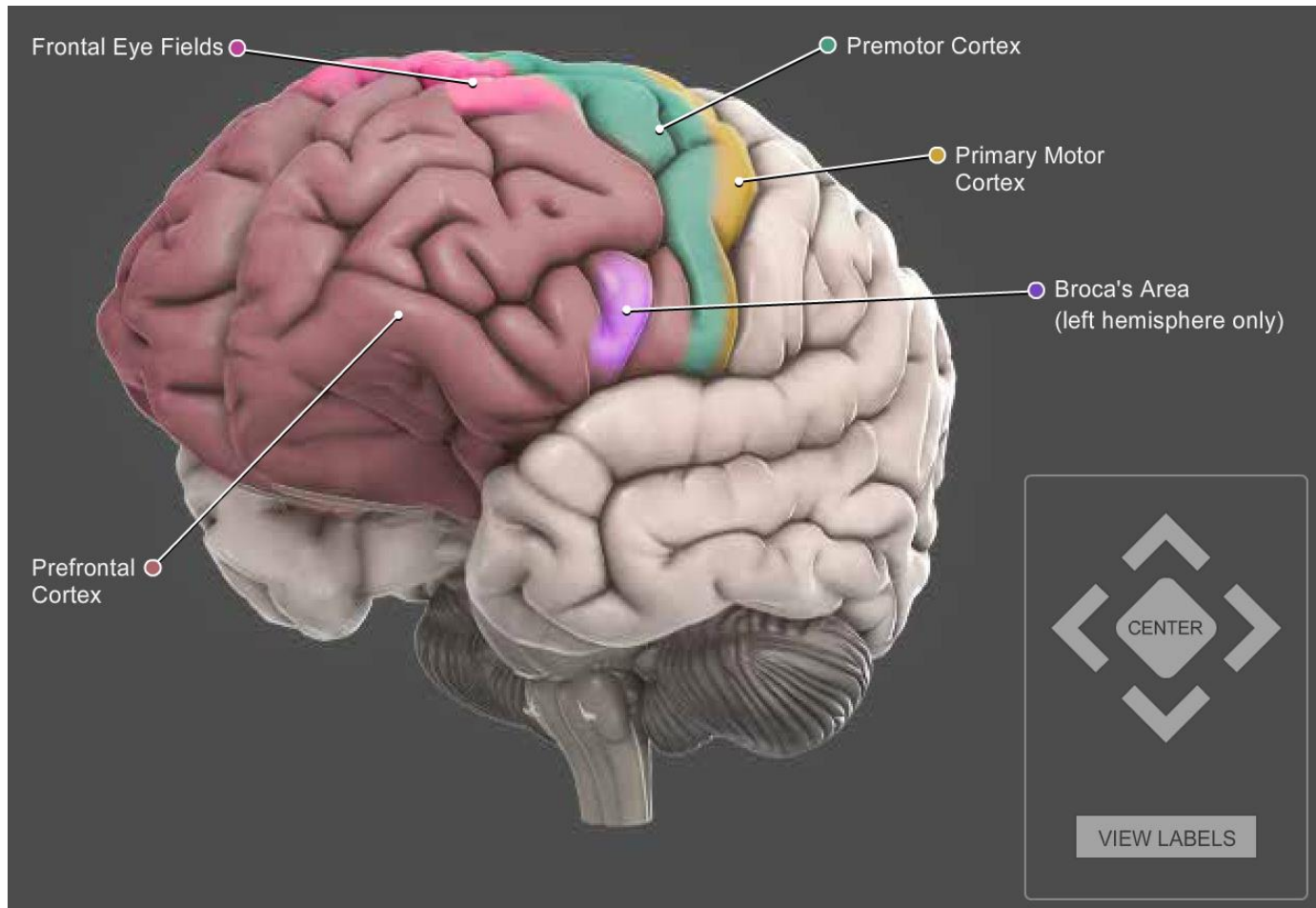


Marc Gosselin

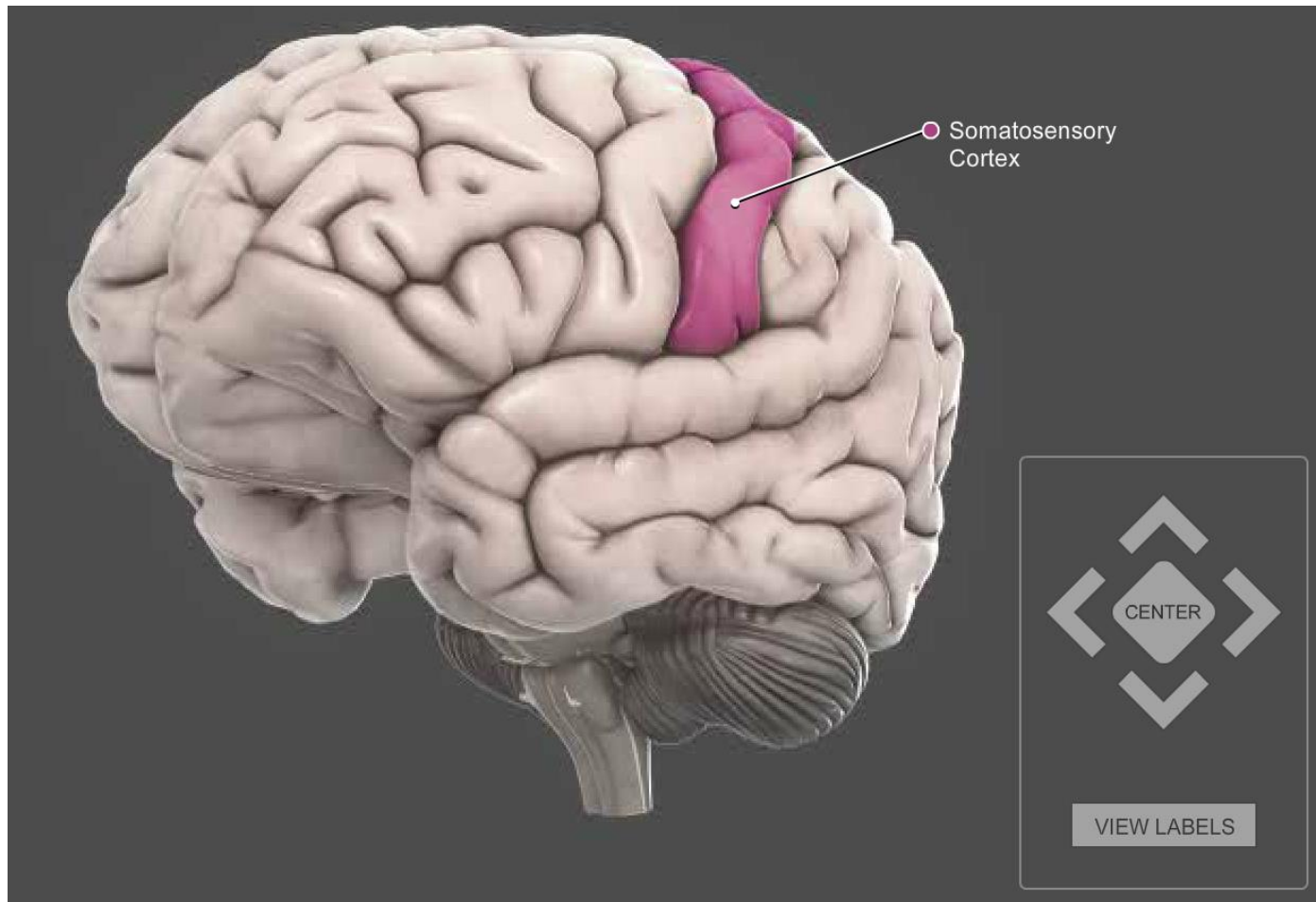
Whole Brain



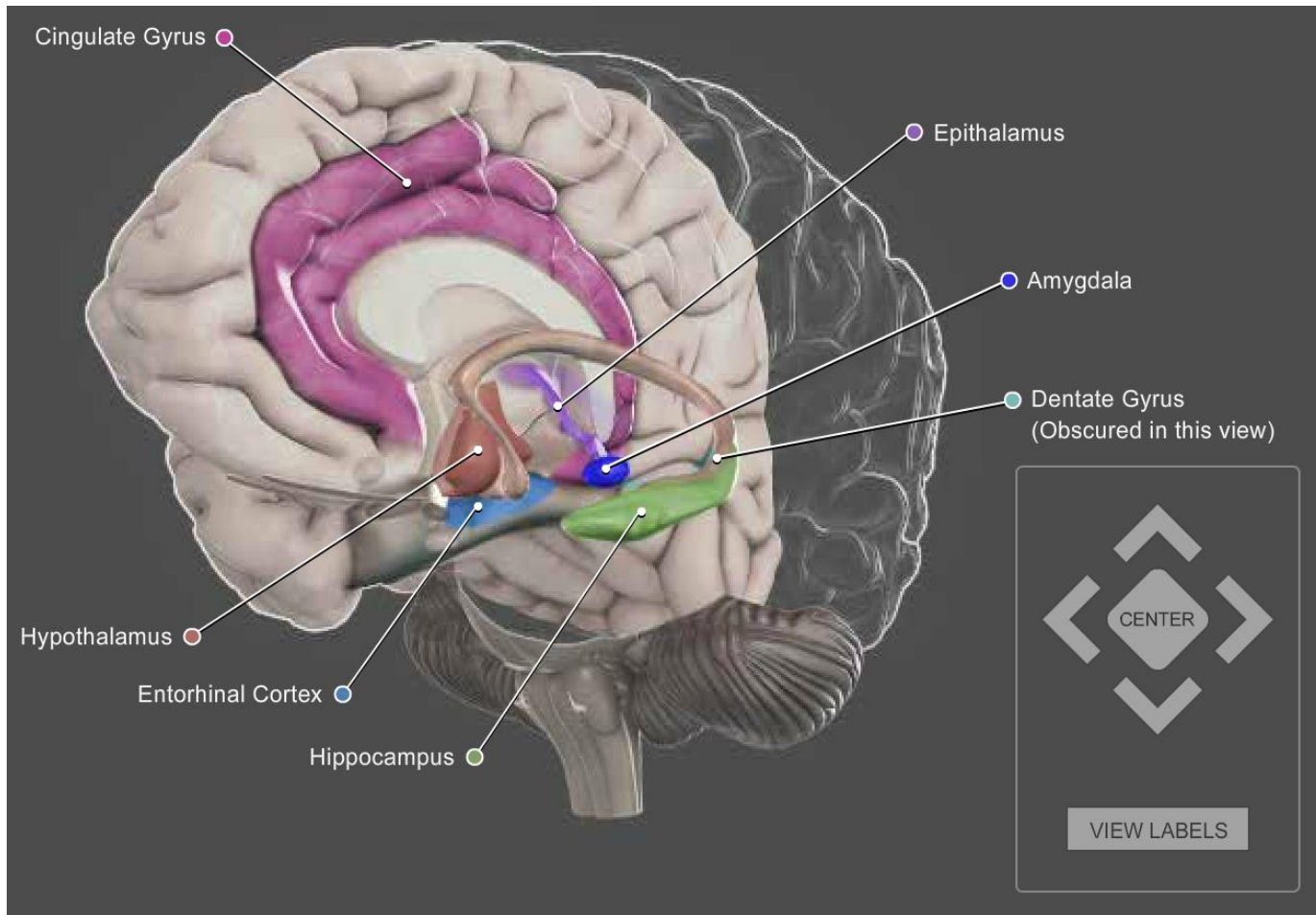
Frontal Lobe



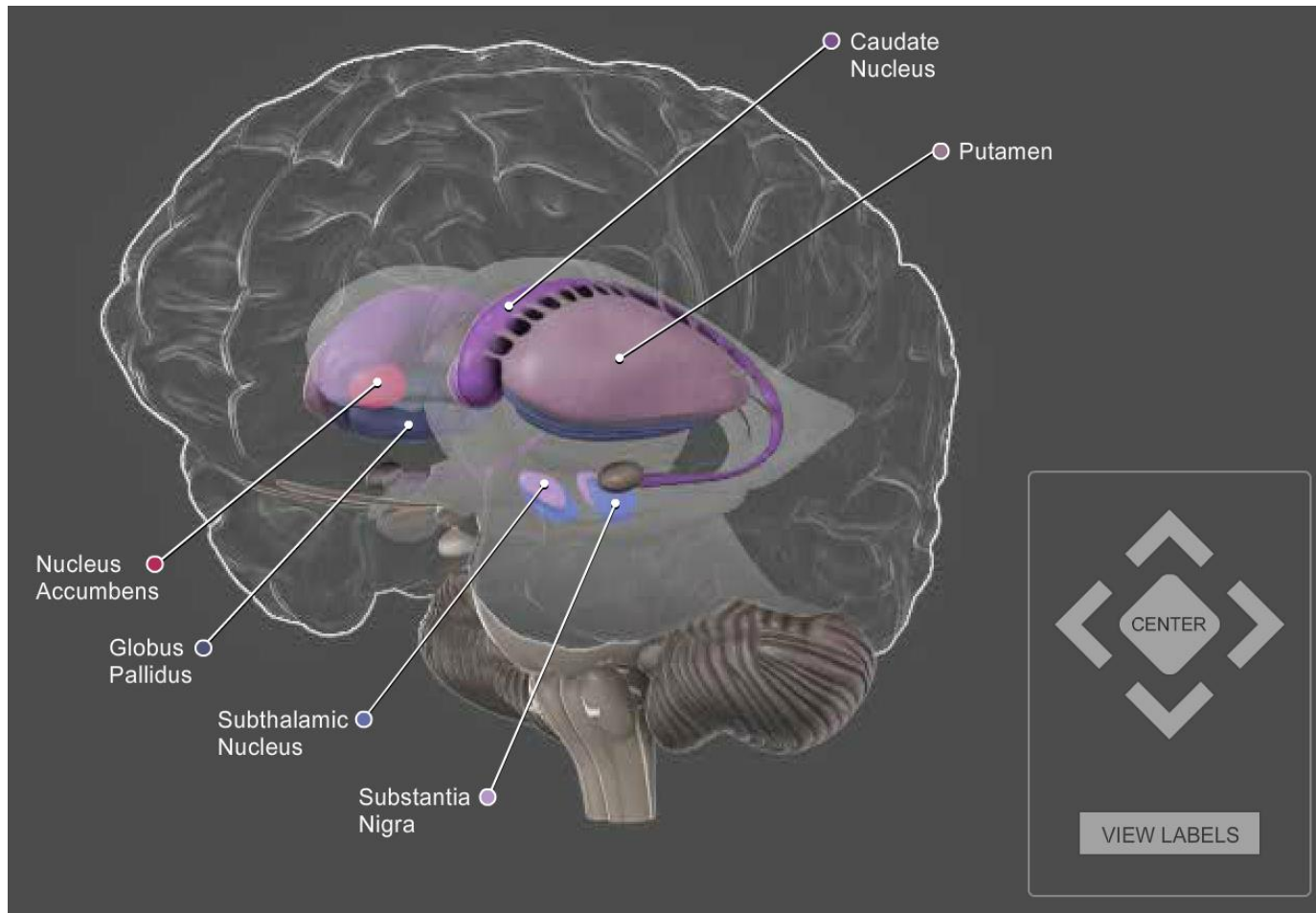
Somatosensory Cortex



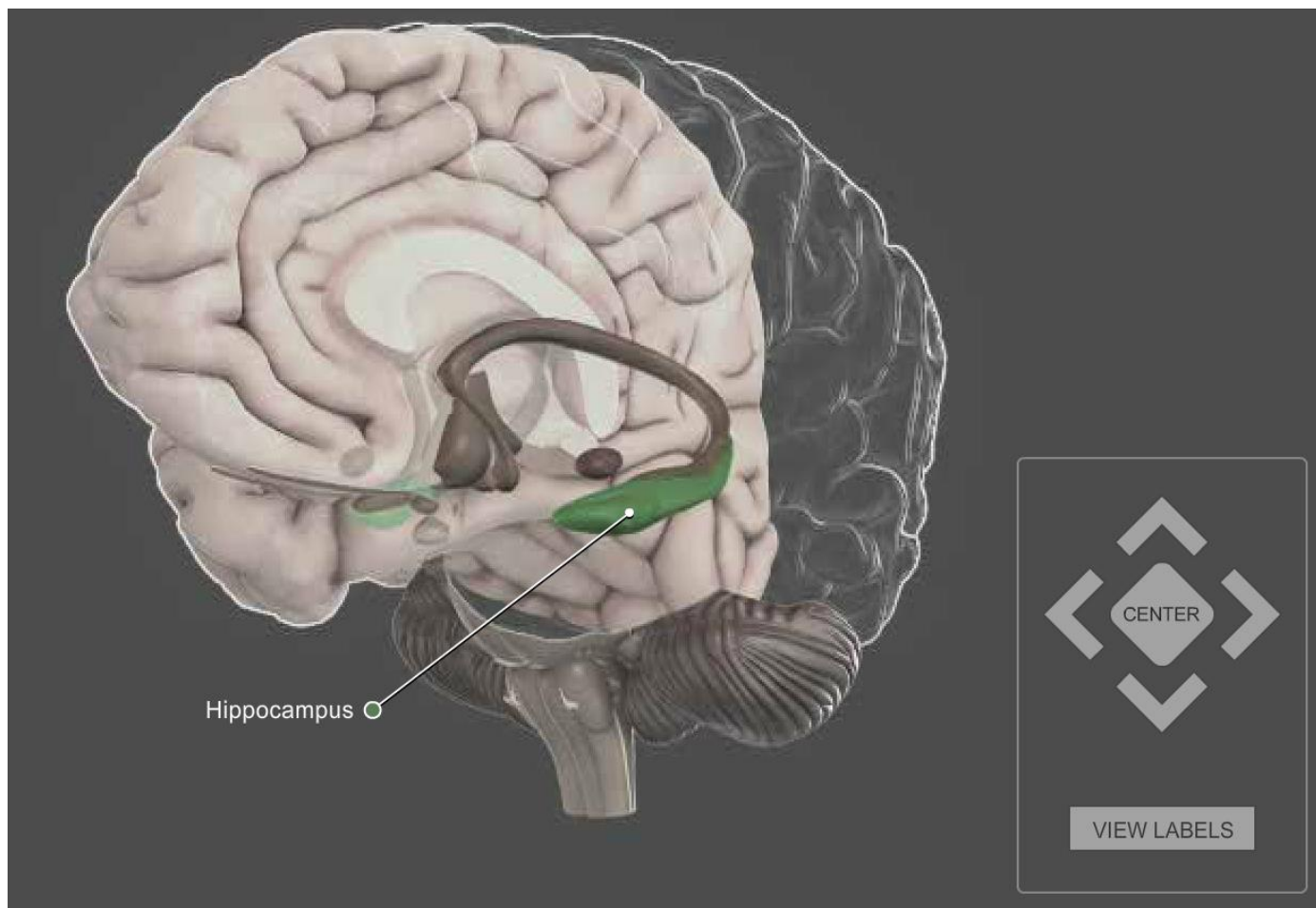
Limbic System



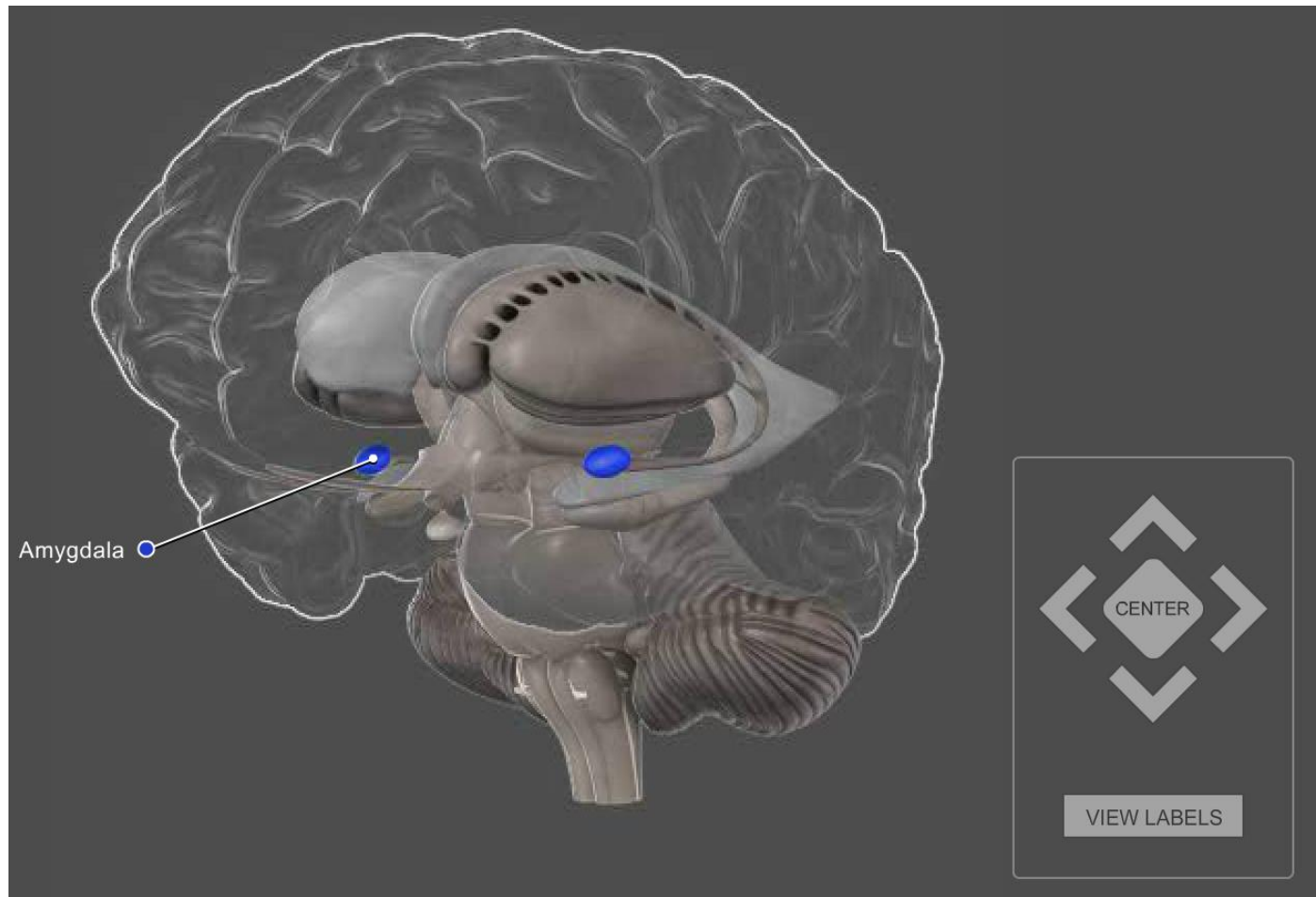
Basal Ganglia (Striatum)



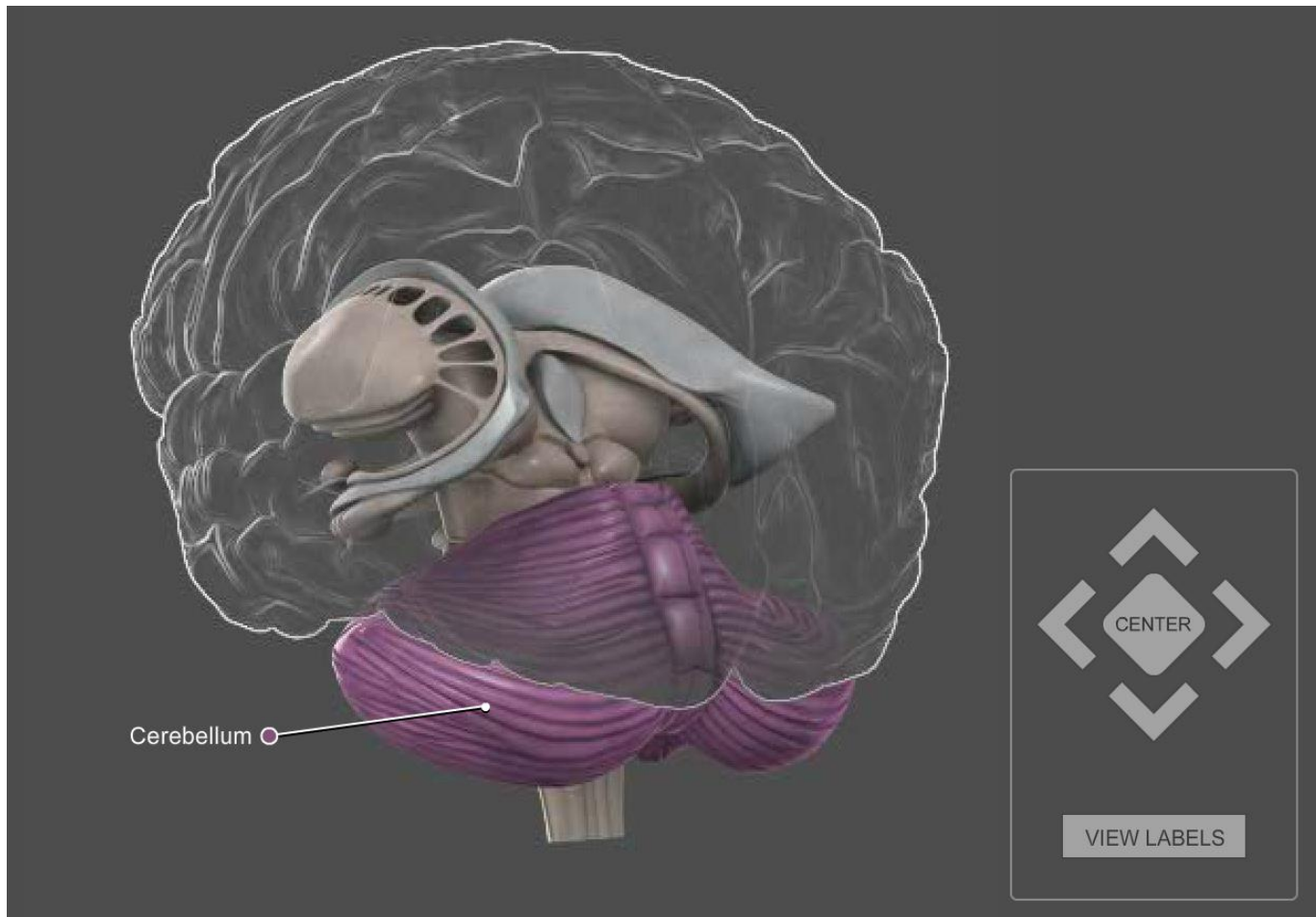
Hippocampus



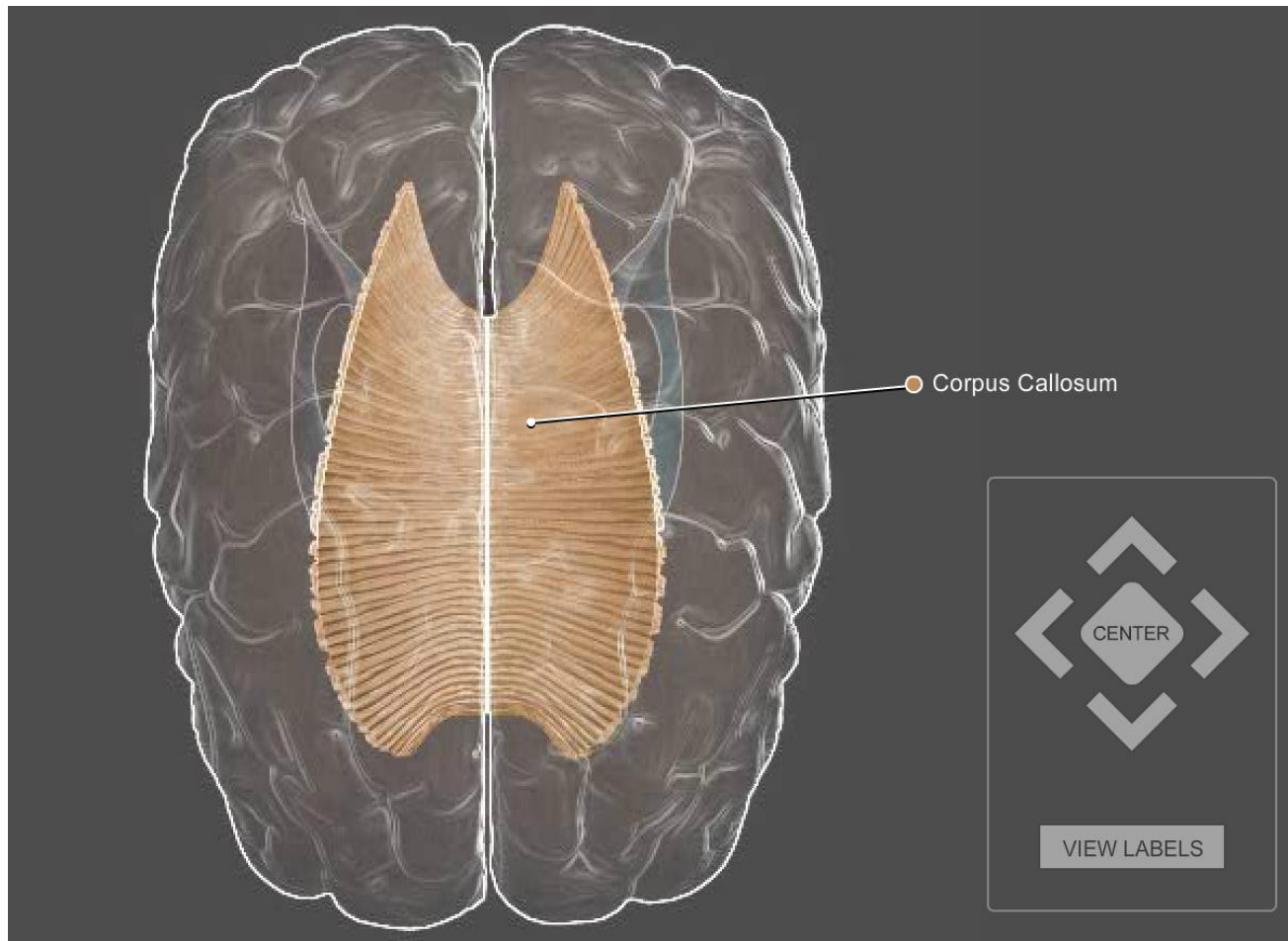
Amygdala



Cerebellum

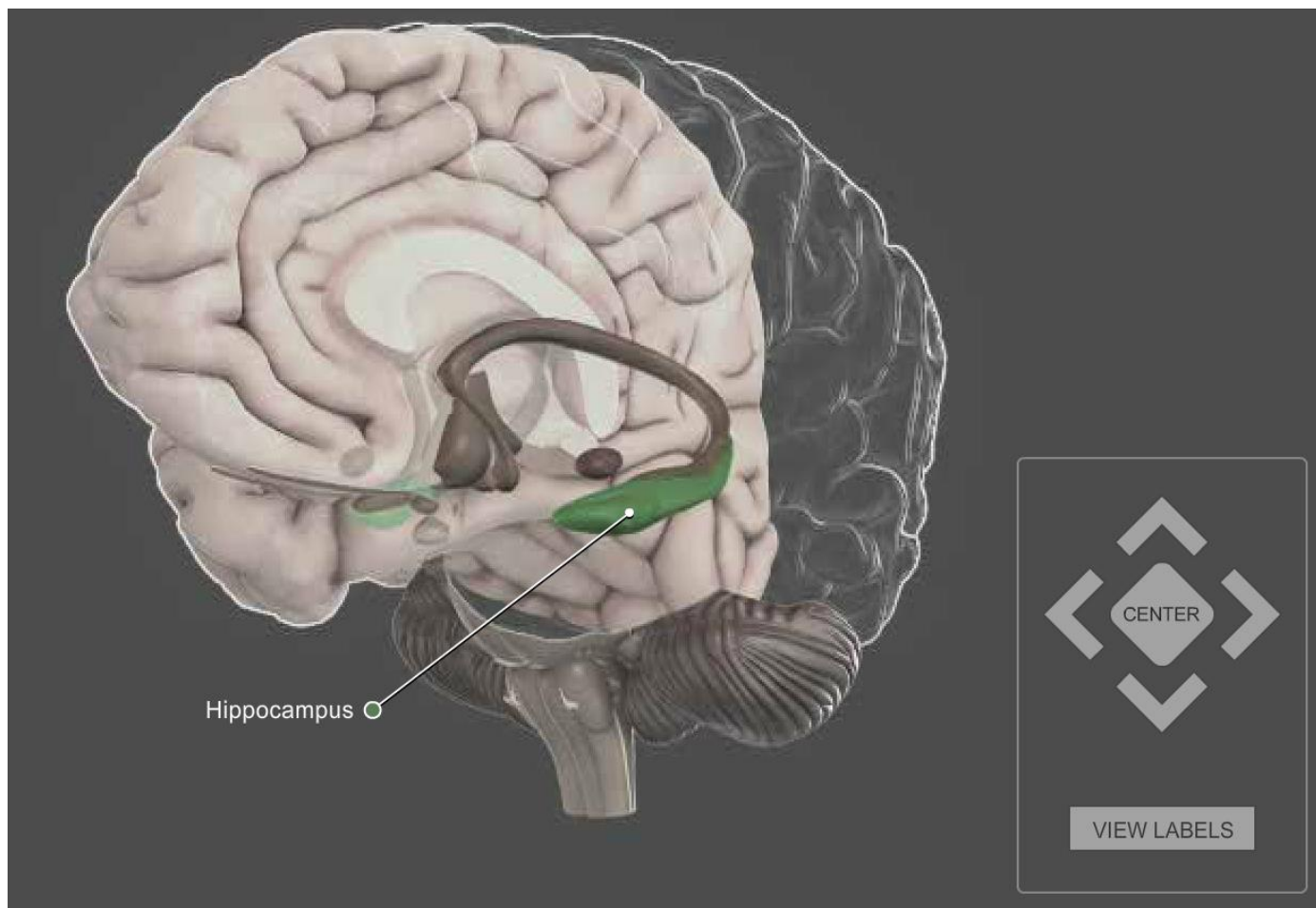


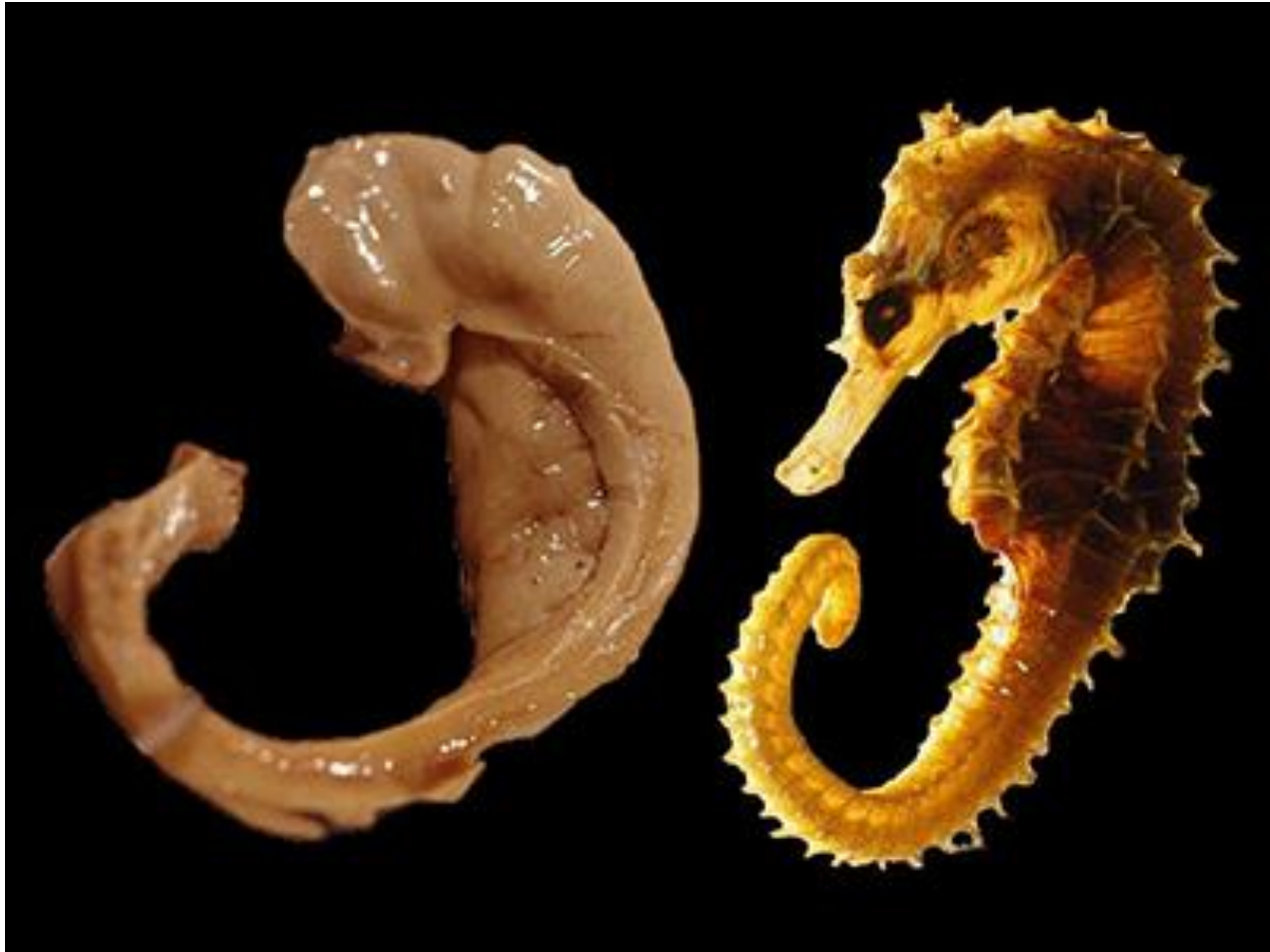
Corpus Callosum



Specific Brain Regions

Hippocampus





Hippocampus

- Involved in the formation and retrieval of **memories**, including **autobiographical memories**¹
 - *Place cells* provide an internal positioning system for the spatiotemporal representation of places, routes and associated experiences²
- Most obvious target for potential effects of childhood maltreatment³

1. Nadel, L, Campbell, J, & Ryan, L. Autobiographical memory retrieval and hippocampal activation as a function of repetition and the passage of time.

2. Moser, EI, Kropff, E & Moser, MB. Place cells, grid cells, and the brain's spatial representation system.

3. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

Hippocampus

- Densely populated with glucocorticoid receptors¹ → **highly susceptible to damage** from excessive levels of glucocorticoids (ex. **cortisol**)²

1. Morimoto, M et al. Distribution of glucocorticoid receptor immunoreactivity and mRNA in the rat brain: An immunohistochemical and in situ hybridization study.

2. Sapolsky, RM, Krey, LC, & McEwen, BS. Prolonged glucocorticoid exposure reduces hippocampal neuron number: Implications for aging.

Hippocampus

- Adults: **smaller** hippocampi in maltreated vs. nonmaltreated¹
 - But smaller effect in females
 - Possibly because of neuroprotective effects of **estrogen**²

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. McEwen, BS. . Stress, sex, and neural adaptation to a changing environment: Mechanisms of neuronal remodeling. Andersen, SL &

Hippocampus

- Childhood/adolescence: evidence less strong of effects¹
 - Might not show until **adolescence** → but studies have commonly combined both groups
 - May have “silent period” between childhood and adolescence with no discernable neurobiological differences²

1. McEwen, BS. . Stress, sex, and neural adaptation to a changing environment: Mechanisms of neuronal remodeling.

2. Andersen, SL & Teicher, MH. Delayed effects of early stress on hippocampal development.

Hippocampus

- May have potential **sensitive exposure periods**¹
 - Most significant: exposure between **3-5 years old**; less degree at 11-13 years old²
 - Early separation stress has much greater effects on synaptic density in hippocampus³
 - Specifically associated with **reduction in volume** of portions of hippocampal that have been shown to be susceptible in both laboratory animals and humans^{1,4-6}

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Andersen, SL et al. Preliminary evidence for sensitive periods in the effect of childhood sexual abuse on regional brain development.

3. Andersen, SL & Teicher, MH. Delayed effects of early stress on hippocampal development

4. Andersen, SL & Teicher, MH. Stress, sensitive periods and maturational events in adolescent depression.

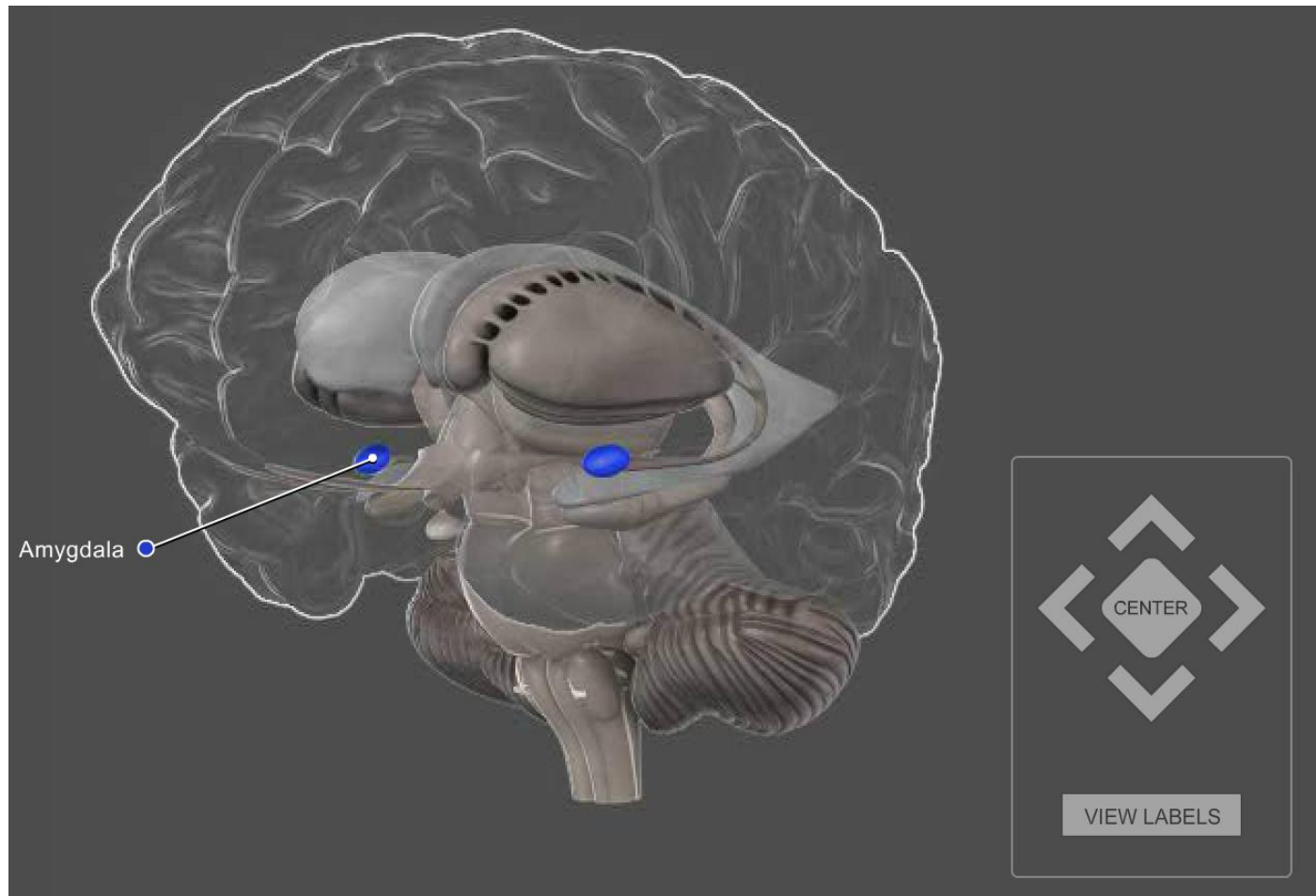
5. Pagliaccio, D et al. Stress-system genes and life stress predict cortisol levels and amygdala and hippocampal volumes in children.

6. Teicher, MH, Anderson, CM, & Polcari, A. Childhood maltreatment is associated with reduced volume in the hippocampal subfields CA3, dentate gyrus, and subiculum.

Hippocampus

- Smaller hippocampal volumes:
 - Mild cognitive impairment/Traumatic brain injury
 - Schizophrenia
 - Major depression
 - Posttraumatic stress disorder
 - Obsessive–compulsive disorder (OCD)
 - Chronic alcoholism
 - Antisocial personality disorder
 - Borderline personality disorder
- Larger hippocampal volumes:
 - Autism
- Preservation of hippocampal volume:
 - Bipolar disorder
 - Anorexia nervosa
 - Panic disorder
 - Attention-deficit/hyperactivity disorder (ADHD)

Amygdala





the amygdala

Amygdala

- Encoding of **implicit emotional memories**¹
- Detecting and responding to **salient stimuli** – ex. facial expressions and potential threats²

1. LeDoux, JE. Emotional memory systems in the brain. *Behavioural Brain Research*. 1993; 58; 69–79

2. Derntl, B et al. General and specific responsiveness of the amygdala during explicit emotion recognition in females and males. *BMC Neuroscience*. 2009; 10: 91

Amygdala

- High density of glucocorticoid receptors on stress-susceptible pyramidal cells¹
- Postnatal developmental trajectory characterized by rapid initial growth → more sustained growth to peak volumes between 9-11 years → gradual pruning thereafter²
- Also **highly susceptible** to exposure to early stress³

1. Sarrieau, A et al. Autoradiographic localization of glucocorticosteroid and progesterone binding sites in the human post-mortem brain.

2. Uematsu, A et al. Developmental trajectories of amygdala and hippocampus from infancy to early adulthood in healthy individuals.

3. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

Amygdala

- Both psychological stressors and stress hormones → stimulate cells → **increase** in volume [hypertrophy]^{1,2}
 - Opposite of effects on hippocampus³
- Hypertrophy, unlike hippocampal hypotrophy, **endures** long after cessation of the stressor⁴

1. Mitra, R et al. Stress duration modulates the spatiotemporal patterns of spine formation in the basolateral amygdala.

2. Vyas, A, Jadhav, S & Chattarji, S. Prolonged behavioral stress enhances synaptic connectivity in the basolateral amygdala.

3. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

4. Vyas, A, Pillai, AG & Chattarji, S. Recovery after chronic stress fails to reverse amygdaloid neuronal hypertrophy and enhanced anxiety-like behavior

Amygdala

- Studies **do not** consistently show changes in maltreatment¹
 - If anything, majority showed a decrease
 - For decrease: had multiple forms of maltreatment; older; worse psychopathology
- Those studies that did show increase: **early** emotional and/or physical neglect

Amygdala

- **Early exposure** to stress sensitizes amygdala
→ **volume reduction** with subsequent exposure to stress (ex. early childhood stress, then later exposed to combat → increased risk to develop **PTSD**¹
- Longer-term study:²
 - **Early** maltreatment → increase in volume
 - **Later** exposure → decrease in volume

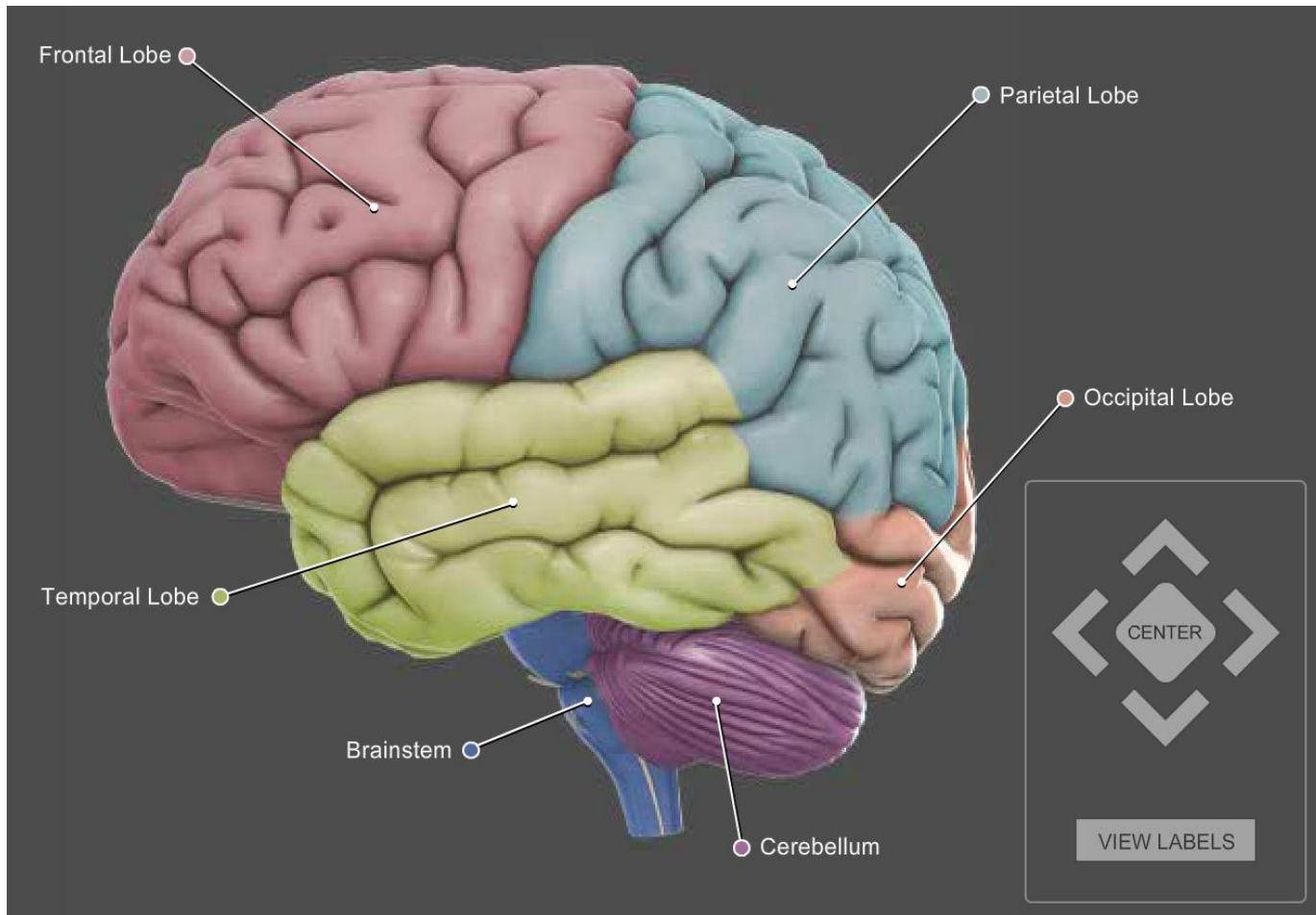
1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Whittle, S et al. Childhood maltreatment and psychopathology affect brain development during adolescence.

Amygdala

- Therefore:
 - **Early** exposure to maltreatment or neglect → **initial increase** in amygdala volume (particularly noticeable during **childhood**)
 - Early exposure may also **sensitize** the amygdala to further stress → result in a substantial **decrease** in amygdala volume (most noticeable in late **adolescence or adulthood**)

Cerebral Cortex



Cerebral Cortex

- Population of stress-susceptible (pyramidal) cells with a high density of glucocorticoid receptors → peaks during **late adolescence/early adulthood**¹
- Glucocorticoid receptors on (glial) cells that are most densely distributed during the neonatal period → gradually **decline**²

1. Sinclair, D et al. Dynamic molecular and anatomical changes in the glucocorticoid receptor in human cortical development.

2. Teicher, MH & Samson, JA. Annual research review: Enduring neurobiological effects of childhood abuse and neglect.

Cerebral Cortex

- Therefore: **two periods** of heightened stress sensitivity¹
 - 1) Infancy to early childhood
 - **Sensory** and **motor** cortical regions develop earlier²
 - 2) Late adolescence to early adulthood³
 - Protracted developmental course of **prefrontal** cortical regions

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Lenroot, RK et al. Differences in genetic and environmental influences on the human cerebral cortex associated with development during childhood and adolescence.

3. Sarrieau, A et al. Autoradiographic localization of glucocorticosteroid and progesterone binding sites in the human post-mortem brain.

Cerebral Cortex

- Overall effects (non region specific)
- Region specific
 - Higher order associations or polysensory cortex
 - Primary and secondary sensory cortex

Cerebral Cortex



Cerebral Cortex

- Overall

- From

- Ear

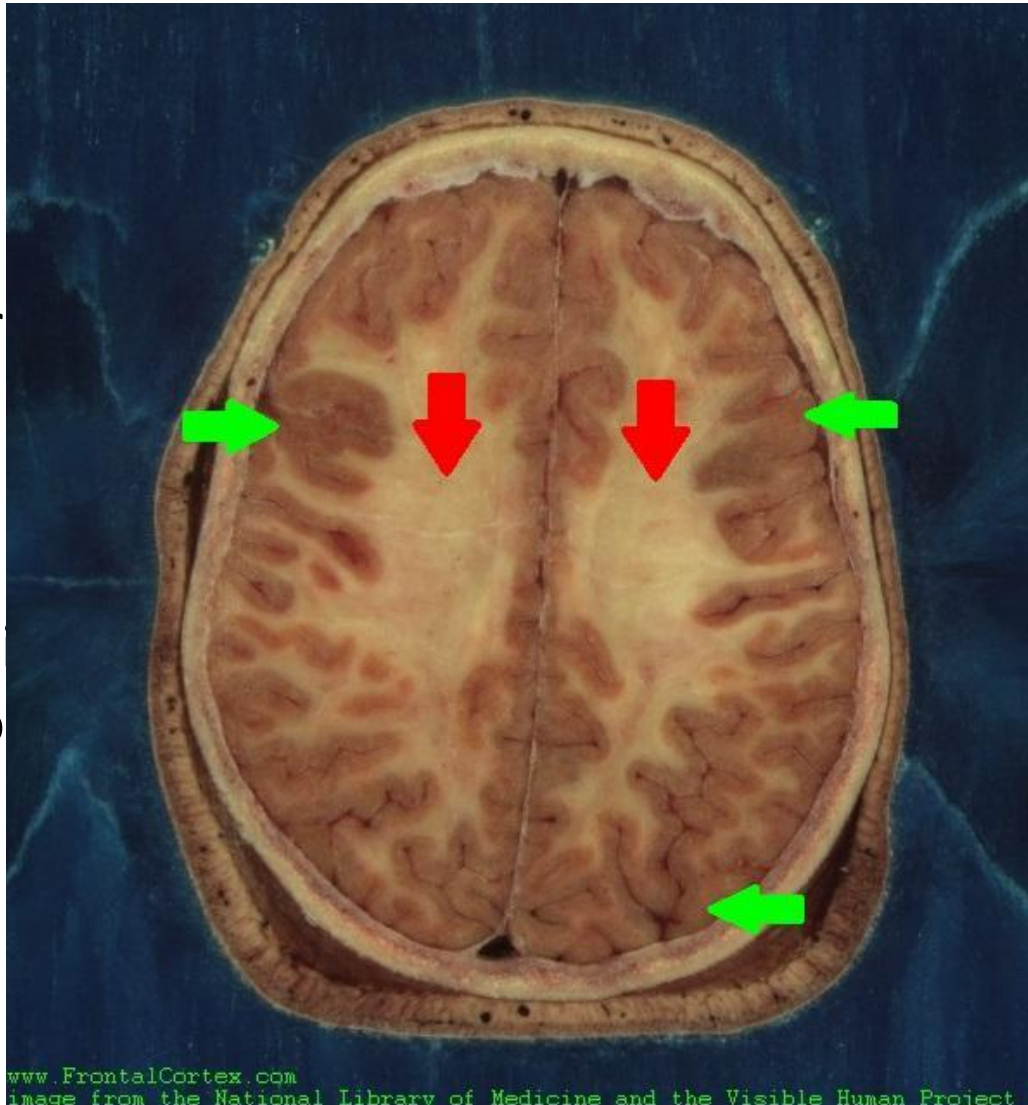
- -

- Ide

- ass

- mo

- -



ct¹
al care:

mly
en 7-33

and adopted

Cerebral Cortex

- Overall effects (non region specific)
 - Other studies have shown similar deleterious effects to children exposed to physical abuse, sexual abuse or witnessing domestic violence¹⁻³

1. Carrion, VG et al.. Attenuation of frontal asymmetry in pediatric posttraumatic stress disorder.

2. De Bellis, MD et al. Developmental traumatology. Part II: Brain development.

3. De Bellis, MD et al. A pilot longitudinal study of hippocampal volumes in pediatric maltreatment-related posttraumatic stress disorder.

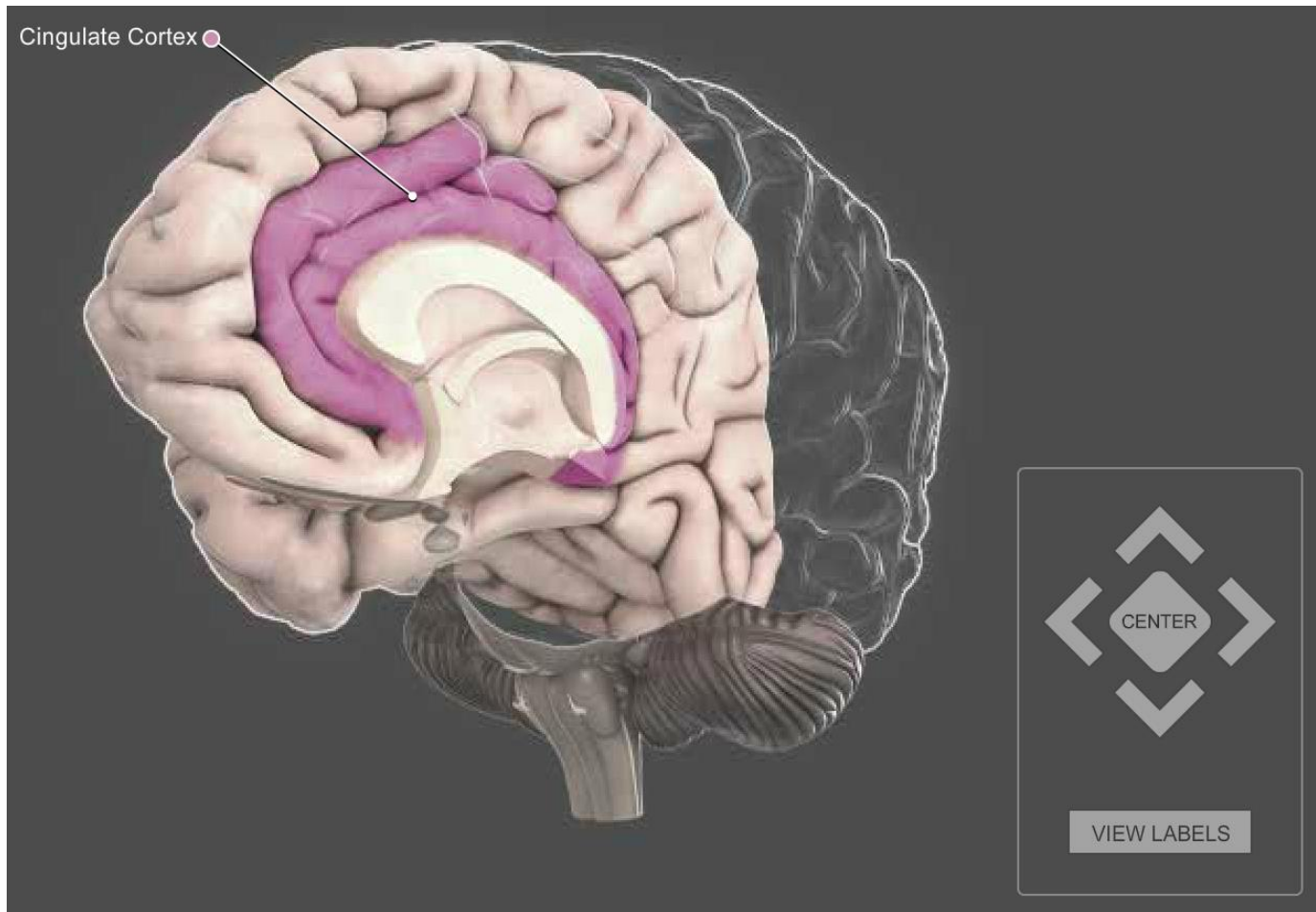
Cerebral Cortex

- Region specific
 - Higher order associations or polysensory cortex^{1,2}
 - Attenuated development of the **anterior cingulate cortex (ACC)** was the most consistent findings → most have reductions on both sides
 - Attenuated **dorsolateral prefrontal cortex**
 - Attenuated **orbitofrontal cortex**
 - Total prefrontal grey matter volume appeared to be most sensitive to maltreatment **between 14-16 years old**

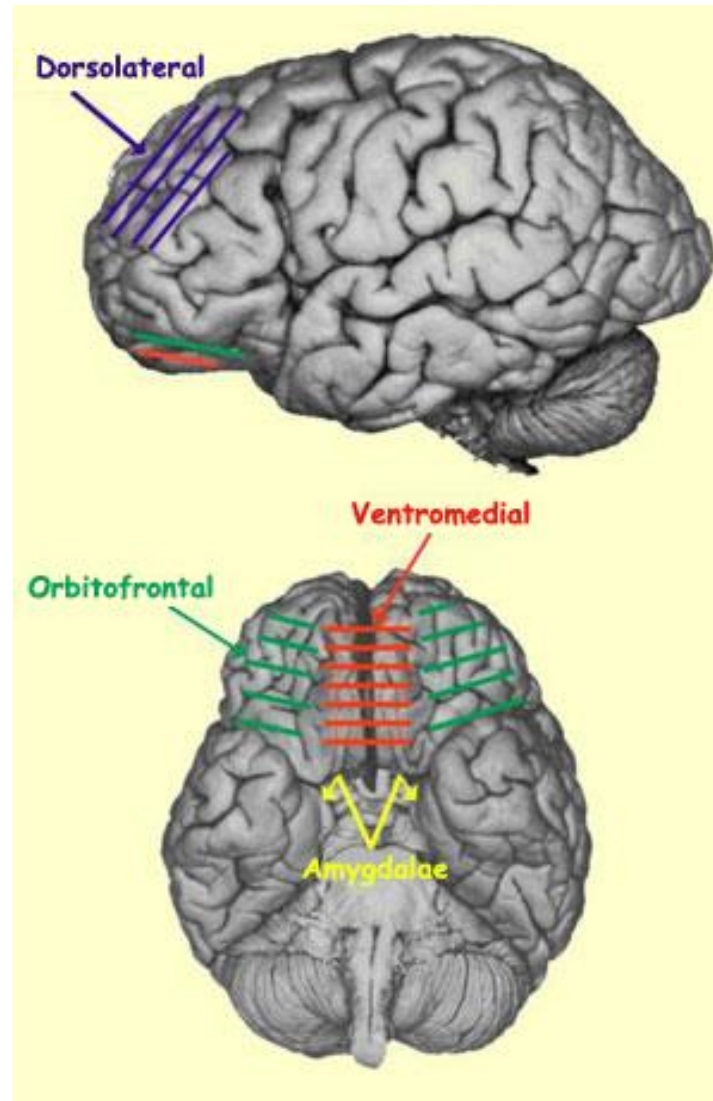
1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Andersen, SL & Teicher, MH. Stress, sensitive periods and maturational events in adolescent depression

Cingulate Cortex



Prefrontal Cortex



The Brain From Top to Bottom

Cerebral Cortex

- Region specific
 - Higher order associations or polysensory cortex
 - These three regions: important role in **decision-making** and **emotional regulation**¹
 - Have a role in **addiction**² → therefore, maltreatment may lead to brain changes that increase risk in addiction
 - Changes in those with a history of sexual abuse and psychotic disorder³

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Koob, GF & Volkow, ND. Neurocircuitry of addiction.

3. Sheffield, JM et al. Reduced gray matter volume in psychotic disorder patients with a history of childhood sexual abuse.

Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex
 - Most reliable correlates of exposure to **severe parental verbal abuse**:¹
 - Grey matter volume alterations in left **auditory** cortex
 - Diminished integrity of the left arcuate fasciculus **language** pathway

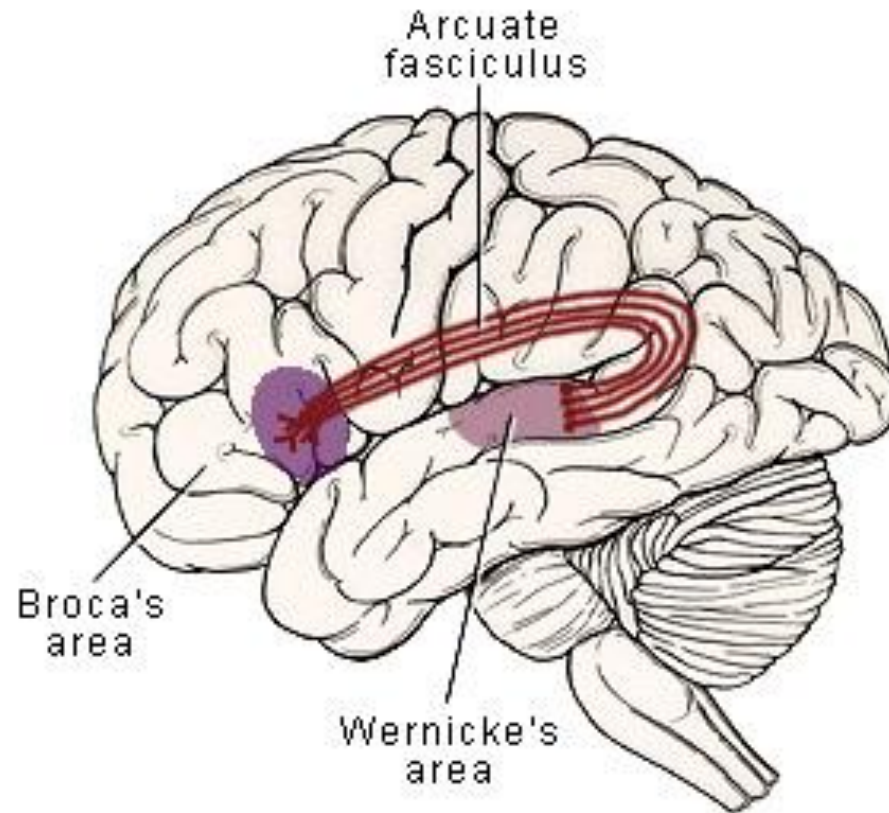
Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex¹
 - 3 fiber tracts²
 - Left arcuate fasciculus: Critically involved in human language³ with **lower verbal comprehension** and **verbal IQ** scores
 - Left cingulum bundle: Correlated with **depressive** and **dissociative** symptoms
 - Left fornix: Associated with symptoms of **anxiety** and **somatization**

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

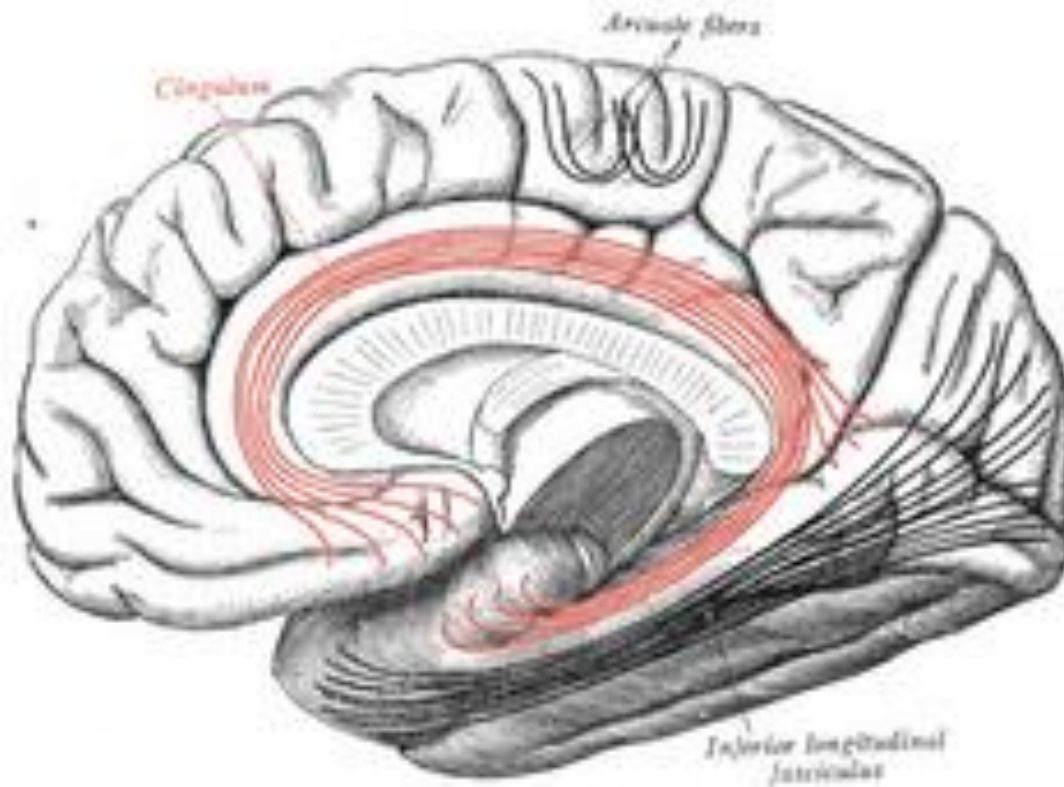
2. Choi, J et al.. Preliminary evidence for white matter tract abnormalities in young adults exposed to parental verbal abuse.

3. Rilling, JK et al. The evolution of the arcuate fasciculus revealed with comparative DTI.



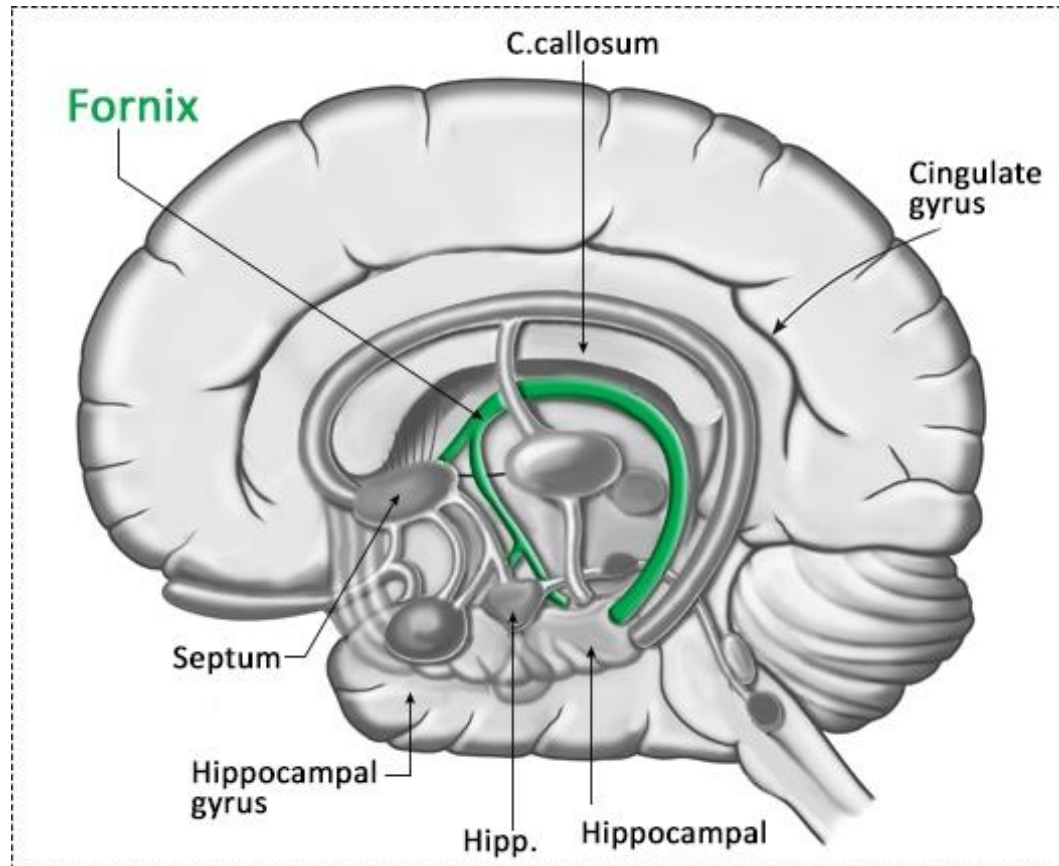
Arcuate Fasciculus

Critically involved in human language with **lower verbal comprehension** and **verbal IQ** scores



Cingulum Bundle

Correlated with **depressive** and **dissociative** symptoms



Fornix

Associated with symptoms of **anxiety** and **somatization**

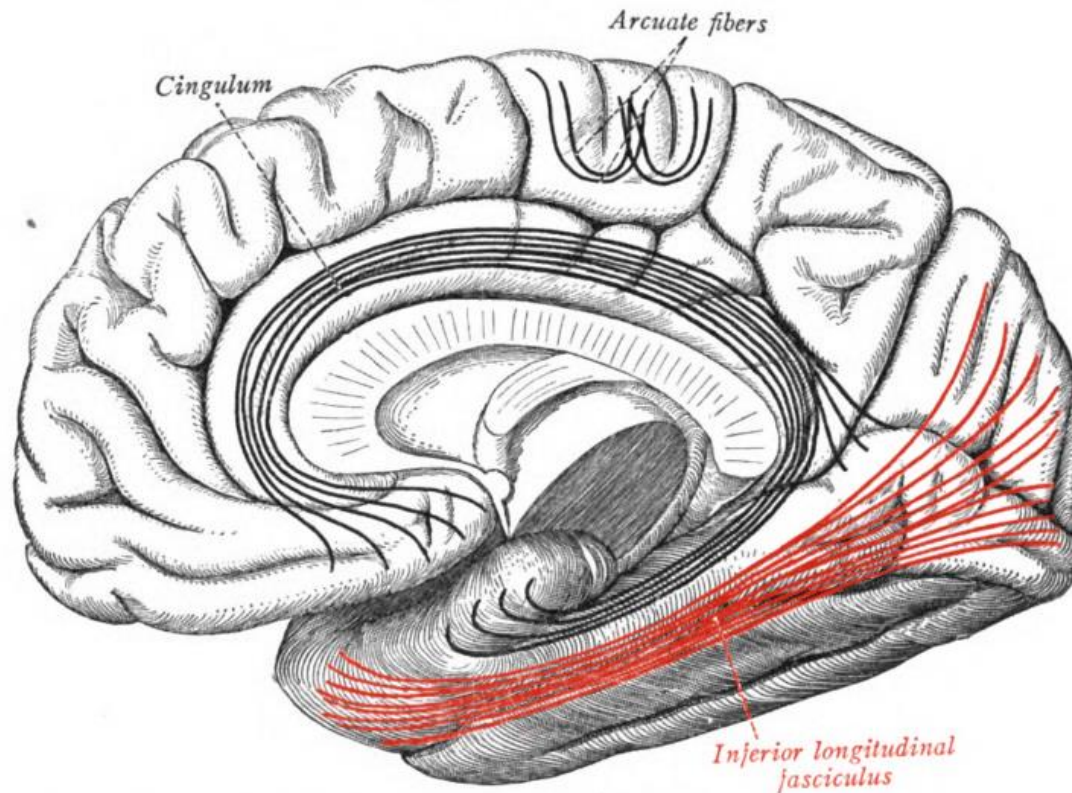
Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex
 - Exposure to **witnessing** multiple episodes of **domestic violence** [especially between ages 7-13]¹
 - Left inferior longitudinal fasciculus: key component of the visual-limbic pathway → subserves **emotional, learning and memory functions** (specific to **vision**)²
 - » Correlated with depression, anxiety, somatization, 'limbic irritability' and neuropsychological measures of processing speed
 - Right lingual gyrus: early processing component of the visual system involved in **visual memory for shapes, faces and letters**
 - » Involved in **nonconscious** processing [especially ages 11-13]³

1. Choi, J et al. Reduced fractional anisotropy in the visual limbic pathway of young adults witnessing domestic violence in childhood.

2. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

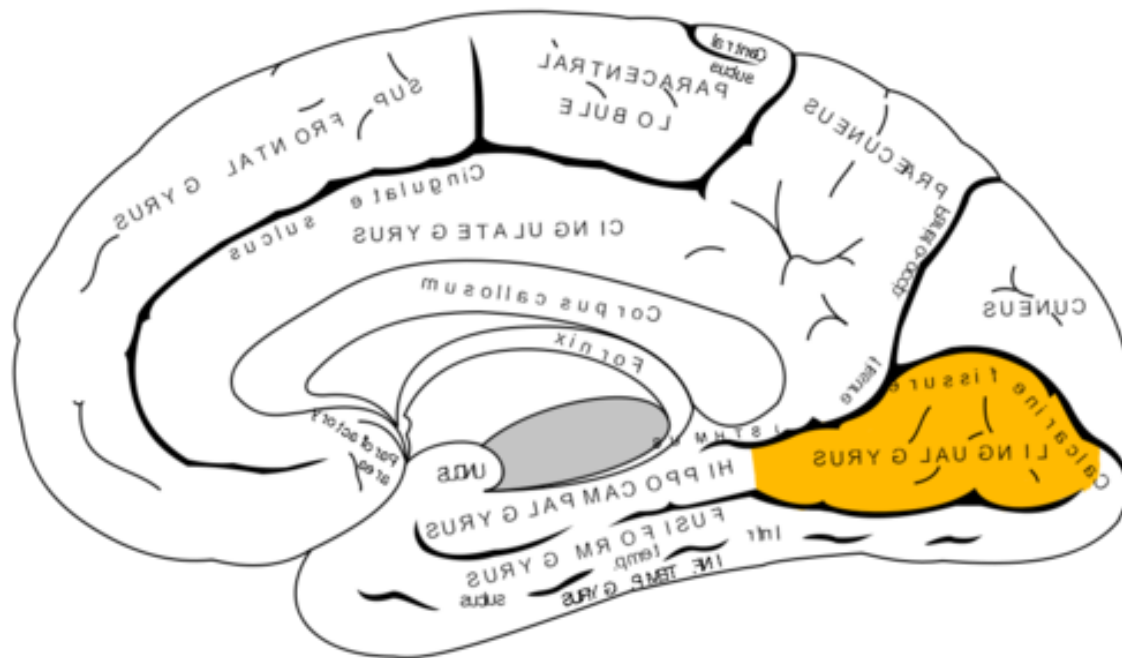
3. Tomoda, A et al. Reduced visual cortex gray matter volume and thickness in young adults who witnessed domestic violence during childhood.



Inferior Longitudinal Fasciculus

Key component of the visual-limbic pathway → subserves **emotional, learning and memory functions** (specific to vision)

Correlated with depression, anxiety, somatization, 'limbic irritability' and neuropsychological measures of processing speed



Lingual Gyrus

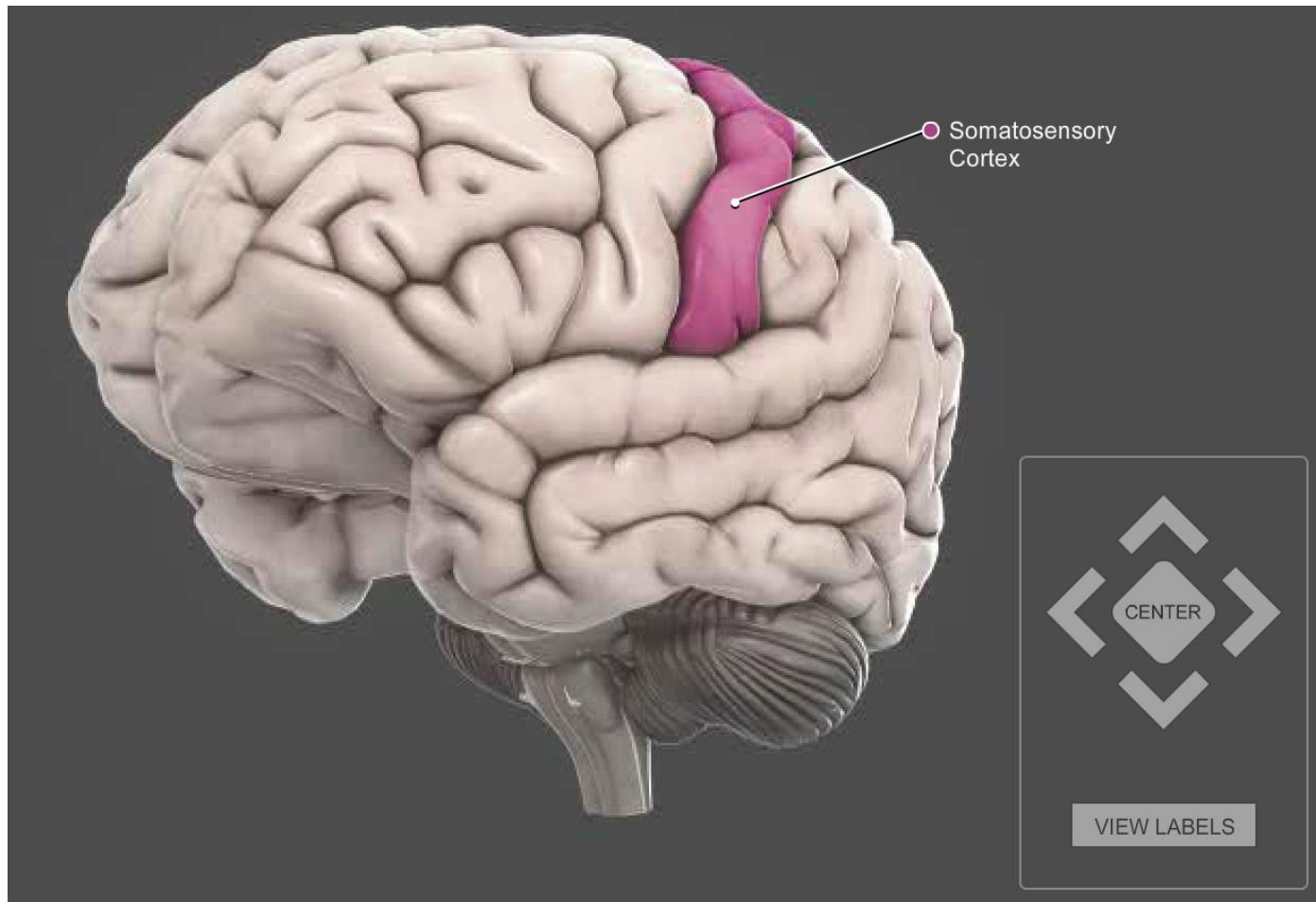
Early processing component of the visual system involved in **visual memory** for shapes, faces and letters

Involved in nonconscious processing [especially ages 11-13]

Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex
 - Exposure to childhood **sexual abuse** was specifically associated with thinning of the portion of **somatosensory cortex** representing the clitoris and surrounding **genital area**¹

Somatosensory Cortex



Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex¹
 - Exposed to **single type of maltreatment**: possible that brain regions and fiber tracts that process, convey adverse sensory input of the abuse may be **specifically modified** by this experience
 - Exposure to **multiple types of maltreatment**: more commonly produce alterations in corticolimbic regions involved in **emotional processing and stress response**

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

Cerebral Cortex

- Region specific
 - Primary and secondary sensory cortex¹
 - Changes in maltreated individuals represent ***modifications or adaptations*** rather than nonspecific damage
 - Neuroplastic cortical adaptations may protectively **shield** a child from the **sensory processing** of the specific abusive experience²
 - » But thinning of the somatosensory cortex may lead to the development of behavioral problems
 - » May lead to **impairments in verbal comprehension, visual recall and emotional responses to witnessed events** later in life

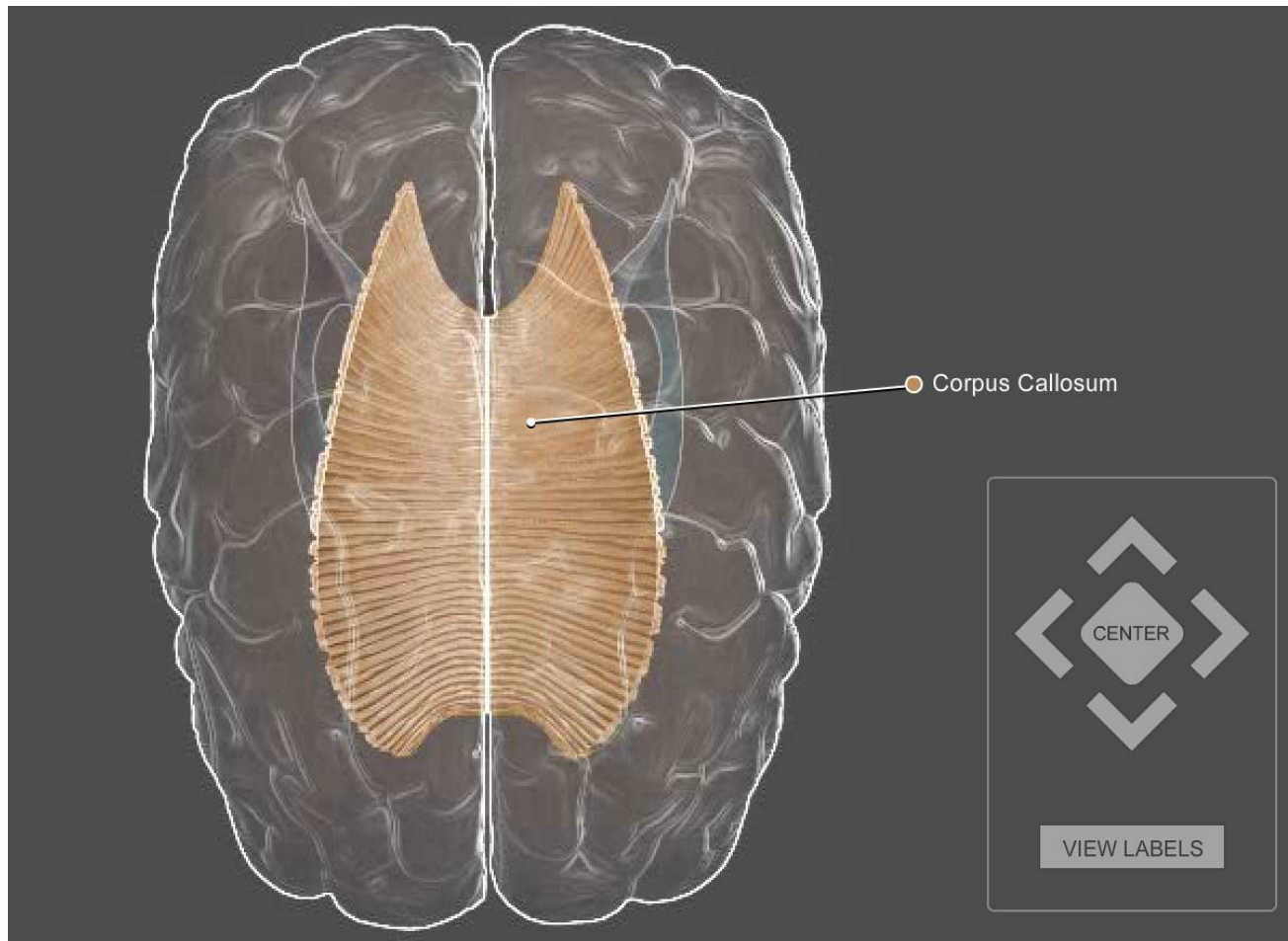
1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Heim, CM et al. Decreased cortical representation of genital somatosensory field after childhood sexual abuse.

Cerebral Cortex

- Region specific
 - Corpus callosum: largest white matter tract¹
 - Critically important role in **inter-hemispheric communication** (particularly between contralateral cortical regions)

Corpus Callosum



Cerebral Cortex

- Region specific
 - Corpus callosum:
 - **Reduced** in maltreated children¹⁻³ and adults^{4,5}
 - Reduced 2x greater in boys than girls^{1-3,6,7}
 - Most susceptible to neglect in males and to sexual abuse in females³
 - Potential reversibility of the effects of early neglect on the corpus callosum⁸

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. De Bellis, MD et al. Developmental traumatology. Part II: Brain development.

3. De Bellis, MD et al. Brain structures in pediatric maltreatment-related posttraumatic stress disorder: A sociodemographically matched study.

4. Teicher, MH et al. Childhood neglect is associated with reduced corpus callosum area.

5. Andersen, SL et al. Preliminary evidence for sensitive periods in the effect of childhood sexual abuse on regional brain development.

6. Teicher, MH et al. Hurtful words: Association of exposure to peer verbal abuse with elevated psychiatric symptom scores and corpus callosum abnormalities.

7. Bellis, MD et al. Brain structures in pediatric maltreatment-related posttraumatic stress disorder: A sociodemographically matched study.

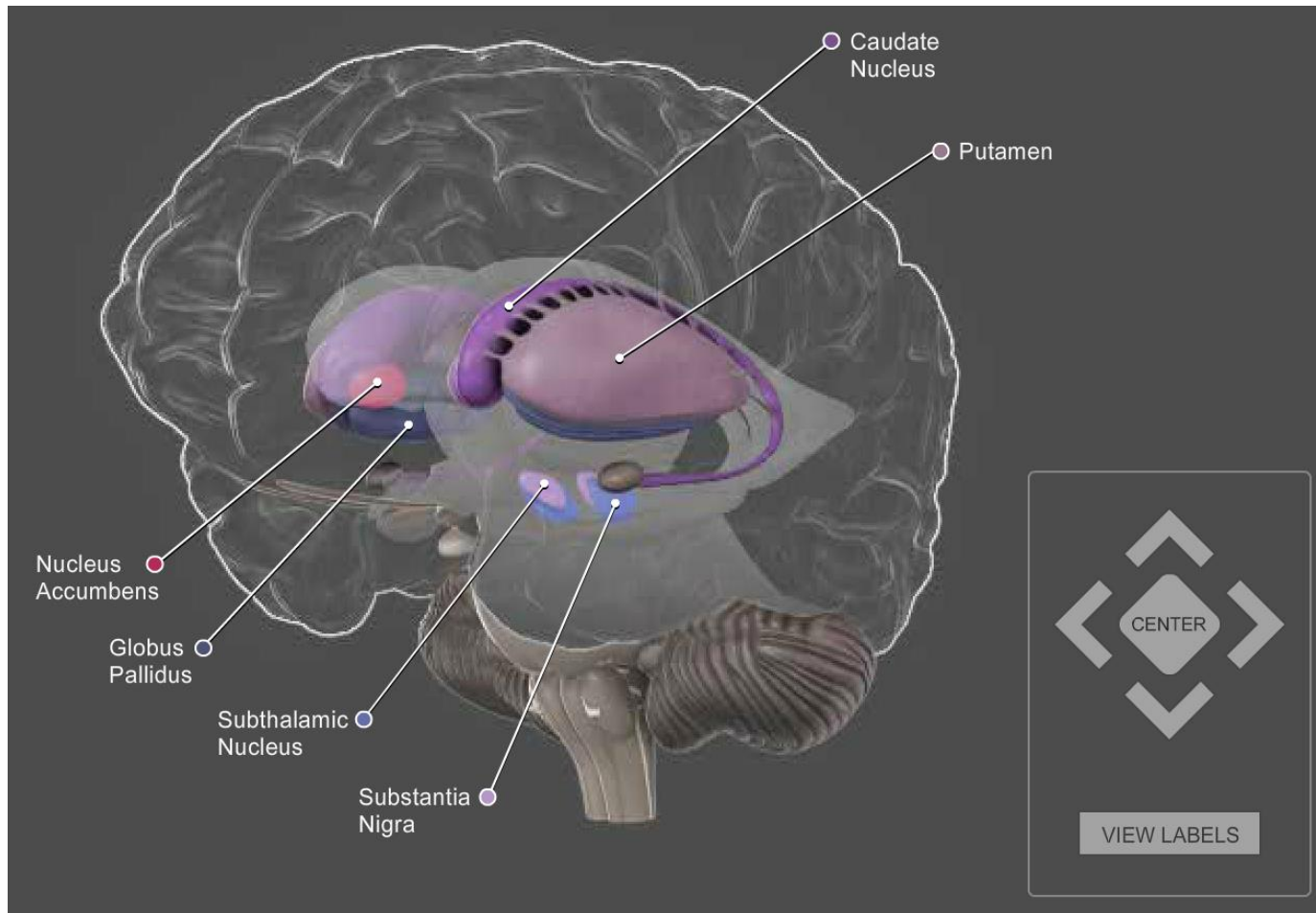
8. Teicher, MH et al. Preliminary evidence for abnormal cortical development in physically and sexually abused children using EEG coherence and MRI.

9. Sheridan, MA et al.. Variation in neural development as a result of exposure to institutionalization early in childhood.

Cerebral Cortex

- Region specific
 - Striatum: **inconsistent** results (sometimes decreased, sometimes no change)
 - May not be sensitive to early life stress
 - Possibly different age and gender effects

Basal Ganglia (Striatum)



Cerebral Cortex

- Region specific
 - Cerebellum: should be **highly susceptible to early** life stressors; research shows usually lower volume in maltreated¹
 - Highest density of glucocorticoid receptors during the neonatal period in rats²
 - Postnatal neurogenesis occurs in cerebellum³
 - Exposure to high levels of glucocorticoids during early development exerted a more persistent effect on cerebellar than hippocampal volume in rats⁴

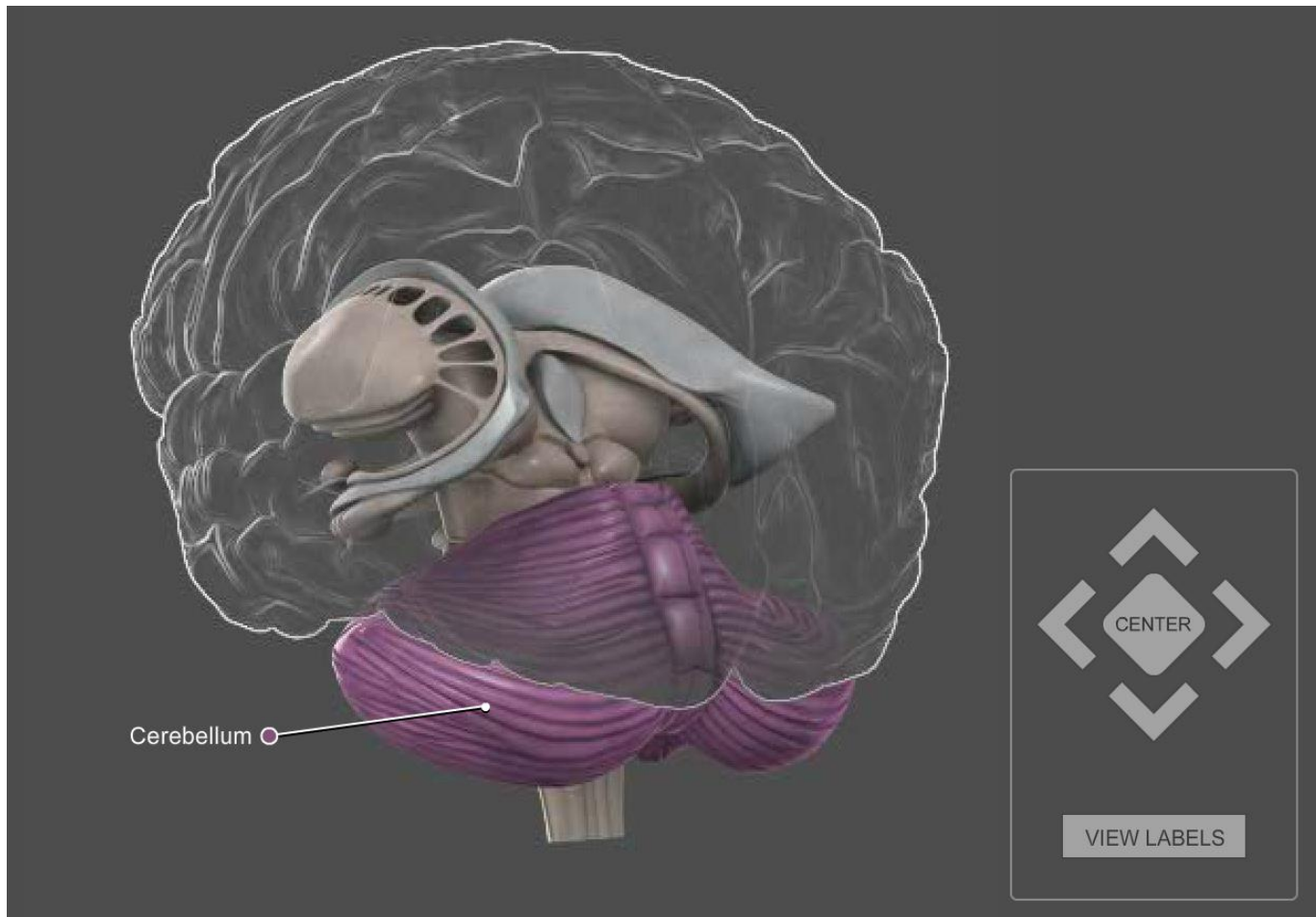
1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Pavlik, A & Buresova, M. The neonatal cerebellum: The highest level of glucocorticoid receptors in the brain.

3. Walton, RM. Postnatal neurogenesis: Of mice, men, and macaques.

4. Ferguson, SA, & Holson, RR. Neonatal dexamethasone on day 7 causes mild hyperactivity and cerebellar stunting.

Cerebellum



Cerebral Cortex

- Region specific
 - Cerebellum: should be highly susceptible to early life stressors; research shows usually lower volume in maltreated¹
 - Vermis in particular seems to be susceptible region
 - » Autism²
 - » Schizophrenia³
 - » Bipolar disorder³
 - » MDD³
 - » ADHD³

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Courchesne, E et al. Hypoplasia of cerebellar vermal lobules VI and VII in autism.

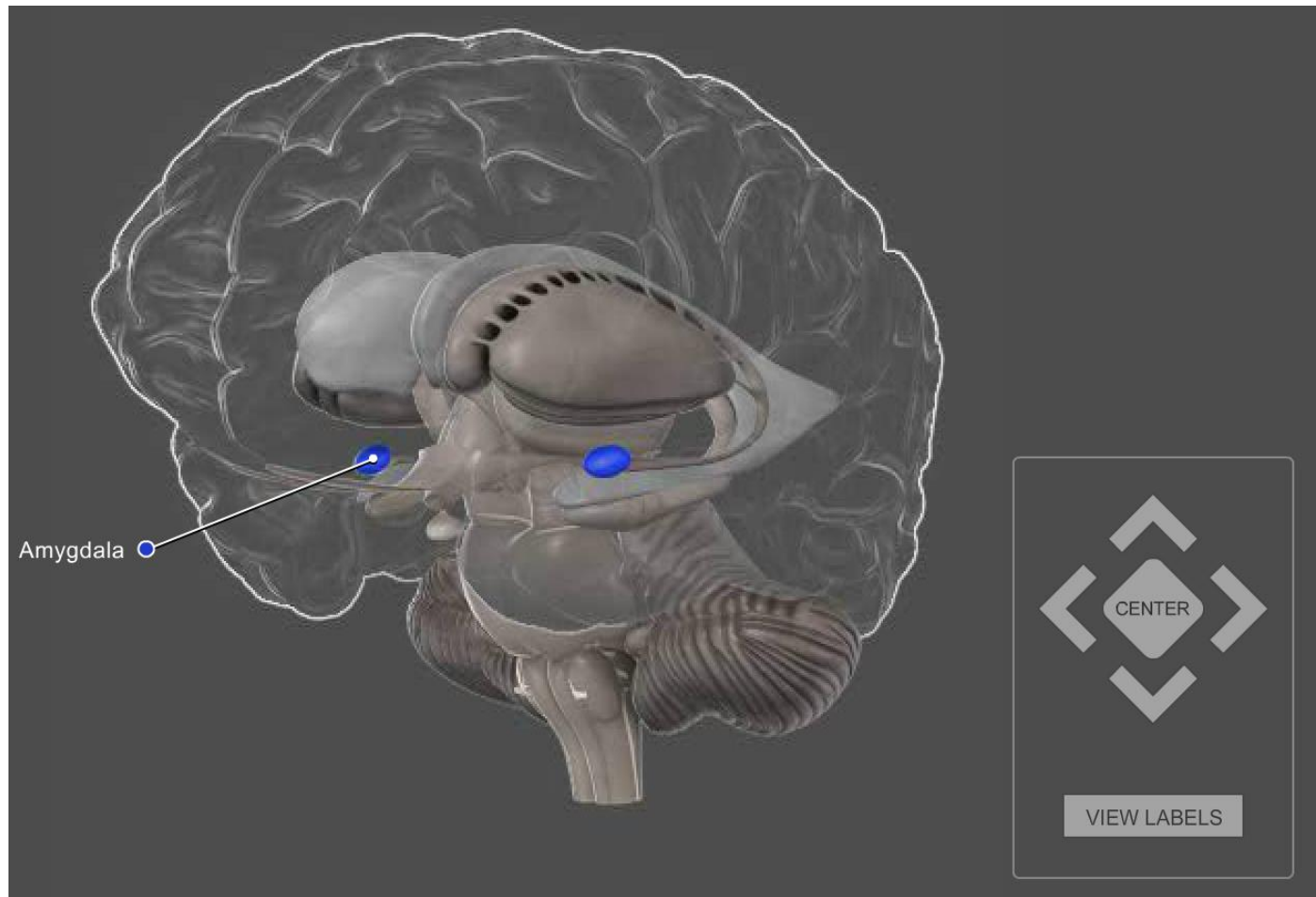
3. Baldacara, L et al. Cerebellum and psychiatric disorders.

Differences in Structure and Function for Those Maltreated

Functional Differences

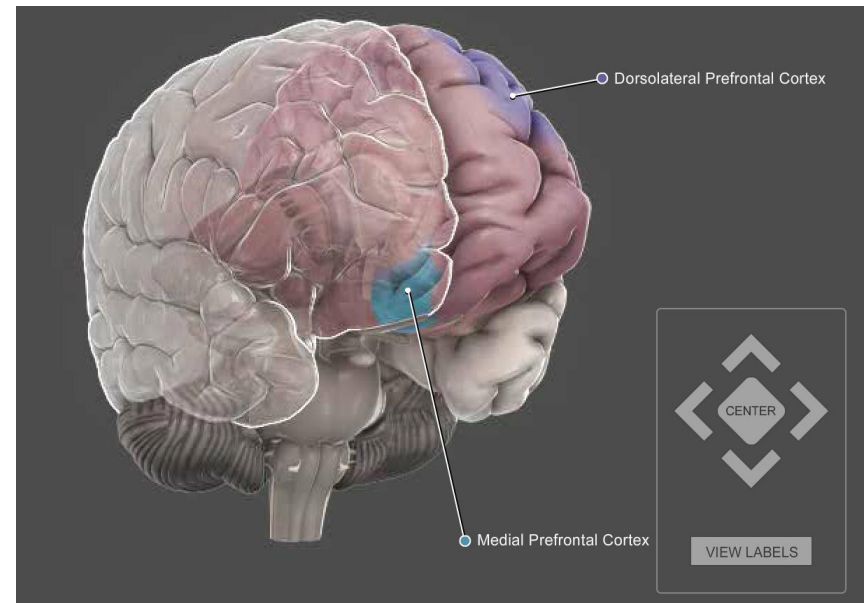
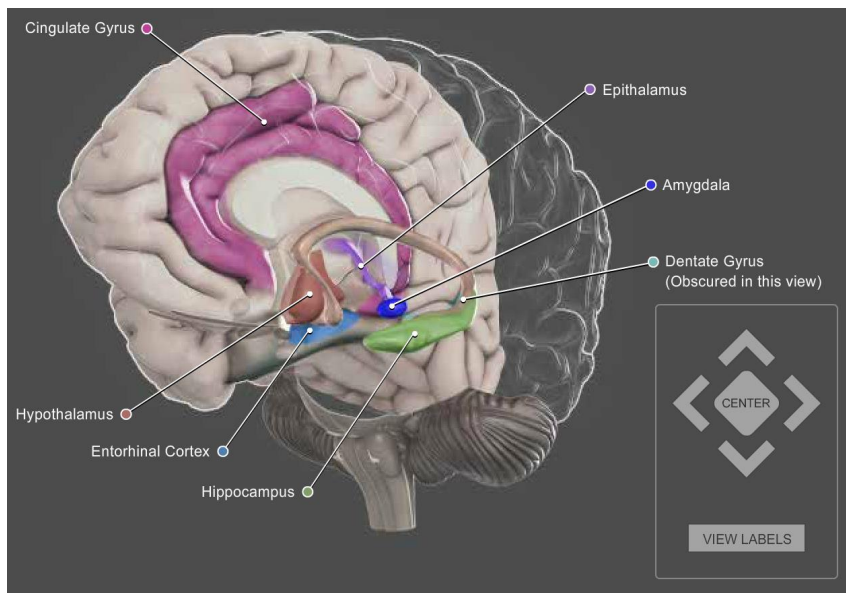
- Differences in response to facial expressions
 - Maltreated = increased amygdala reactivity to emotional faces

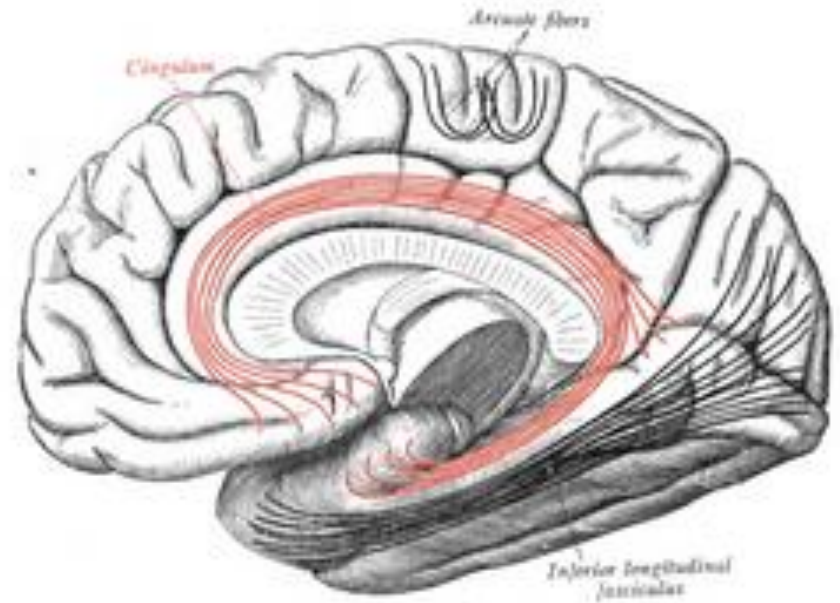
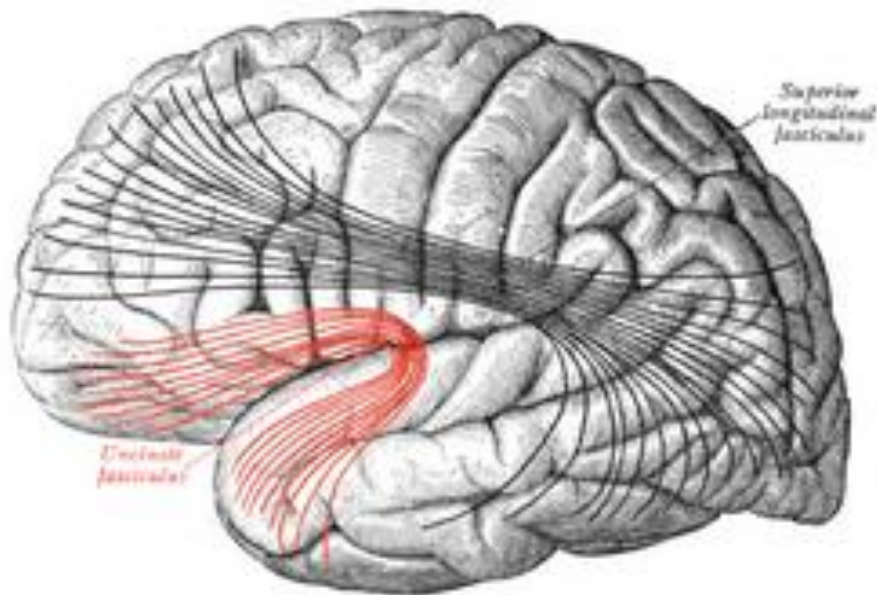
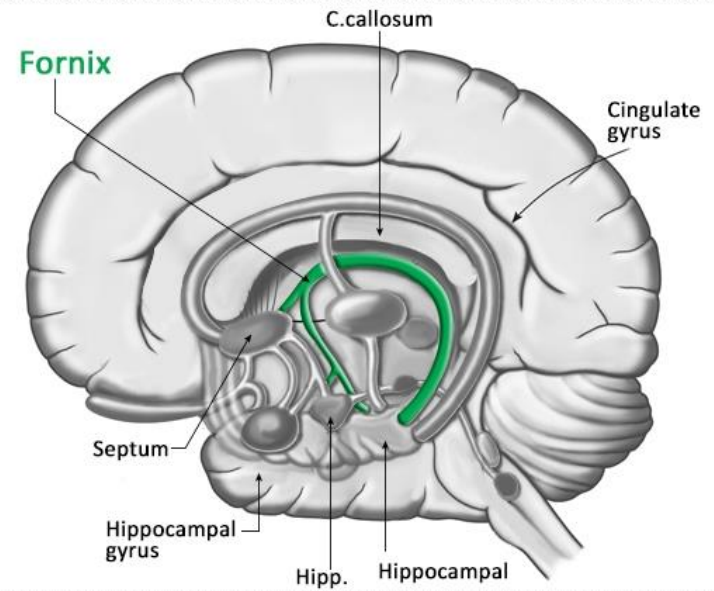
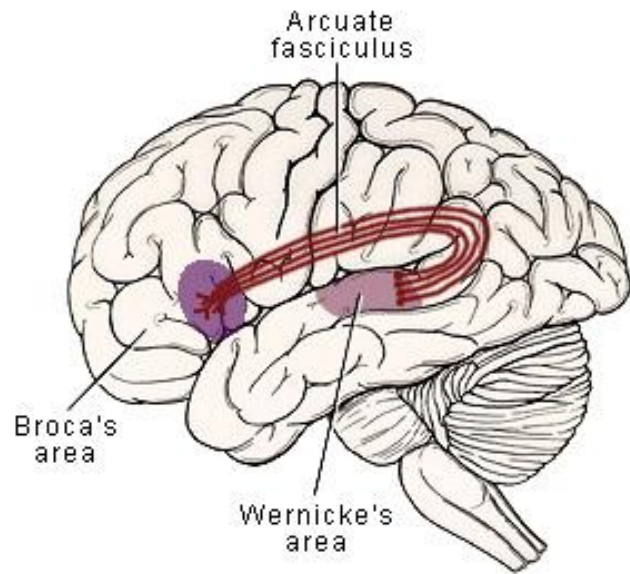
Amygdala



Functional Differences

- Altered response to threatening stimuli
 - Thalamus, visual cortex, anterior cingulate cortex, ventromedial prefrontal cortex, amygdala and hippocampus
 - Decreased integrity of fiber tracts
 - inferior longitudinal fasciculus, superior longitudinal fasciculus/arcuate fasciculus, uncinate fasciculus, cingulum bundle and fornix.
 - Therefore: **most of the regions previously discussed** are also involved in **threat detection**





Functional Differences

- Therefore: changes in brain regions may be **adaptive modifications** to trauma → **enhanced threat detection, more rapid recognition of fearful stimuli**^{1,2}
 - Circuit: **rapidly** responsive **nonconscious** subcortical path and a **slower conscious** cortical path to the amygdala
 - Maltreatment-related alterations in sensory cortex: diminish the influence of the conscious component favoring **rapid but less nuanced** response via the subcortical pathway

1. Teicher, MH & Samson, JA. Annual research view: Enduring neurobiological effects of childhood abuse and neglect.

2. Masten, CL et al. Recognition of facial emotions among maltreated children with high rates of post-traumatic stress disorder.

Functional Differences

- Differences in sensitive exposure periods of different regions
 - Range: hippocampus (3-5 years old) to prefrontal cortex (14-16 years old)
 - Therefore: Because multiple different regions involved in these circuits → **can have multiple different reasons for same overall outcome**

Functional Differences

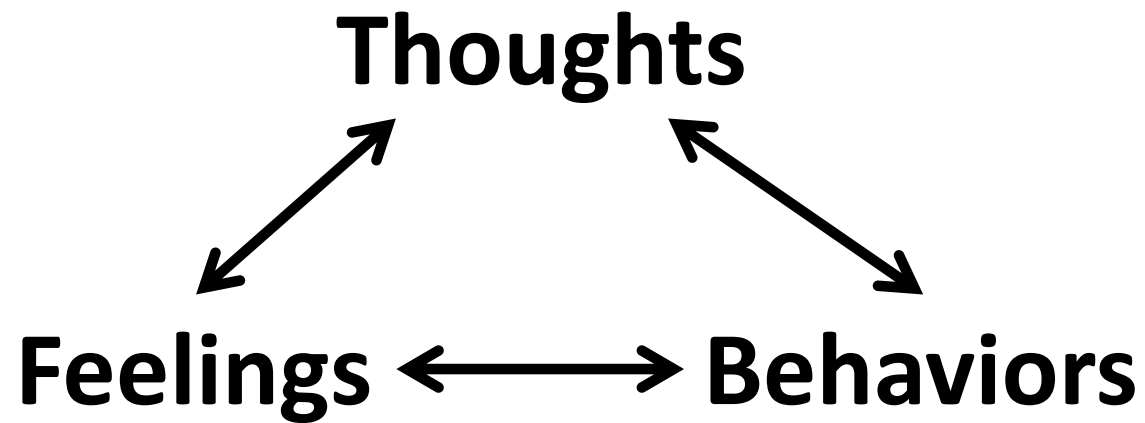
- Reward anticipation
 - Areas involved: Many of the same ones we have just covered
 - Again, many of these regions affected in those with maltreatment
 - Overall: **reduced response to anticipatory reward**
 - May be adaptive: if decrease belief that an action will be rewarded, more likely to **avoid** (like avoiding things that lead to maltreatment)

Network Architecture

- Maltreated individuals have **diminished** capacity to:
 - Regulate impulses and emotions
 - Accurately attribute thoughts and intentions to others
 - Be mindful of oneself in a social context
- Maltreated individuals have **increased**
 - Experience of internal emotions and cravings
 - Greater tendency to think about oneself and to engage in self-centered mental imagery

Network Architecture

- Therefore: neurobiological explanations for why psychotherapy for these individuals focuses on:
 - Enhance emotional regulation
 - Correct misconceptions about self and others
 - Diminish focus on internal feelings and to reduce harmful self-centered thinking



Summary

Summary

- There are many effects that come with maltreatment, including both direct and indirect effects of domestic violence
- While there are clear associations between chronic trauma and the development of PTSD, maltreatment has far ranging consequences both in terms of cognitive development and the increased risk of a large variety of mental health disorders
- The focus here has been primarily “organic” changes that come with being a part of domestic violence → however, we are increasingly learning about the direct and indirect psychological consequences of these events

References

- Andersen, SL & Teicher, MH. Delayed effects of early stress on hippocampal development. *Neuropsychopharmacology*. 2004; 29: 1988–1993
- Andersen, SL & Teicher, MH. Stress, sensitive periods and maturational events in adolescent depression. *Trends in Neurosciences*. 2008; 31: 183–191
- Anderson, CM et al. Abnormal T2 relaxation time in the cerebellar vermis of adults sexually abused in childhood: Potential role of the vermis in stress-enhanced risk for drug abuse. *Psychoneuroendocrinology*. 2002;27: 231–244
- Andersen, SL et al. Preliminary evidence for sensitive periods in the effect of childhood sexual abuse on regional brain development. *Journal of Neuropsychiatry and Clinical Neurosciences*. 2008; 20: 292–301
- Arnone, D et al. Meta-analysis of magnetic resonance imaging studies of the corpus callosum in bipolar disorder. *Acta Psychiatrica Scand*. 2008;118: 357–362.

References

- Baldacara, L et al. Cerebellum and psychiatric disorders. *Revista Brasileira de Psiquiatria*. 2008;30: 281–289
- Ball, JS & Links, PS. Borderline personality disorder and childhood trauma: Evidence for a causal relationship. *Current Psychiatry Reports*. 2009;11: 63–68
- Baloch, HA, Brambilla, P, & Soares, JC. Corpus callosum abnormalities in pediatric bipolar disorder. *Expert Review of Neurotherapeutics*, 2009;9: 949–955.
- Benedetti, F et al. Disruption of white matter integrity in bipolar depression as a possible structural marker of illness. *Biological Psychiatry*. 2011; 69: 309– 317
- Benedetti, F et al. Adverse childhood experiences influence white matter microstructure in patients with bipolar disorder. *Psychological Medicine*. 2014; 44: 3069–3082.
- Bucker, J et al. Childhood maltreatment and corpus callosum volume in recently diagnosed patients with bipolar I disorder: Data from the Systematic Treatment Optimization Program for Early Mania (STOP-EM). *Journal of Psychiatric Research*. 2014; 48: 65–72
- De Bellis, MD et al. Developmental traumatology. Part II: Brain development. *Biological Psychiatry*. 1999; 45: 1271–1284.
- De Bellis, MD et al. A pilot longitudinal study of hippocampal volumes in pediatric maltreatment-related posttraumatic stress disorder. *Biological Psychiatry*. 2001; 50: 305–309.
- De Bellis, MD et al. Brain structures in pediatric maltreatment-related posttraumatic stress disorder: A sociodemographically matched study. *Biological Psychiatry*. 2002;52: 1066–1078.
- de Bellis, MD et al. Neuropsychological findings in childhood neglect and their relationships to pediatric PTSD. *Journal of the International Neuropsychological Society*. 2009;15: 868–878
- Bendall, S et al. Childhood trauma and psychotic disorders: A systematic, critical review of the evidence. *Schizophrenia Bulletin*. 2008; 34: 568–579
- Bremner, JD et al. Magnetic resonance imaging-based measurement of hippocampal volume in posttraumatic stress disorder related to childhood physical and sexual abuse—a preliminary report. *Biological Psychiatry*. 1997; 41: 23–32
- Brown, DW et al. Adverse childhood experiences and the risk of premature mortality. *American Journal of Preventive Medicine*. 2009; 37: 389–396.
- Brown, DW et al. Adverse childhood experiences are associated with the risk of lung cancer: A prospective cohort study. *BMC Public Health*. 2010;10: 20

References

- Carrion, VG et al.. Attenuation of frontal asymmetry in pediatric posttraumatic stress disorder. *Biological Psychiatry*. 2001;50: 943–951.
- Choi, J et al. Preliminary evidence for white matter tract abnormalities in young adults exposed to parental verbal abuse. *Biological Psychiatry*. 2009;65: 227–234.
- Choi, J et al. Reduced fractional anisotropy in the visual limbic pathway of young adults witnessing domestic violence in childhood. *NeuroImage*. 2012; 59: 1071–1079
- Clark, K et al. White matter integrity, language, and childhood onset schizophrenia. *Schizophrenia Research*. 2012;138: 150–156
- Courchesne, E et al. Hypoplasia of cerebellar vermal lobules VI and VII in autism. *New England Journal of Medicine*. 1988;318: 1349–1354
- Craig, MC et a. Altered connections on the road to psychopathy. *Molecular Psychiatry*. 2009; 14: 946– 953, 907.

References

- Danese, A et al. Childhood maltreatment predicts adult inflammation in a life-course study. *Proceedings of the National Academy of Sciences of the United States of America*. 2010; 104: 1319–1324.
- Danese, A et al. Adverse childhood experiences and adult risk factors for age-related disease: Depression, inflammation, and clustering of metabolic risk markers. *Archives of Pediatrics and Adolescent Medicine*. 2009;163: 1135–1143
- Dannlowski, U et al. Limbic scars: Long-term consequences of childhood maltreatment revealed by functional and structural magnetic resonance imaging. *Biological Psychiatry*. 2012; 71: 286–293.
- Derntl, B et al. General and specific responsiveness of the amygdala during explicit emotion recognition in females and males. *BMC Neuroscience*. 2009; 10: 91
- Dong, M et al. Insights into causal pathways for ischemic heart disease: Adverse childhood experiences study. *Circulation*. 2004;110: 1761–1766
- Edmiston, EE et al. Corticostriatal-limbic gray matter morphology in adolescents with self-reported exposure to childhood maltreatment. *Archives of Pediatrics and Adolescent Medicine*. 2011; 165: 1069–1077
- Egeland, B. Taking stock: Childhood emotional maltreatment and developmental psychopathology. *Child Abuse & Neglect*. 2009;33:22-26.

References

- Frazier, JA et al. White matter abnormalities in children with and at risk for bipolar disorder. *Bipolar Disorders*. 2007;9: 799–809
- Ferguson, SA, & Holson, RR. Neonatal dexamethasone on day 7 causes mild hyperactivity and cerebellar stunting. *Neurotoxicology and Teratology*. 1999; 21: 71– 76
- Genes to Cognition. Accessed at <http://www.g2conline.org/> on March 26, 2016.
- Geuze, E, Vermetten, E & Bremner, JD. MR-based in vivo hippocampal volumetrics: 2. Findings in neuropsychiatric disorders. *Molecular Psychiatry*. 2005; 10: 160– 184.
- Gould, F et al. The effects of child abuse and neglect on cognitive functioning in adulthood. *Journal of Psychiatric Research*. 2012; 4: 500–506.
- Green, JG et al. Childhood adversities and adult psychiatric disorders in the national comorbidity survey replication I: Associations with first onset of DSM-IV disorders. *Archives of General Psychiatry*. 2010; 67: 113–123.
- Gurrera, RJ et al. The uncinate fasciculus and extraversion in schizotypal personality disorder: A diffusion tensor imaging study. *Schizophrenia Research*. 2007;90: 360–362.

References

- Hamilton, LS et al. Reduced white matter integrity in attention-deficit hyperactivity disorder. *NeuroReport*. 2008;19: 1705–1708
- Hayes, JP. (2009). Parenting Assessment in Abuse, Neglect and Permanent Wardship Cases. In Principles and practice of child and adolescent forensic mental health. American Psychiatric Publishing.
- Heim, CM et al. Decreased cortical representation of genital somatosensory field after childhood sexual abuse. *American Journal of Psychiatry*. 2013;170: 616–623.
- Ito, Y et al. Increased prevalence of electrophysiological abnormalities in children with psychological, physical, and sexual abuse. *Journal of Neuropsychiatry and Clinical Neurosciences*. 1993; 5: 401–408
- Kawashima, T et al. Uncinate fasciculus abnormalities in recent onset schizophrenia and affective psychosis: A diffusion tensor imaging study. *Schizophrenia Research*. 2009; 110, 119–126.
- Kitzmann KM et al. Child witnesses to domestic violence: a meta-analytic review. *J Consult Clin Psychol*. 2003; 71(2):339–52
- Keyes KM et al. Childhood maltreatment and the structure of common psychiatric disorders. *Br J Psychiatry*. 2012;200(2):107–15.
- Klika, JB & Herrenkohl, TI. A review of developmental research on resilience in maltreated children. *Trauma Violence & Abuse*. 2013, 14(3): 222-234
- van der Kolk, B & Greenberg, MS. (1987). The psychobiology of the trauma response: Hyperarousal, constriction, and addiction to traumatic reexposure. In B. van der Kolk (Ed.), *Psychological trauma* (pp. 63–87). Washington, DC: American Psychiatric Press.
- Koob, GF & Volkow, ND. Neurocircuitry of addiction. *Neuropsychopharmacology*. 2010; 35: 217–238.
- de Kwaasteniet, B et al. Relation between structural and functional connectivity in major depressive disorder. *Biological Psychiatry*. 2013;74: 40–47.

References

- Lai, CH & Wu, YT. Alterations in white matter micro- integrity of the superior longitudinal fasciculus and anterior thalamic radiation of young adult patients with depression. *Psychological Medicine*. 2014; 44: 2825–2832
- LeDoux, JE. Emotional memory systems in the brain. *Behavioural Brain Research*. 1993; 58:, 69–79
- Lenroot, RK et al. Differences in genetic and environmental influences on the human cerebral cortex associated with development during childhood and adolescence. *Human Brain Mapping*, 2009;30: 163–174.
- Levendosky AA, Bogat GA, & Martinez-Torteya C. PTSD symptoms in young children exposed to intimate partner violence. *Violence Against Women*. 2013; 19(2):187–201
- Luders, E et al. Positive correlations between corpus callosum thickness and intelligence. *NeuroImage*. 2007; 37: 1457–1464.
- Luders, E et al. Decreased callosal thickness in attention-deficit/hyperactivity disorder. *Biological Psychiatry*. 2009;65: 84–88.

References

- Masten, CL et al. Recognition of facial emotions among maltreated children with high rates of post-traumatic stress disorder. *Child Abuse and Neglect*. 2008; 32: 139–153
- McCain NM, Mustard F, & Shanker S. Early Years Study 2. Putting science into action. Toronto, ON: Council for Early Child Development; 2007. Accessed 2014 Sept 24, available from: http://earlylearning.ubc.ca/media/publications/early_years_study_2.pdf
- McEwen, BS. . Stress, sex, and neural adaptation to a changing environment: Mechanisms of neuronal remodeling. *Annals of the New York Academy of Sciences*. 2010; 1204(Suppl 1): E38–E59
- McIntosh, AM et al. White matter tractography in bipolar disorder and schizophrenia. *Biological Psychiatry*. 2008; 64, 1088–1092.
- Melicher, T et al. White matter changes in first episode psychosis and their relation to the size of sample studied: A DTI study. *Schizophrenia Research*. 2015;162: 22–28
- Mitra, R et al. Stress duration modulates the spatiotemporal patterns of spine formation in the basolateral amygdala. *Proceedings of the National Academy of Sciences of the United States of America*. 2005; 102: 9371–9376.
- Morimoto, M et al. Distribution of glucocorticoid receptor immunoreactivity and mRNA in the rat brain: An immunohistochemical and in situ hybridization study. *Neuroscience Research*. 1996; 26: 235– 269.
- Moser, EI, Kropff, E & Moser, MB. Place cells, grid cells, and the brain's spatial representation system. *Annual Review of Neuroscience*. 2008; 31: 69–89

References

- Nadel, L, Campbell, J, & Ryan, L. Autobiographical memory retrieval and hippocampal activation as a function of repetition and the passage of time. *Neural Plasticity*. 2007;2007: 90472.
- Nanni, V, Uher, R & Danese, A. Childhood maltreatment predicts unfavorable course of illness and treatment outcome in depression: A meta-analysis. *American Journal of Psychiatry*. 2012;169: 141–151.
- Norman, RE et al. The long-term health consequences of child physical abuse, emotional abuse, and neglect: A systematic review and meta-analysis. *PLoS Medicine*, 2012;9: e1001349.
- Odgers, CL & Jaffee, SR. Routine versus catastrophic influences on the developing child. *Annual Review Public Health*. 2013;34: 29-48.
- Odhayani, AA, Watson, WJ & Watson, L. Behavioral consequences of child abuse. *Canadian Family Physician*. 2013;59:831-836.
- Pagliaccio, D et al. Stress-system genes and life stress predict cortisol levels and amygdala and hippocampal volumes in children. *Neuropsychopharmacology*. 2014; 39: 1245–1253.
- Pavlik, A & Buresova, M. The neonatal cerebellum: The highest level of glucocorticoid receptors in the brain. *Brain Research*. 1984;314: 13–20
- Phan, KL et al. Preliminary evidence of white matter abnormality in the uncinate fasciculus in generalized social anxiety disorder. *Biological Psychiatry*. 2009;66: 691–694
- Price, LH et al. Telomeres and early-life stress: An overview. *Biological Psychiatry*, 2013;73: 15–23
- Rilling, JK et al. The evolution of the arcuate fasciculus revealed with comparative DTI. *Nature Neuroscience*. 2008; 11: 426–428.
- Rinne-Albers, MA et al. Neuroimaging in children, adolescents and young adults with psychological trauma. *Eur Child Adolesc Psychiatry*. 2013;22:745-755.

References

- Sarrieau, A et al. Autoradiographic localization of glucocorticosteroid and progesterone binding sites in the human post-mortem brain. *Journal of Steroid Biochemistry*. 1986; 25: 717–721.
- Sapolsky, RM. Stress, glucocorticoids, and damage to the nervous system: The current state of confusion. *Stress*. 1996;1: 1–19.
- Sapolsky, RM, Krey, LC, & McEwen, BS. Prolonged glucocorticoid exposure reduces hippocampal neuron number: Implications for aging. *Journal of Neuroscience*. 1985; 5: 1222–1227
- Sheffield, JM et al. Reduced gray matter volume in psychotic disorder patients with a history of childhood sexual abuse. *Schizophrenia Research*. 2013;143: 185–191.
- Sheridan, MA et al.. Variation in neural development as a result of exposure to institutionalization early in childhood. *Proceedings of the National Academy of Sciences of the United States of America*,. 2012;109: 12927–12932.
- Sinclair, D et al. Dynamic molecular and anatomical changes in the glucocorticoid receptor in human cortical development. *Molecular Psychiatry*, 2011;16: 504–515.
- Smyke, AT et al. A new model of foster care for young children: The Bucharest early intervention project. *Child and Adolescent Psychiatric Clinics of North America*. 2009; 18: 721–734.
- Spitzer, C et al. Gender-specific association between childhood trauma and rheumatoid arthritis: A case-control study. *Journal of Psychosomatic Research*. 2013;74: 296–300
- Stein, MB et al. Hippocampal volume in women victimized by childhood sexual abuse. *Psychological Medicine*. 1997; 27: 951–959

References

- Teicher, MH. Wounds that time won't heal: The neurobiology of child abuse. *Cerebrum*. 2000; 4: 50–67
- Teicher, MH. Scars that won't heal: The neurobiology of child abuse. *Scientific American*. 2002; 286: 68–75
- Teicher, MH & Samson, JA. Childhood maltreatment and psychopathology: A case for ecophenotypic variants as clinically and neurobiologically distinct subtypes. *American Journal of Psychiatry*. 2013;170: 1114–1133.
- Teicher, MH, Anderson, CM, & Polcari, A. Childhood maltreatment is associated with reduced volume in the hippocampal subfields CA3, dentate gyrus, and subiculum. *Proceedings of the National Academy of Sciences of the United States of America*. 2012; 109: E563–E572.
- Teicher, MH et al. Early childhood abuse and limbic system ratings in adult psychiatric outpatients. *Journal of Neuropsychiatry and Clinical Neurosciences*, 1993;5: 301–306
- Teicher, MH et al. Preliminary evidence for abnormal cortical development in physically and sexually abused children using EEG coherence and MRI. *Annals of the New York Academy of Sciences*. 1997;821: 160–175.
- Teicher, MH et al. Childhood neglect is associated with reduced corpus callosum area. *Biological Psychiatry*. 2004; 56: 80–85.
- Teicher, MH et al. Hurtful words: Association of exposure to peer verbal abuse with elevated psychiatric symptom scores and corpus callosum abnormalities. *American Journal of Psychiatry*. 2010; 167: 1464–1471.
- Tomoda, A et al.. Exposure to parental verbal abuse is associated with increased gray matter volume in superior temporal gyrus. *NeuroImage*. 2011;54(Suppl 1): S280–S286.
- Tomoda, A et al. Reduced visual cortex gray matter volume and thickness in young adults who witnessed domestic violence during childhood. *PLoS ONE*. 2012; 7: e52528.
- Torchalla, I et al. The association between childhood maltreatment subtypes and current suicide risk among homeless men and women. *Child Maltreat*. 2012;17(2): 132–143.
- Treadway, MT et al. Early adverse events, HPA activity and rostral anterior cingulate volume in MDD. *PLoS ONE*. 2009; 4: e4887.
- Tsavoussis, A et al. Child-witnessed domestic violence and its adverse effects on brain development: A call for societal self-examination and awareness. *Frontiers in Public Health*. 2014;2:1–5.

References

- Uematsu, A et al. Developmental trajectories of amygdala and hippocampus from infancy to early adulthood in healthy individuals. *PLoS ONE*. 2012; 7: e46970.
- Vyas, A, Pillai, AG & Chattarji, S. Recovery after chronic stress fails to reverse amygdaloid neuronal hypertrophy and enhanced anxiety-like behavior. *Neuroscience*. 2004; 128: 667– 673
- Vyas, A, Jadhav, S & Chattarji, S. Prolonged behavioral stress enhances synaptic connectivity in the basolateral amygdala. *Neuroscience*. 2006;143: 387–393.
- Walton, RM. Postnatal neurogenesis: Of mice, men, and macaques. *Veterinary Pathology*. 2012; 49: 155–165
- Whittle, S et al. Childhood maltreatment and psychopathology affect brain development during adolescence. *Journal of the American Academy of Child and Adolescent Psychiatry*. 2013;52: 940–952, e941
- Wolfers, T et al. Lower white matter microstructure in the superior longitudinal fasciculus is associated with increased response time variability in adults with attention-deficit/hyperactivity disorder. *Journal of Psychiatry and Neuroscience*. 2015; 40: 140154.

Questions?

psmartin@buffalo.edu