



Katrina Kully-Martens, MEd
PhD Candidate

E6 - Prenatal Alcohol Exposure and Social Behavior Function

Challenges Associated with Navigating
the Social World with FASD

Parker Holman, MS, MEd
PhD Candidate



Acknowledgments



Dr. Joanne Weinberg

Professor and Distinguished University Scholar, Emerita

Dr. Tamara Bodnar
Immune Function,
Inflammation



Dr. Charlis Rainecki
Early-Life Adversity



Dr. Ni Lan
Stress Signaling,
Placenta, Fetal
Brain



National Institute
on Alcohol Abuse
and Alcoholism



Parker Holman
Social Behavior



Vivian Lam
Stress & Mental
Health Outcomes



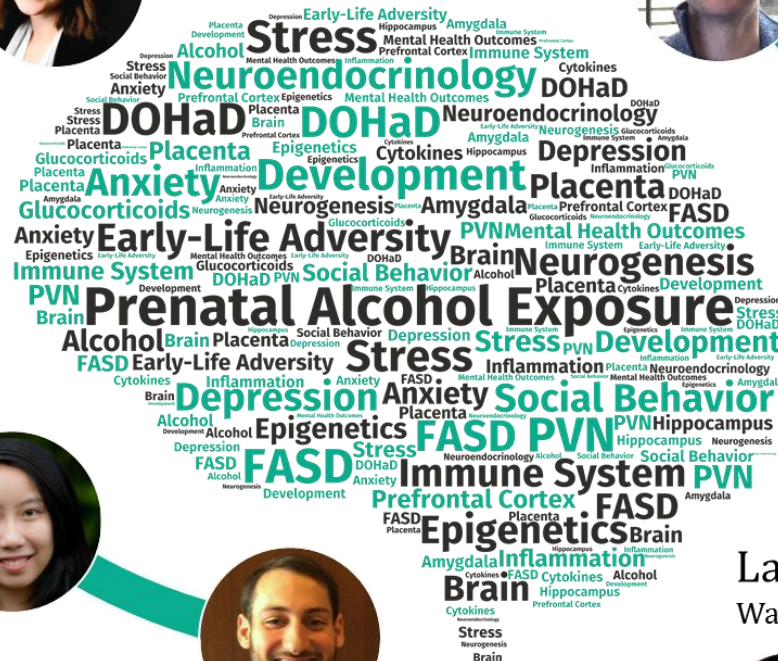
Alexandre Lussier
Gene Expression,
Epigenetic Mechanisms



Lab Managers:

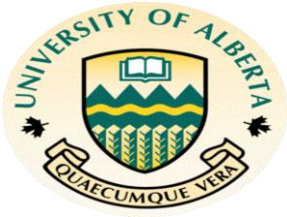
Wayne Yu

Linda Ellis



• Funding:

- NIH/NIAAA F31 AA023151
- KBHN (NeuroDevNet)



Acknowledgements



FASD RESEARCH LAB

University of Alberta & Glenrose Rehabilitation Hospital
Edmonton, AB, Canada

- Dr. Carmen Rasmussen
- Dr. Jacqueline Pei
- Dr. Gail Andrew
- Research Staff: Lauren Baugh, Michael-Anne LaFrance, Kathryn Kryska
- Graduate Students: Allison McNeil, Elise Regehr



SSHRC  CRSH



Social Sciences and
Humanities Research
Council of Canada

Conseil de recherches
en sciences humaines
du Canada

Canada



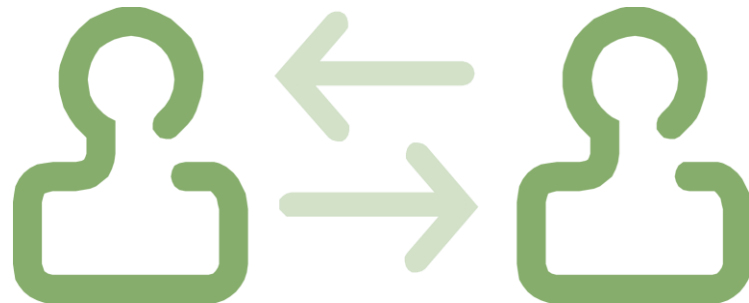
Bourses d'études
supérieures du Canada
Vanier
Canada Graduate
Scholarships

Outline

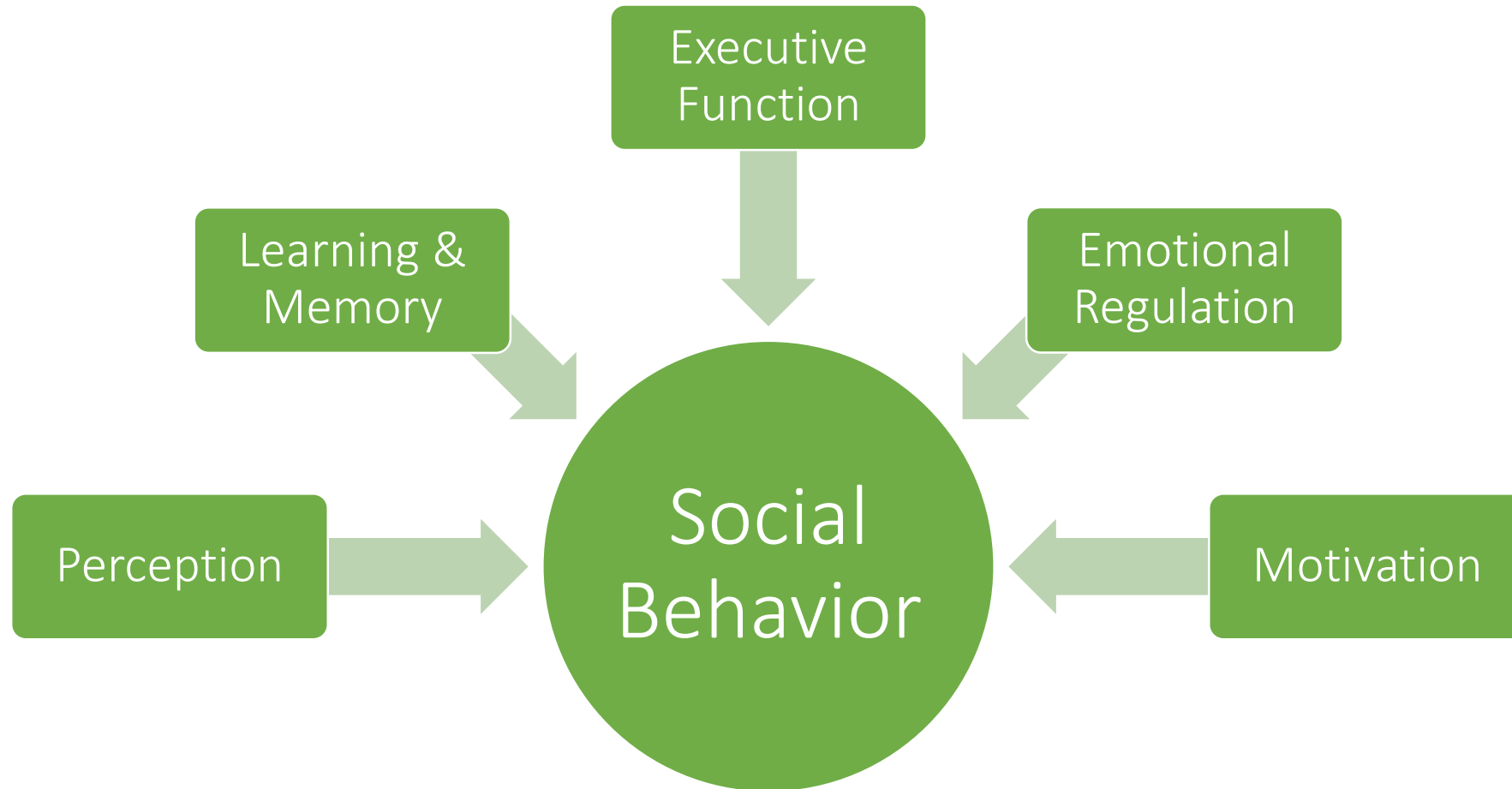
- Provide an overview of how prenatal alcohol exposure (PAE) can alter social cognition and behavior development, with a particular focus on social problem solving skills and social perspective taking
- Discuss findings from an animal model of PAE, including potential neural mechanisms of social behavior dysfunction
- Examine the relationship between resilience, mental health, and social competence in adolescents with FASD

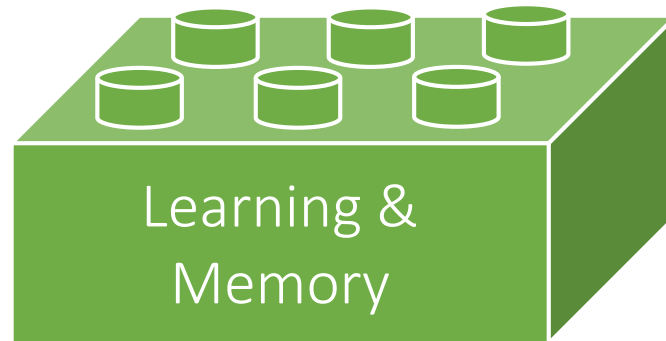
How do we define Social Behavior?

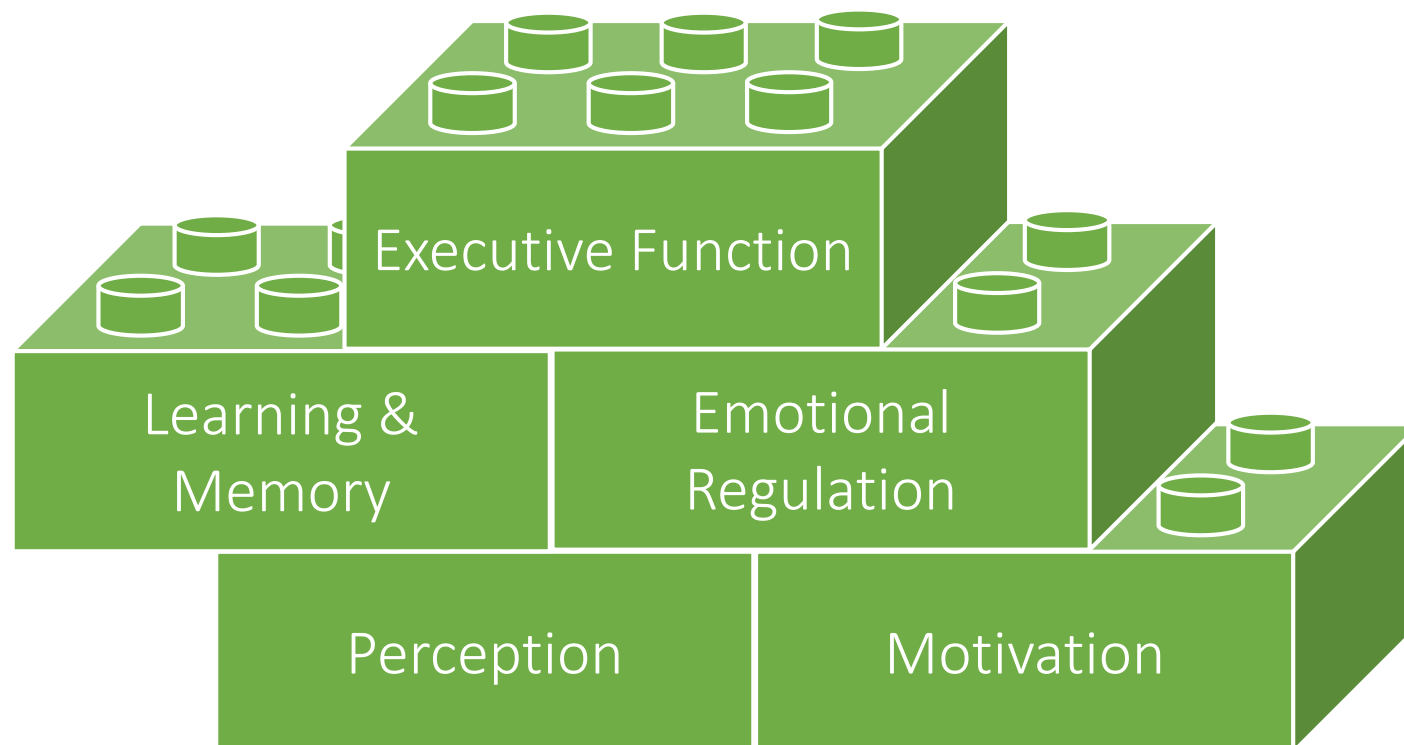
- “Social Skills”
 - Initiate and manage social interactions
 - Establish and maintain friendship(s)
 - Express and interpret emotions
 - Use interpersonal/communication skills
 - Inhibit inappropriate/negative social behaviors



How do we define **Social Behavior**?

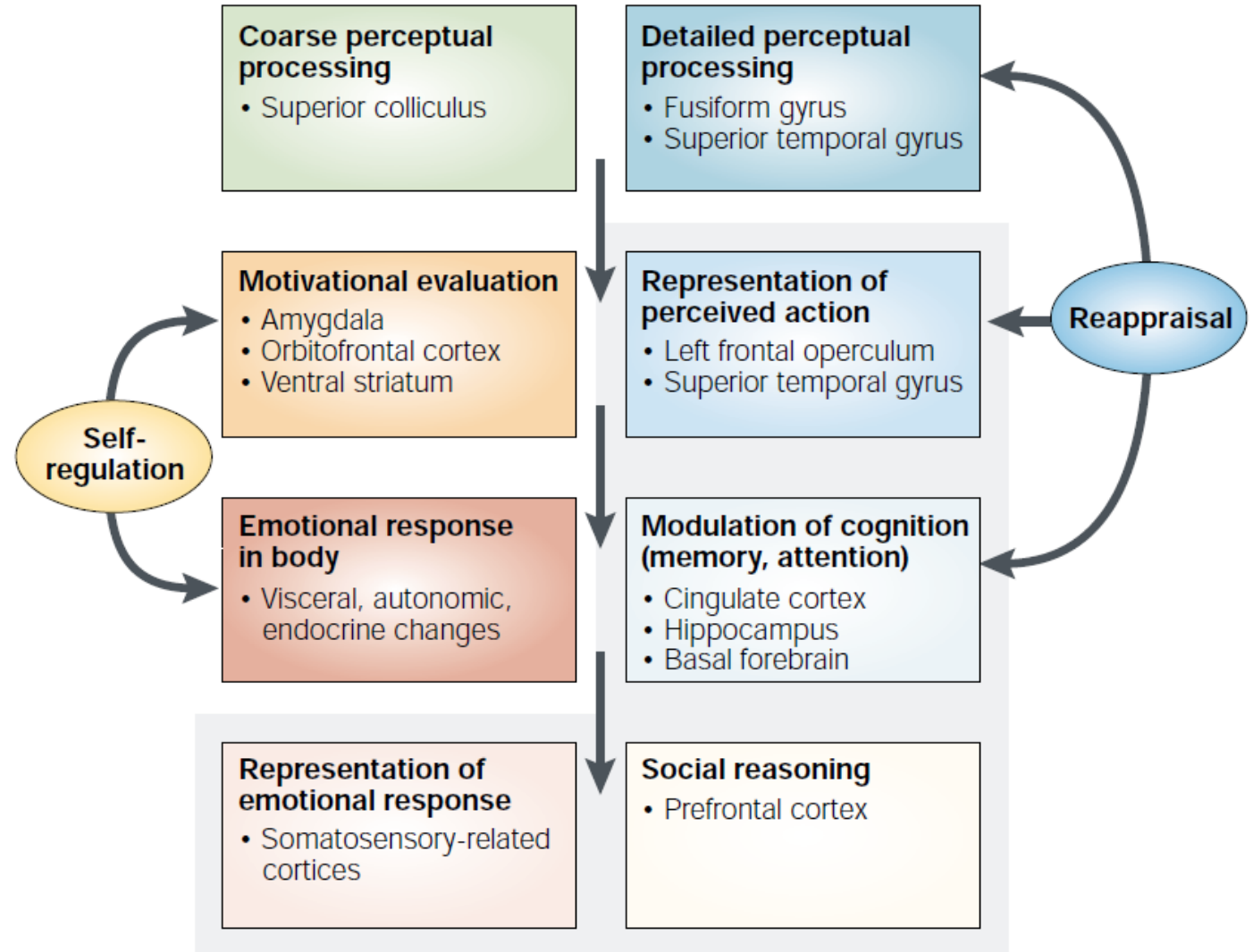






Social Behavior

Human Social Cognition



Why study social behavior in FASD?

- Social behavior deficits are a pervasive feature across the entire continuum of FASD



FAS

Partial FAS

ARBD and ARND

- Deficits also observed in 'sub-clinical' alcohol-exposed groups ('PAE')
- Impact can become more pronounced with age (adolescence)

Why study social behavior in FASD?

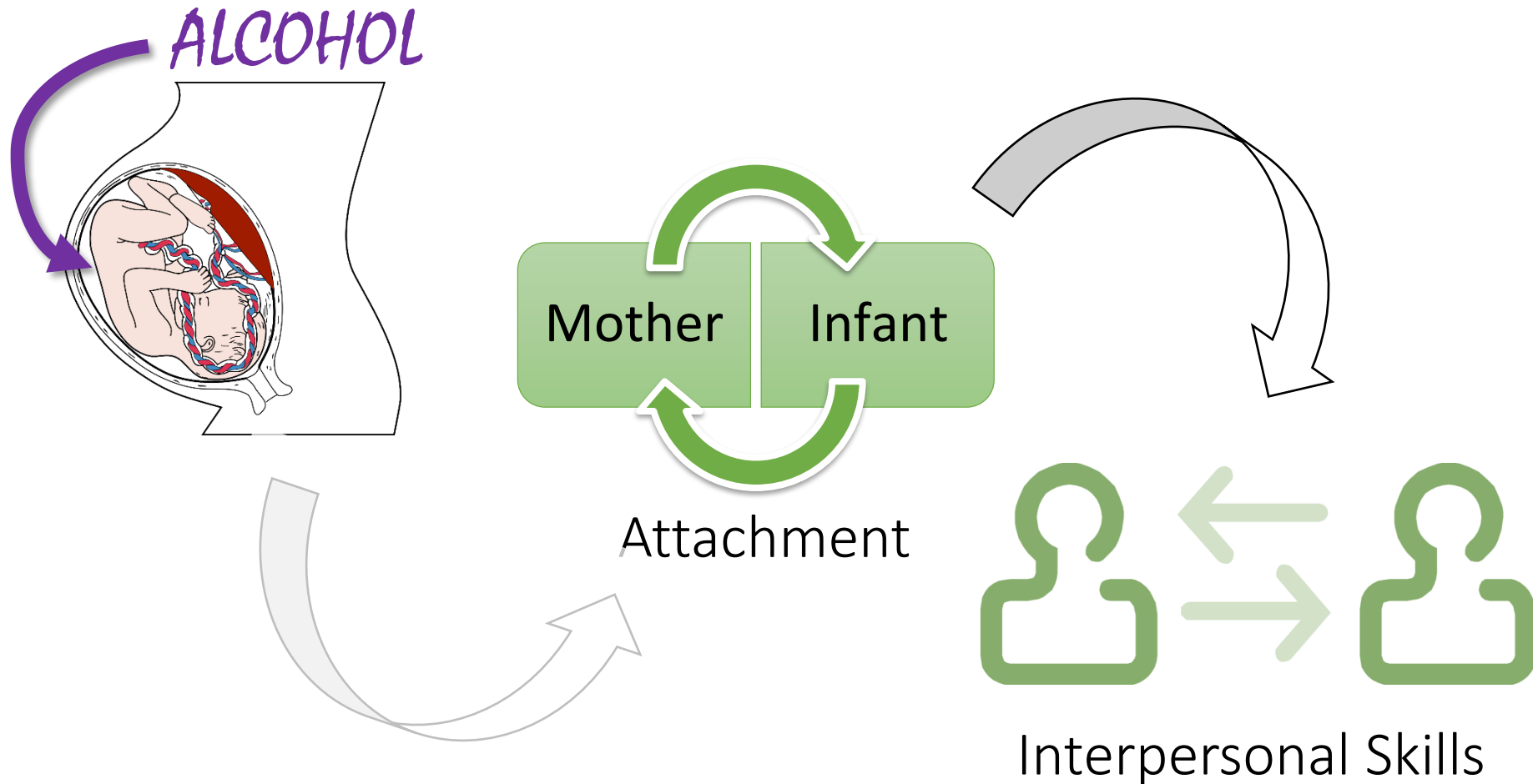
- Impaired social behavior in individuals with FASD has widespread implications for other domains and may contribute to:

- Difficulties in school
- Social rejection
- Trouble with the law
- Later mental health problems
- Independent living & work

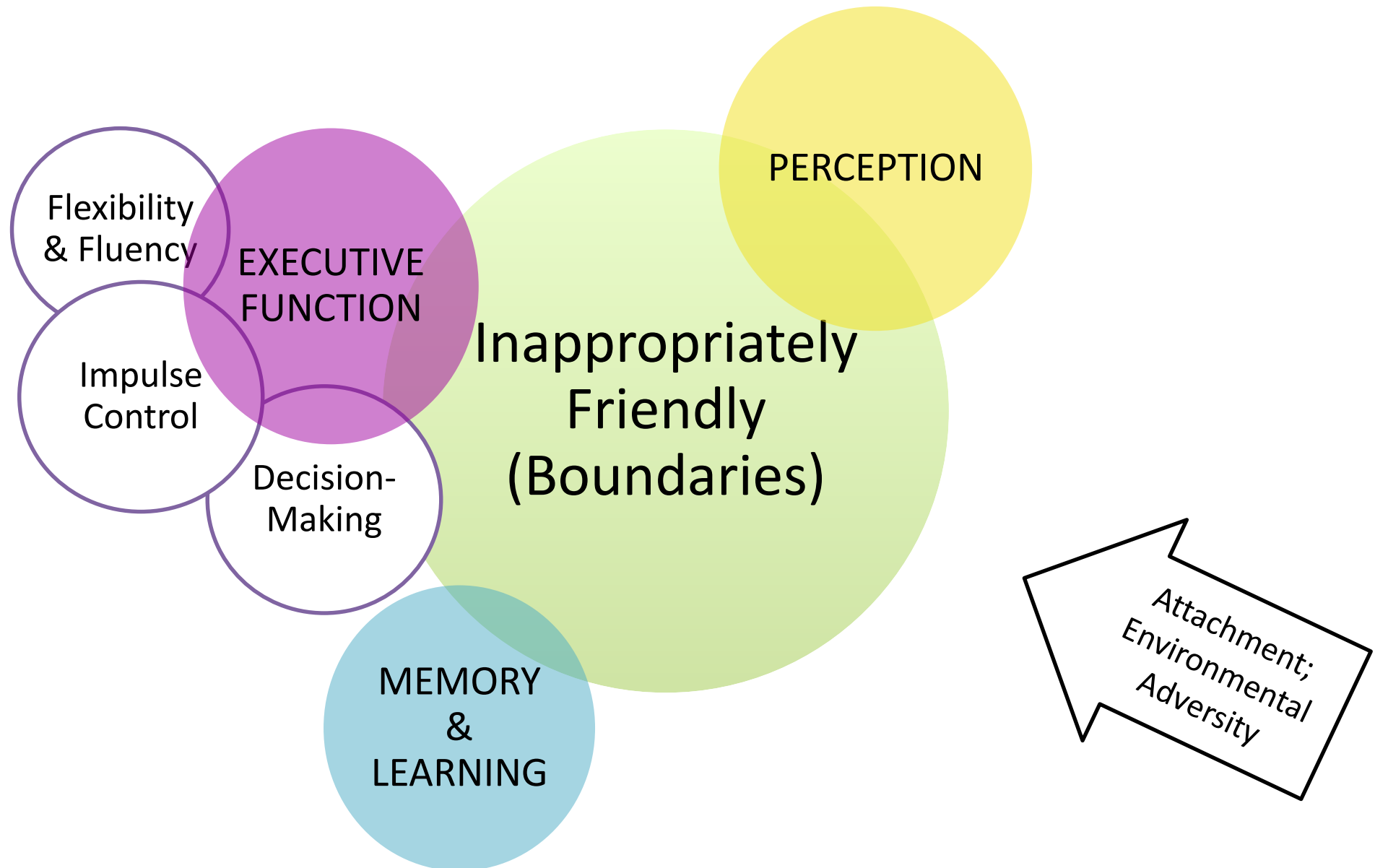


“Secondary Disabilities”

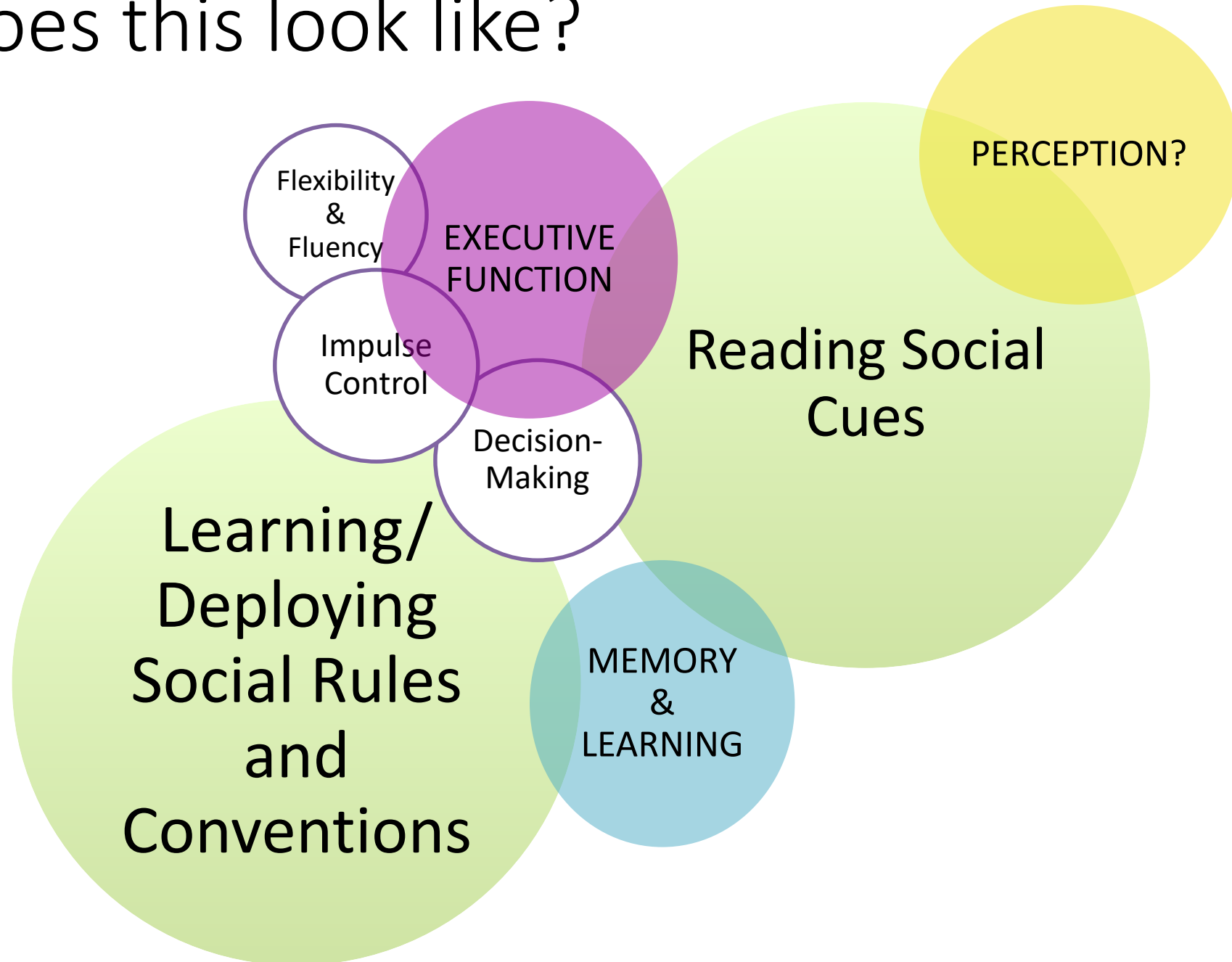
Early environment shapes development of social behavior



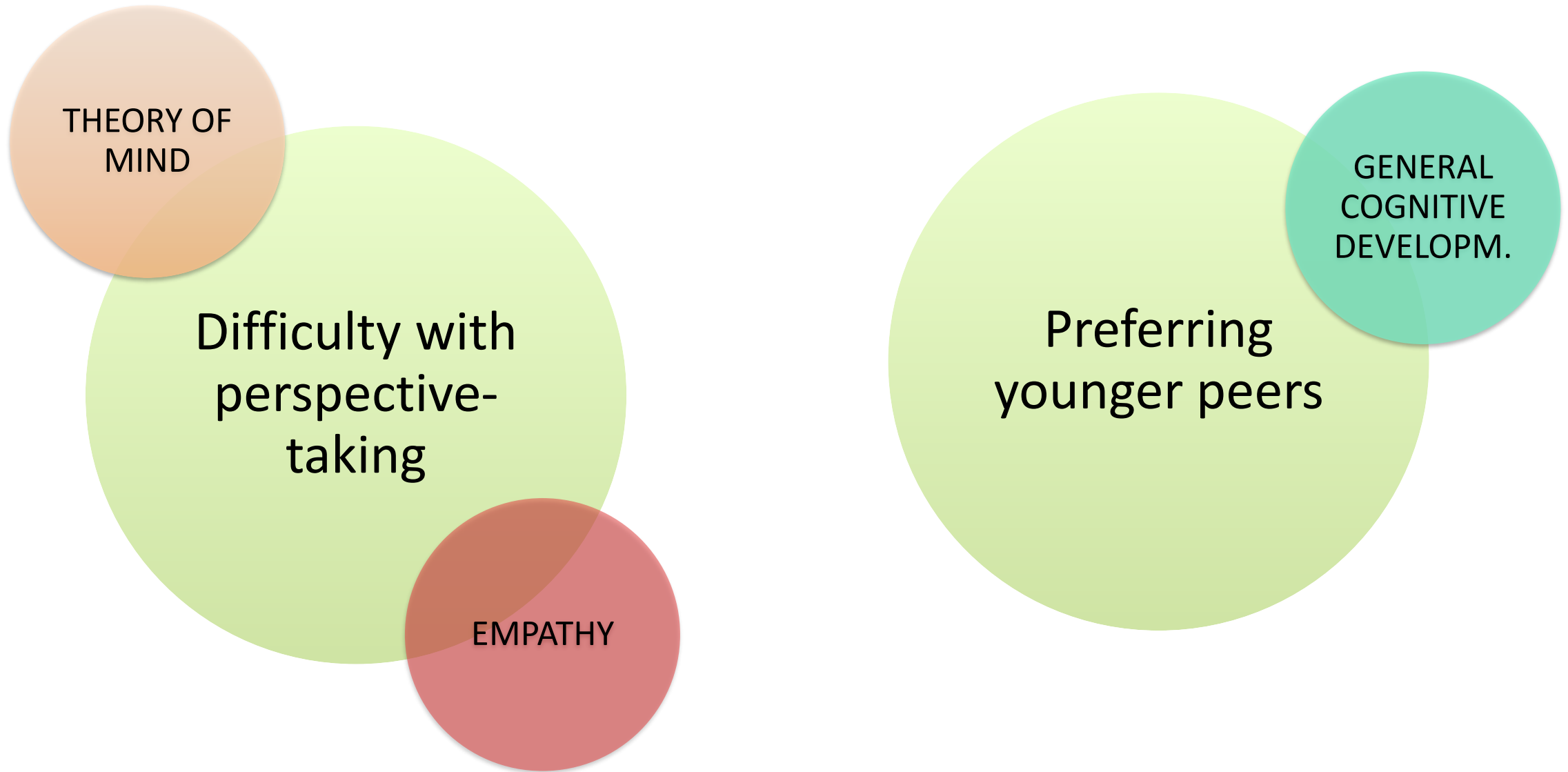
What does this look like?



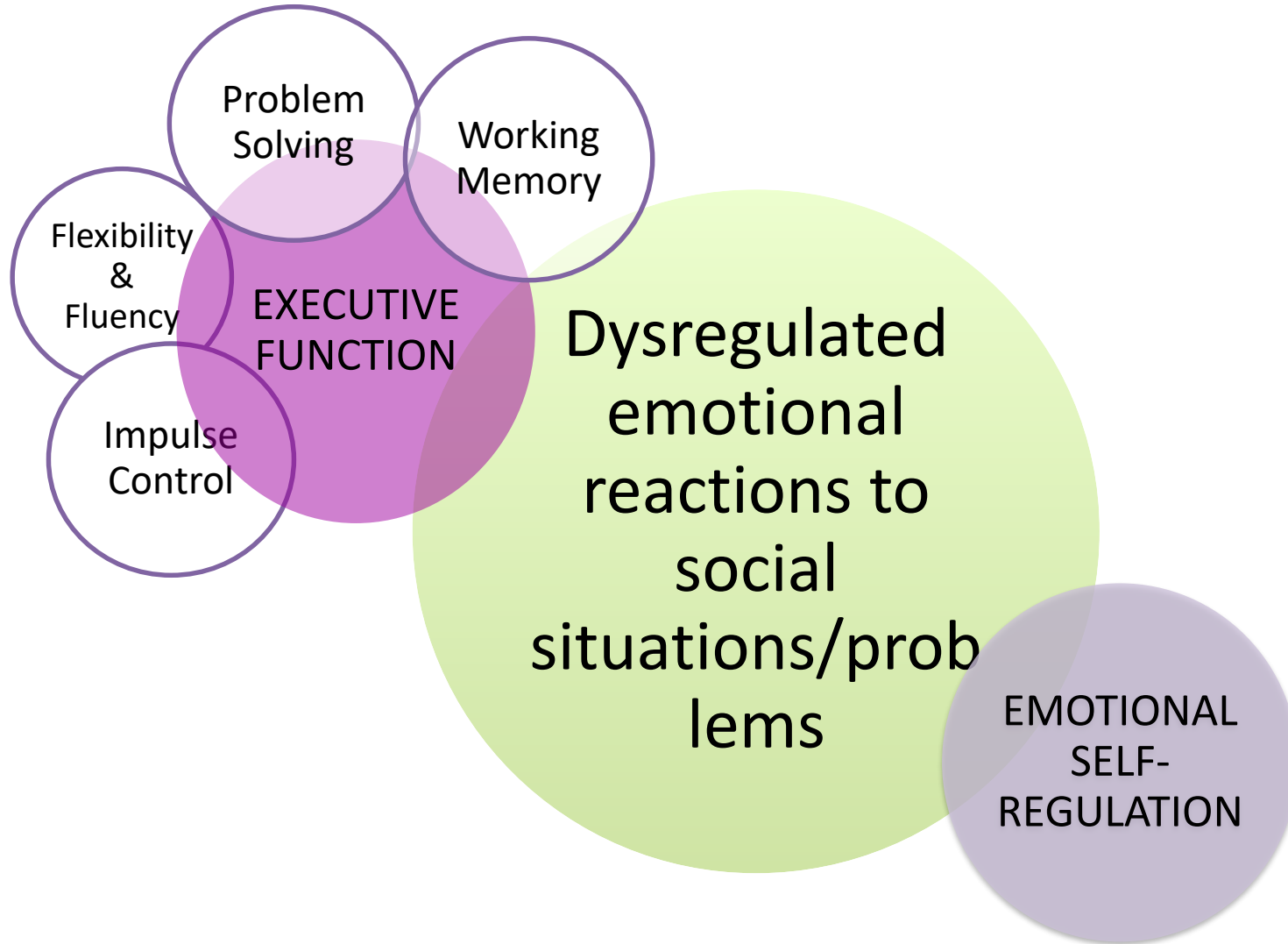
What does this look like?



What does this look like?



What does this look like?





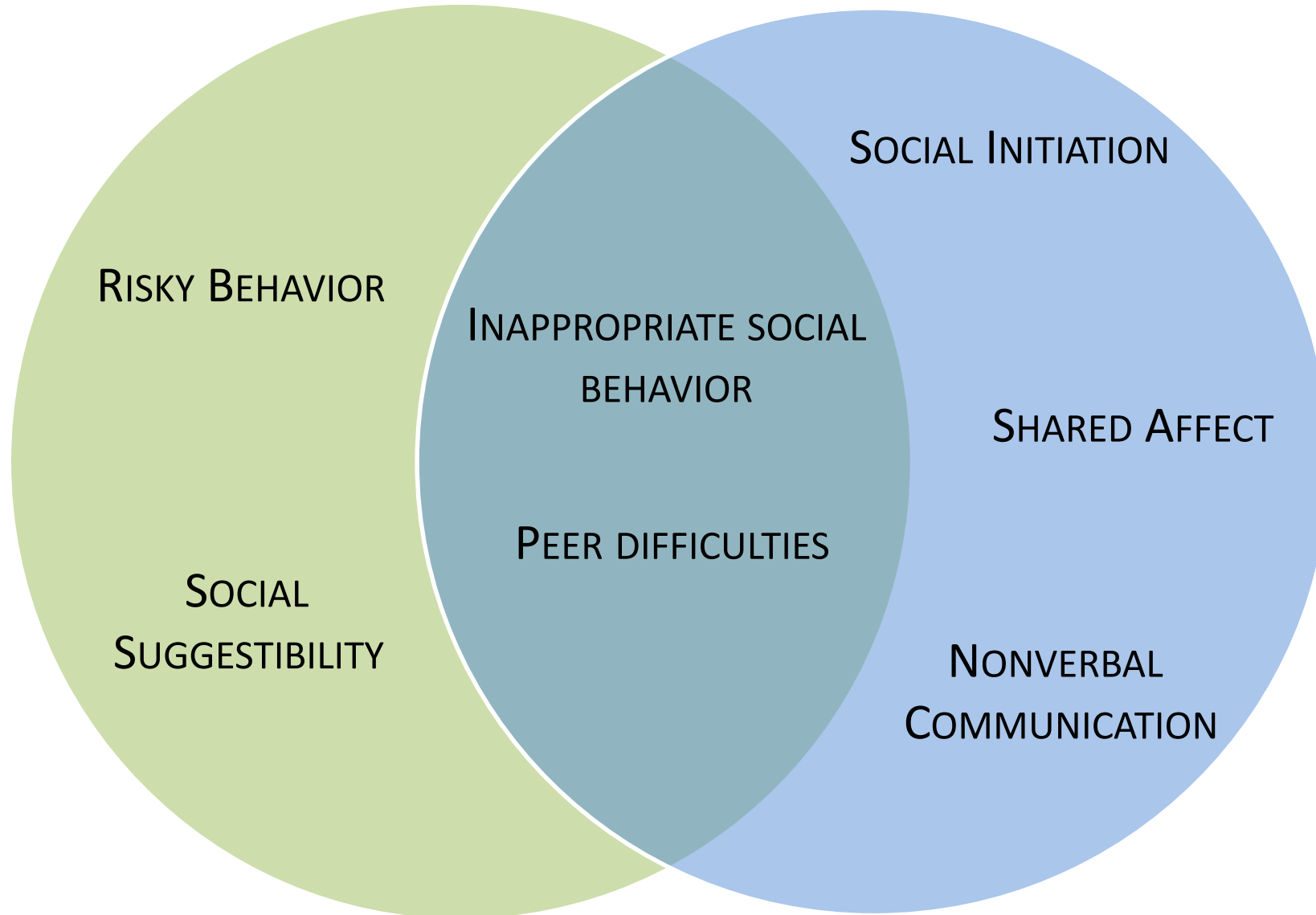
Social Strengths

- Interest in others
 - Enjoy being social (Duquette et al., 2006)
- Caregivers have described their children with FASD as:
 - Being friendly, talkative, and charming (James et al., 2010)
 - Having a loving/caring personality and determination and energy (Breen & Burns, 2012)
- Mean scores in the normative average range on the SSRS: Communication, Assertion, Engagement (Regehr, 2015)



FASD

ASD



Children with FASD show changes in social behaviors across development

Infancy	Periadolescence	Adolescence / Adulthood
<ul style="list-style-type: none">• Disruption in Mother-Infant Dyad (O'Connor et al. 1992; 2006)• Disrupted sleeping/feeding rhythms; Increased irritability (Coles & Platzman 1993)• Less “social monitoring” behaviors (Jirikowic et al. 2016)	<ul style="list-style-type: none">• Low scores on social behavior assessments (Thomas 1998; O'Connor et al. 2006; Regehr 2015; Stevens et al. 2015)• Impulsivity (McGee et al. 2008)• High scores on “problem behavior” assessments (Stevens et al. 2015)	<ul style="list-style-type: none">• Low scores on social behavior assessments (Moore & Riley 2015)• Increased risk for interactions with criminal justice system (Streissguth et al. 1996; MacPherson & Chudly 2007)

Animal models corroborate clinical findings

Infancy	Periadolescence	Adolescence / Adulthood
<ul style="list-style-type: none">• Increased latency to nipple attach (Subramanian 1992)• Altered suckling behavior (Barron, Kelly, & Riley 1991)• Inability to elicit retrieval by mother (Ness & Franchina 1990)• Altered pattern of ultrasonic vocalizations (Marino et al. 2002)	<ul style="list-style-type: none">• Altered play behavior (Hamilton et al. 2010; Lawrence et al. 2008)• Decreased social investigation (Mooney & Varlinskaya 2011)	<ul style="list-style-type: none">• Reduced social interactions (Hellemans et al. 2010)• Sexually dimorphic effects of social memory encoding and duration (Kelly, Leggett, & Cronise 2009)

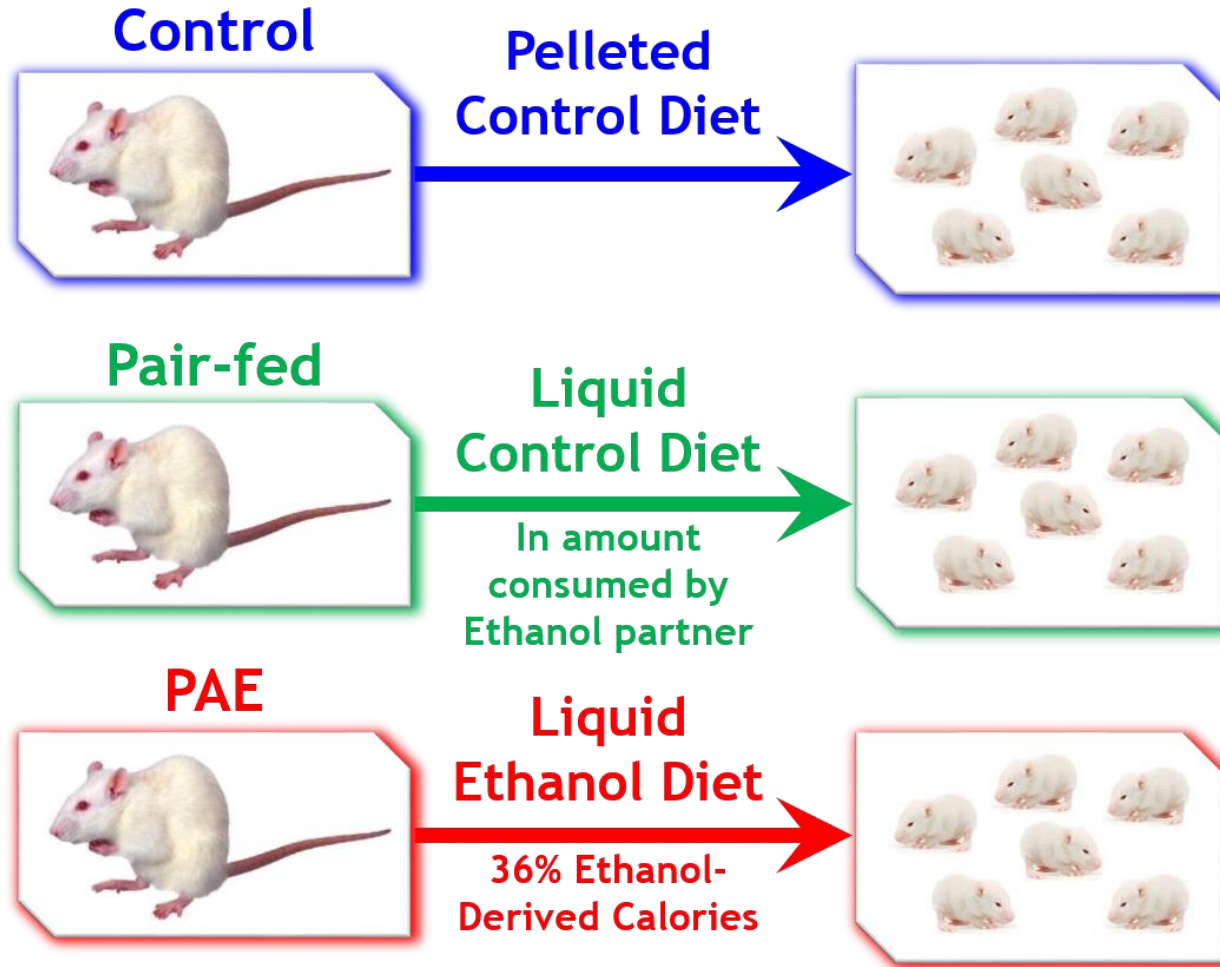
Why use animal models?



- Control for many variables:
 - Dose of alcohol
 - Timing of alcohol exposure
 - Environmental factors
 - Sex of offspring
- Ability to look at mechanism
 - Investigate changes in behavior and correlate with changes in brain



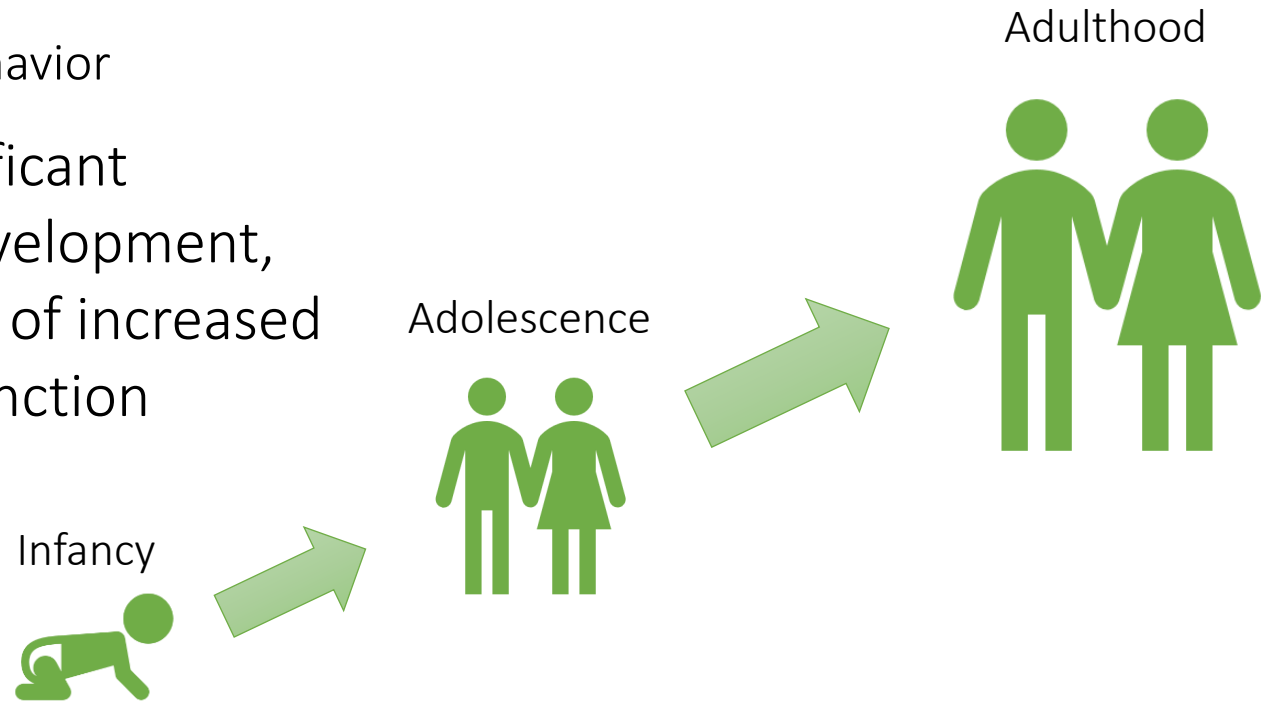
Animal model of prenatal alcohol exposure



- Diets maintained throughout gestation
(human 1st- and 2nd-trimester equivalent)
- Peak blood alcohol
≈150 mg/dL
- Offspring Ages at Testing:
 - Early Adolescence (~P30)
 - Late Adolescence (~P45)

Adolescence and Social Behavior Development

- Adolescence is a critical developmental period in which significant behavioral, cognitive, physiological and neurological changes occur, including sexual maturity (puberty)
 - Sex Hormones – Organize brain and behavior
- Maturational changes can have significant consequences for social behavior development, making adolescence a unique period of increased vulnerability to social behavior dysfunction



Infancy



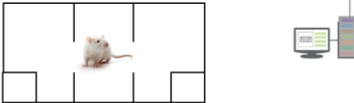
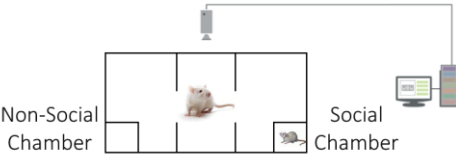


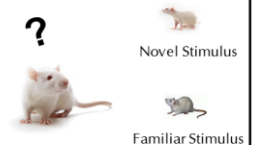
Adolescence



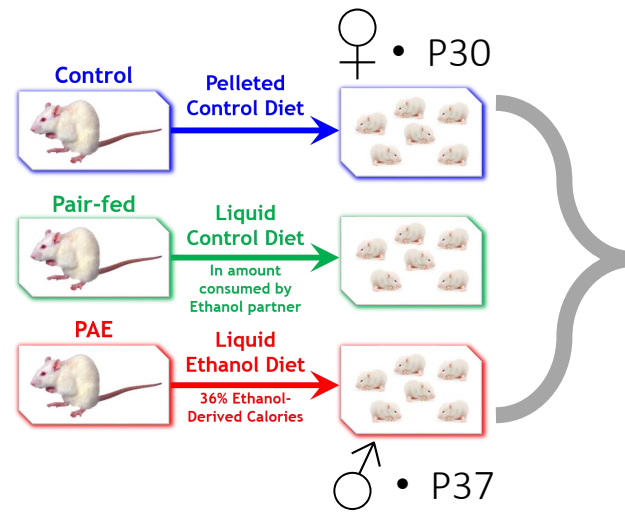
Adulthood



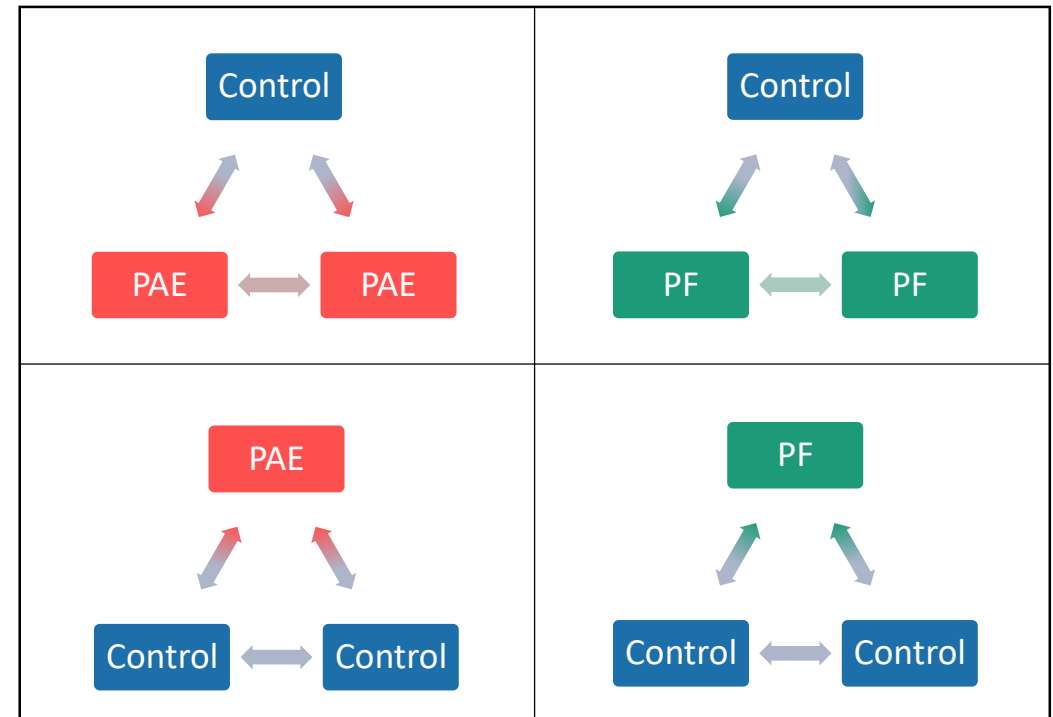
Effects of PAE on social behavior during pre-pubertal and pubertal periods of adolescent development

Social Motivation	Social Play	Social Recognition	Social Discrimination
<p data-bbox="280 656 420 714">Habituation (5 min)</p>  <p data-bbox="280 935 420 992">Testing (10 min)</p> 		<p data-bbox="1324 635 1541 735">Trial 1 2 min</p> <p data-bbox="1388 739 1477 761">↓ 18 min</p> <p data-bbox="1324 765 1541 865">Trial 2 2 min</p> <p data-bbox="1388 869 1477 891">↓ 18 min</p> <p data-bbox="1324 895 1541 995">Trial 3 2 min</p> <p data-bbox="1388 998 1477 1019">↓ 18 min</p> <p data-bbox="1324 1023 1541 1123">Trial 4 2 min</p> <p data-bbox="1388 1128 1477 1149">↓ 18 min</p> <p data-bbox="1324 1153 1541 1253">Trial 5 2 min</p> <p data-bbox="1592 835 1847 892">Learning Phase Familiar Social Stimulus</p> <p data-bbox="1592 1142 1847 1199">Testing Phase Novel Social Stimulus</p>	<p data-bbox="2114 635 2280 664">Learning Phase</p>  <p data-bbox="2178 835 2229 849">5 min</p> <p data-bbox="2089 892 2191 963">15 min Retention Period</p> <p data-bbox="2178 878 2229 992">↓</p> <p data-bbox="2089 1028 2305 1056">Discrimination Phase</p> <p data-bbox="2102 1078 2140 1106">?</p>  <p data-bbox="2178 1220 2229 1235">5 min</p>

Play behavior preferences in adolescent rats with or without prenatal alcohol exposure



Play Behavior – Triads



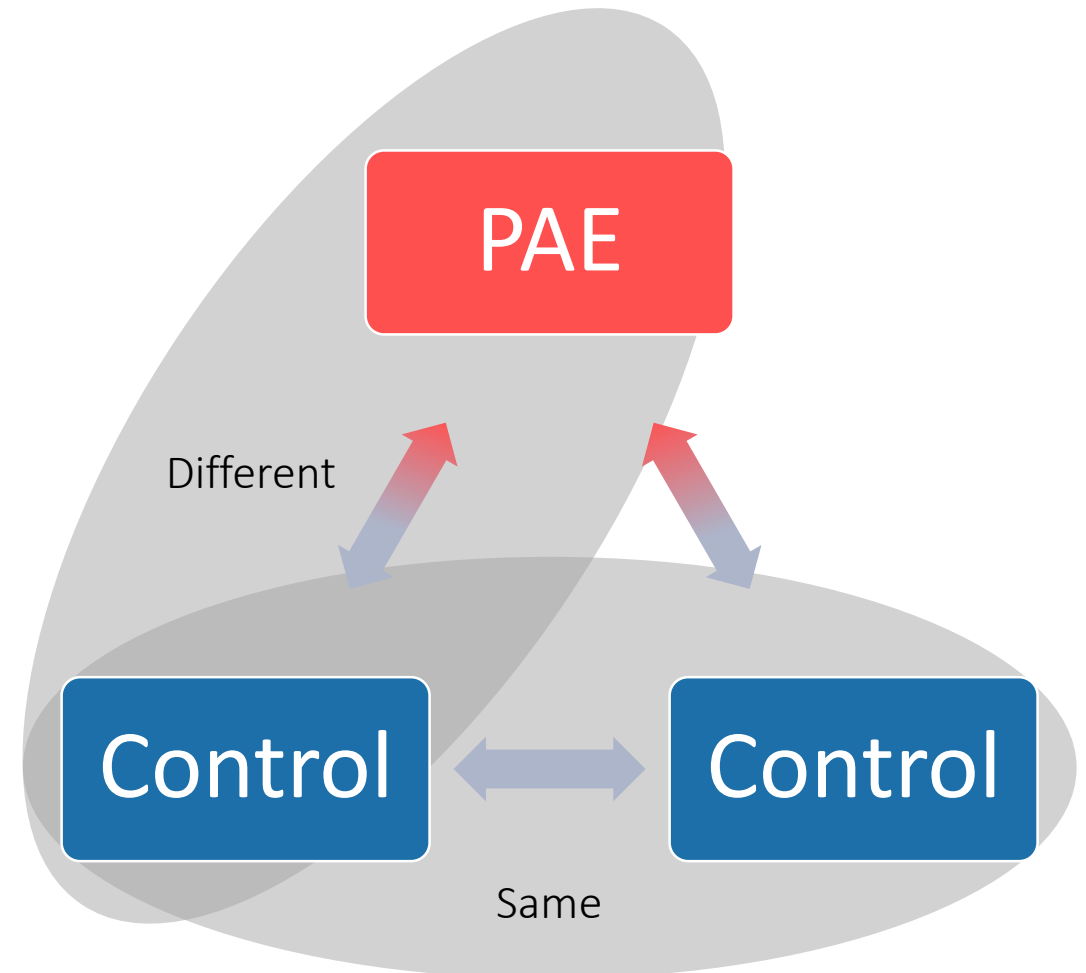
Samantha Baglot

Erin Morgan

Play behavior preferences in adolescent rats with or without prenatal alcohol exposure



Play Behavior – Triads



Samantha Baglot

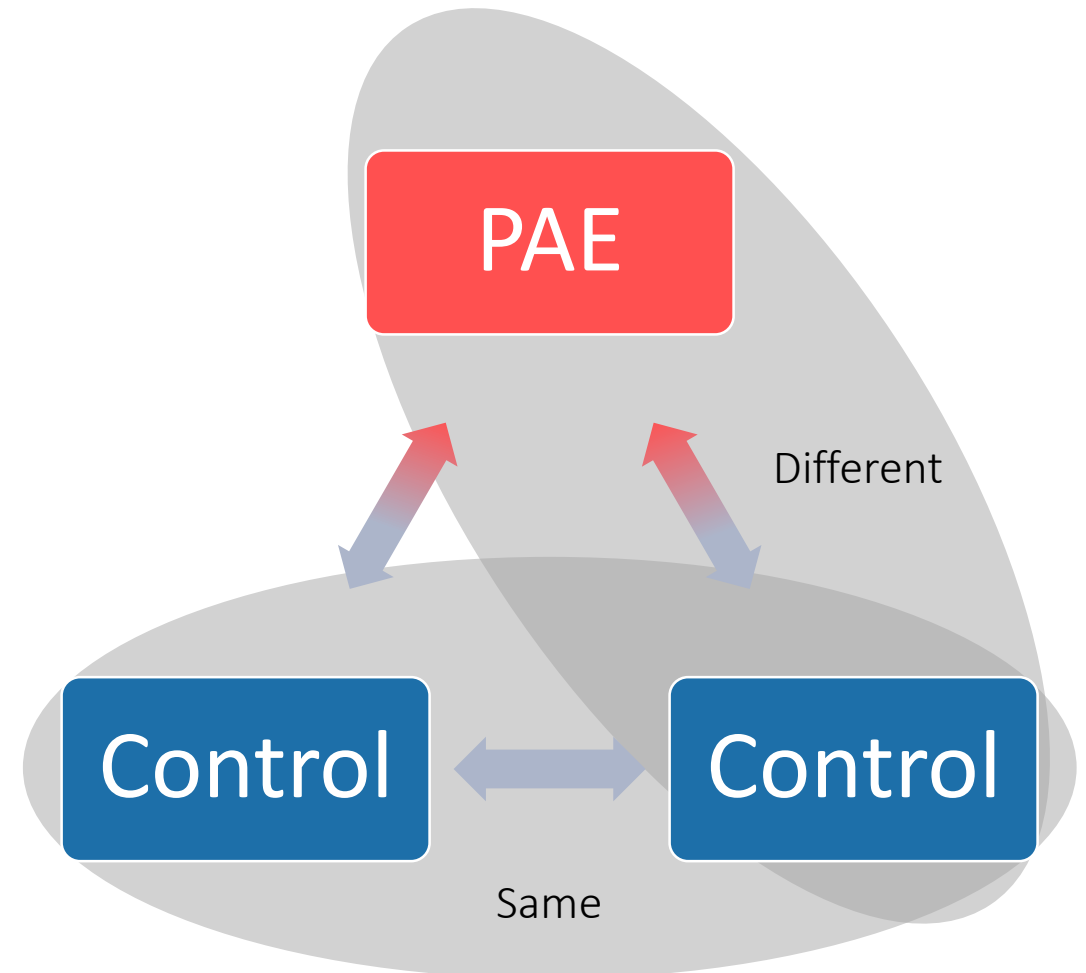


Erin Morgan

Play behavior preferences in adolescent rats with or without prenatal alcohol exposure



Play Behavior – Triads



Samantha Baglot



Erin Morgan

Summary

- Social motivation **not** altered in animals follow prenatal alcohol exposure
- In play triads, control and PAE animals bias their play towards controls in sexually dimorphic ways
- Normal development of social recognition memory is delayed in PAE adolescent males
- Social discrimination is impaired in adolescent males

Social Behavior & Oxytocin

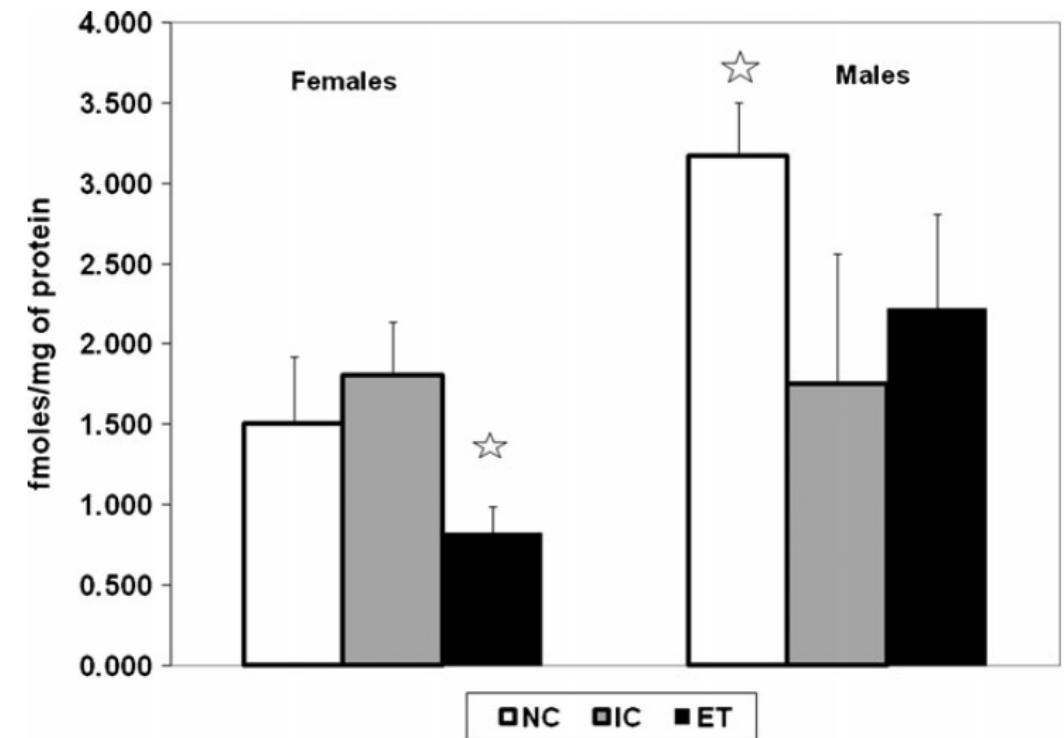
- Oxytocin
 - Uterine contractions
 - Lactation
 - Emotional feelings and responses
 - Social behaviors
 - Maternal behaviors and social bonds



Oxytocin and Prenatal Alcohol Exposure

- Adult rats show reduced OT receptor binding following PAE in amygdala
- Similar reductions in OT-IR cells observed in PVN & SON of adult female Mandarin voles following PAE (He, Zhang & Guo 2012)

Mean Oxytocin Receptor Binding in Amygdala



Summary

- Prenatal alcohol exposure (PAE) produces long-term changes in social behavior

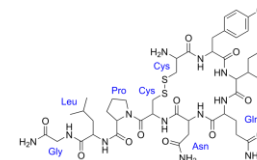
- Delays social behavior development
- Impairs social recognition memory
 - Particularly in complex social context



- PAE effects on social behavior are specific & sexually dimorphic
- PAE alters oxytocin system in the brain

- Next Steps

- Investigate oxytocin's potential to “rescue” social discrimination deficits following PAE



Questions???



Questions???



Intervention

	Children's Friendship Training Program (Keil et al., 2010; O'Connor et al., 2006; 2012)	Social Skills Intervention System (SSIS) (Regehr, 2015)
Type	Adapted established protocol for FASD	Non-adapted but customized manualized intervention
Target Age	Elementary school	Elementary school
Format	Group (child or parent)	Individual (child only)
Scope	<ul style="list-style-type: none">• Child-focused friendship training• Caregiver education	<ul style="list-style-type: none">• Direct skill teaching, role playing, video modeling for variety of social skills & problem behaviors
Research	<ul style="list-style-type: none">• Delayed treatment control• Significant gains in social knowledge, skills, problem behavior	<ul style="list-style-type: none">• Decrease in problem behaviors, but not dissimilar to contrast group
Community Translation	<ul style="list-style-type: none">• More effective than a community-based social skills intervention	No

Intervention

- Identify risk factors: Universal and population-specific
- Universal and population-specific factors that lead to more positive outcomes
- How can adverse outcomes be prevented?

Risk & Resiliency

- Resiliency = *competent* development during the course of negotiating, adapting to, or managing significant sources of stress and trauma
- Requires exposure to risk
- Protective factors
 - Individual, relational, contextual assets that facilitate adaptation
- Two dominant models:
 - Ecological
 - Constructivist

Risk & Resiliency



Key Protective Factors (Universal)



INTERNAL

Social Competence

Autonomy

Problem Solving

Sense of Purpose

Self Regulation



External

Caregiver Relationship

Peer Relationships

School Relationships

High Expectations

Opportunity

Resiliency: Key Points

- Requires exposure to risk
- Dynamic process, not outcome
 - Varies over lifespan
 - Not something that is necessarily ever ‘achieved’
 - Measurement?
 - Longitudinal research
- Primarily studied in the context of (otherwise) normative development
 - What about psychopathology?

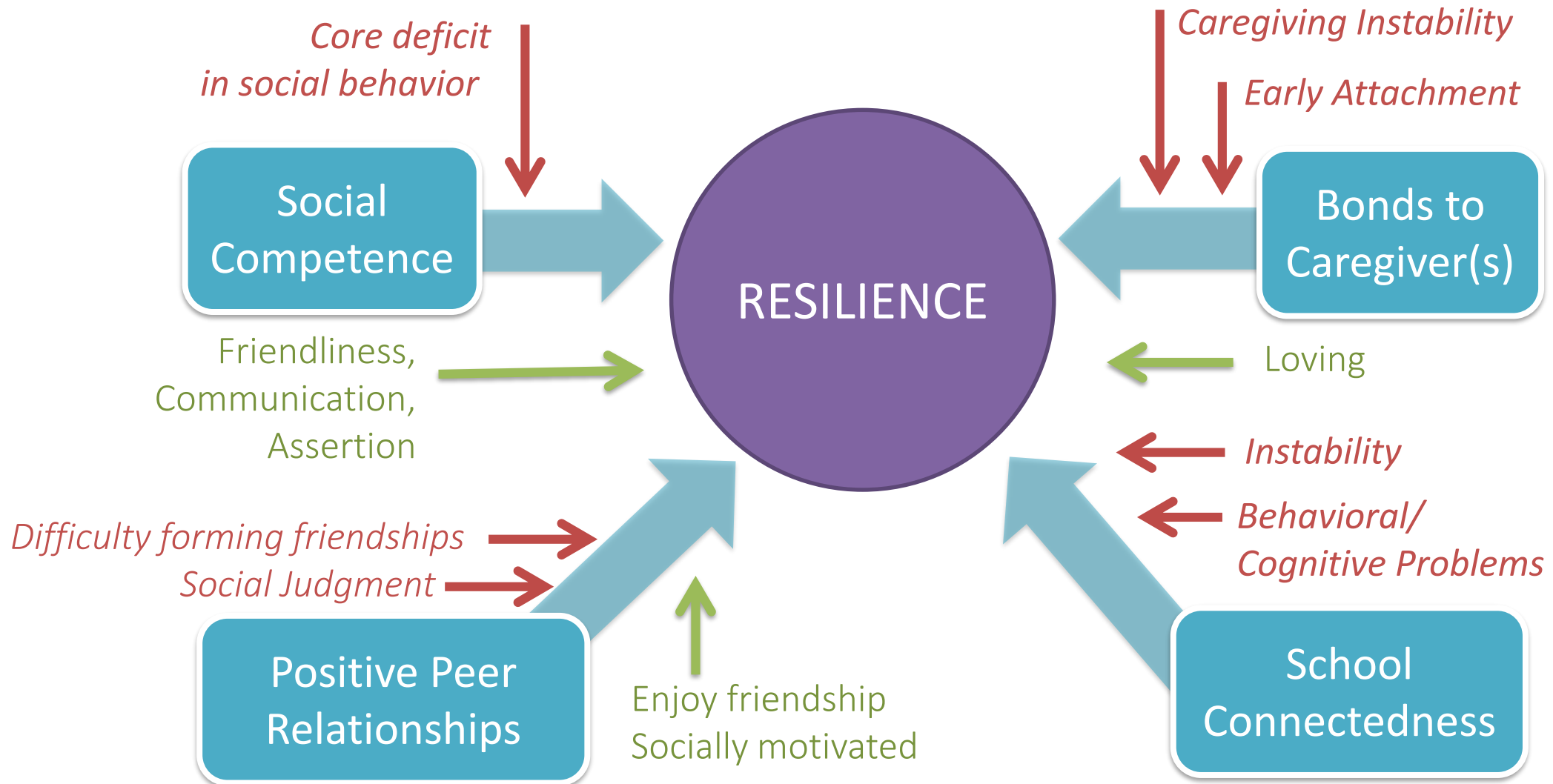


Risk, Resiliency, & FASD

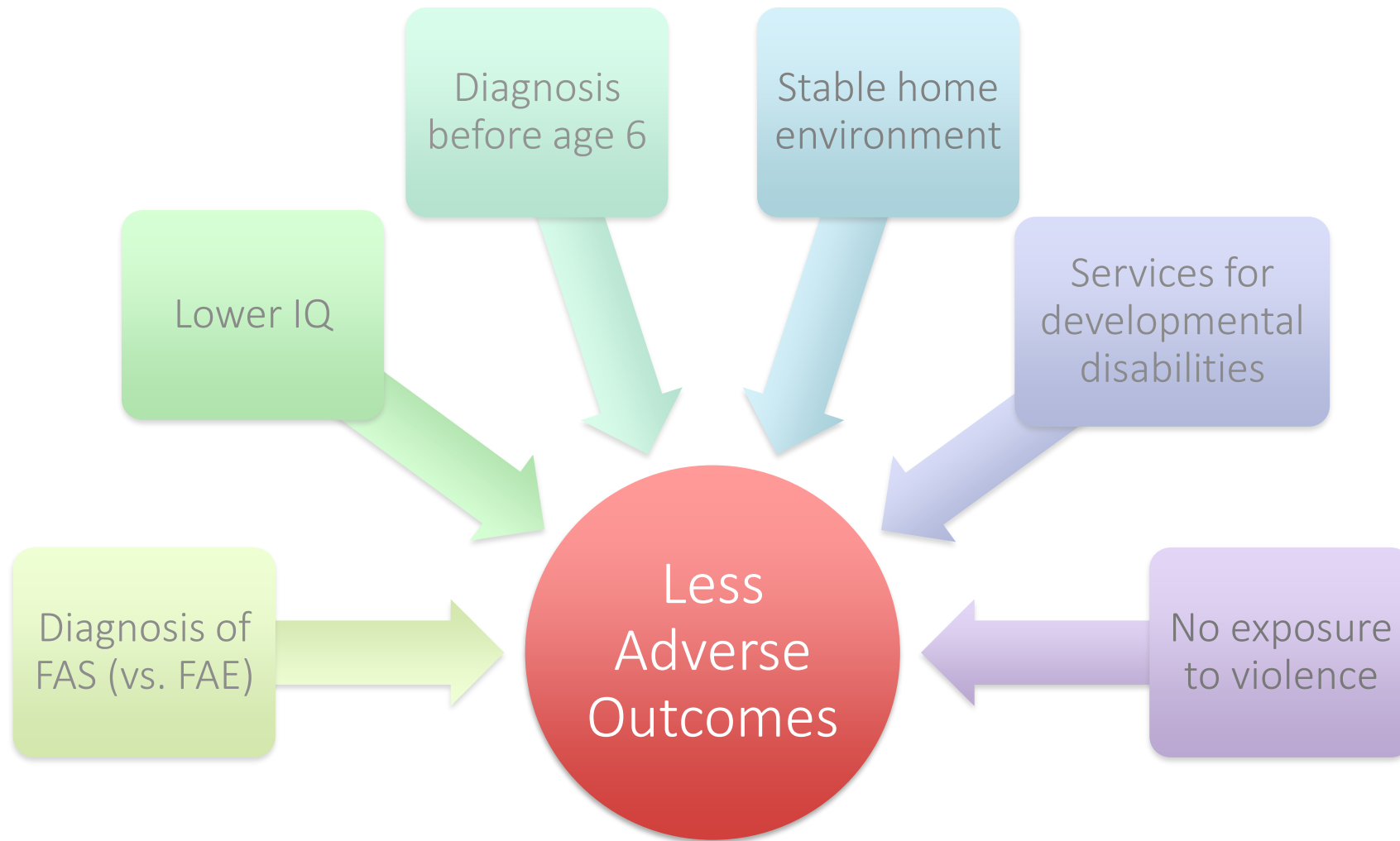
- “Double Jeopardy”
- Heterogeneity suggests resilience operating



Social Protective Factors



Key Protective Factors – FASD Specific



Conceptualizing Resilience in FASD

- Reconceptualization as a process
- Protective factors → resources (process-oriented)
- Outcomes
 - Point in time
 - What *is* competent development?
 - Within-group analysis of typical trajectories
 - Condition-specific vs. normative
 - Which domains should be considered?

Resilience & FASD

- Cohort of children, youth, and emerging adults assessed for FASD at the Glenrose Rehabilitation Clinic (Edmonton, AB)
- Tracking trajectories: mental health/behavior, adaptive behavior, executive function, working memory, etc.
- Child and Youth Resiliency Measure (CYRM)

From Research to Practice: Assessment

- Explicit consideration/quantification of resources
 - Resilience-informed assessment
- Hidden resilience in case conceptualization
- Consideration of strengths

From Research to Practice: Intervention

- Carmichael Olson, 2015
 - Identify commonly occurring areas of impairment ✓
 - Adapt existing EBPs
 - Appropriate to developmental stage
 - Social skills intervention for adolescents?
 - Incorporate an understanding of risk and protective factors

From Research to Practice: Intervention

- Key universal protective factor: Peer relationships; School Connectedness
 - E.g., School-based peer-mediated intervention
 - Directly or indirectly target social skills
- Key universal protective factor: Caregiver relationships
 - Prioritizing stability
 - Modified Parent-Child Interaction Therapy?
 - Families Moving Forward

Guiding Questions

- What is 'competent' (social) functioning for an individual with FASD? Normative, or 'better than expected?' If you work directly with kids with FASD, what are you aiming for?
- If resilience is a process, when does it begin?
- What unanswered questions do you still have?