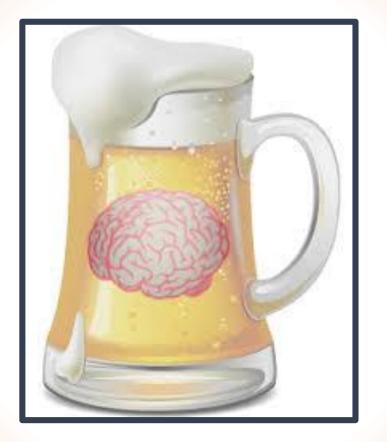
Neuropsychology of Fetal Alcohol Syndrome: Methods and Findings



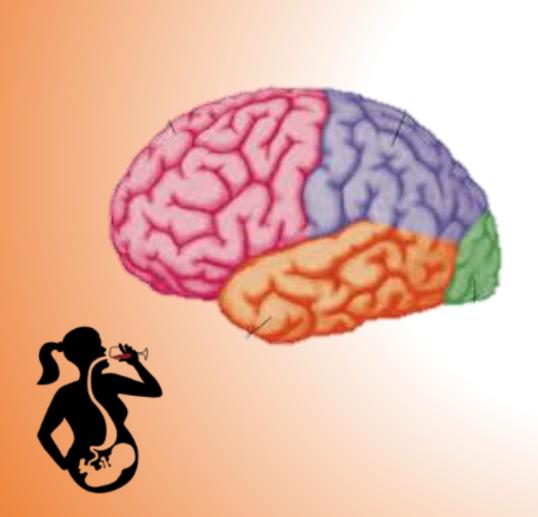
Anne Uecker PhD Child Neuropsychologist Edmund N Ervin Pediatric Center MaineGeneral Medical Center (Thayer Center for Health, Waterville)

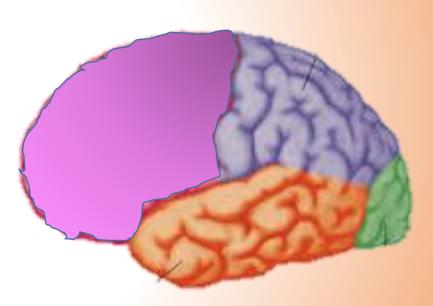
OBJECTIVES

- Identify neuropsychological (information processing deficits) associated with FASDs
- Become familiar with brain imaging findings in individuals with FASDs
- Identify disabilities that arise in the day to day functioning of individuals with FASDs
- Develop a sense of what's available and how to approach interventions for individuals impacted by FASDs.









Transgenerational Findings

--recent animal model findings

- Impact on information processing neural networks in not only alcohol impacted children, but has currently been shown to last for three generations.
- Increased risk for alcoholism, also currently shown for three generations.



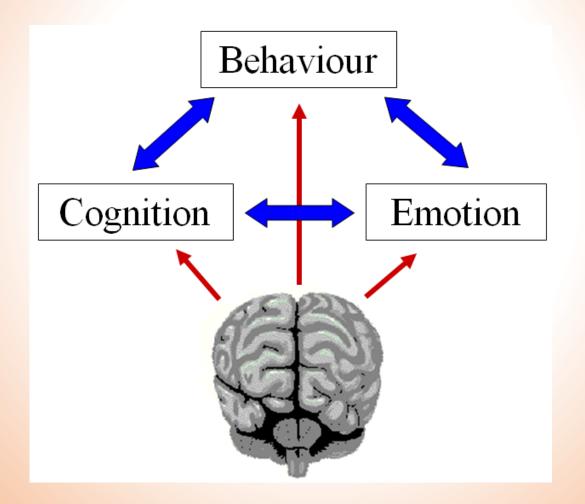




FASDs considered both a birth defect and a developmental disability

- <u>Birth defect</u>: a physical or biochemical abnormality that is present at birth and that may be inherited or the result of environmental influence.
- <u>Developmental Disability</u>: Significant long-term problems. They may be physical, such as blindness. They may affect mental ability, such as learning disorders. Or the problem can be both physical and mental. The problems are usually life-long, and can affect everyday living. Often there is no cure, but treatment can help the symptoms.

Neuropsychology of FASDs

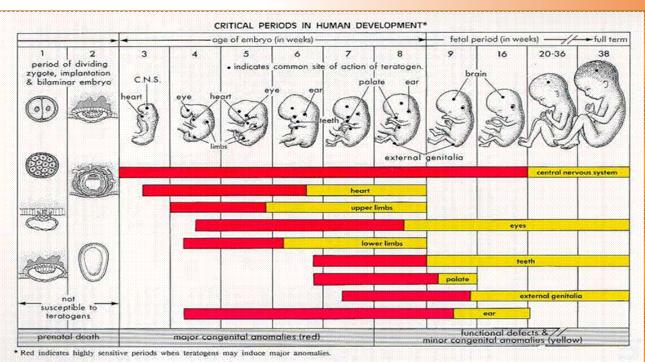




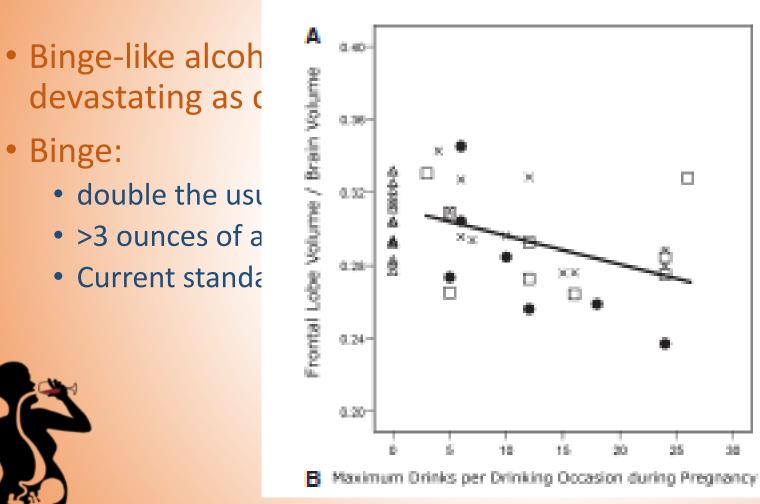
Fetal Alcohol Spectrum Disorder

- Amount/dose
- Timing
- Prenatal environment (care, nutrition)
- Genetics
- Subsequent postnatal environment





Binge Exposure



n spurt may be as ancy.





• Binge:

- 501 parent-children dyads in a Michigan-based university hospital.
- Children assessed at ages 6 to 7 years of age.
- Alcohol use during pregnancy:
- 25 % denied
- 64% Low levels of alcohol use
- 13% Moderate/heavy use
- The table emphasizes the complexity of factors involved.

• Sood et al 2001

TABLE 1. Demographic Characteristics by Alcohol Exposure Group								
Characseeistic				P Value				
			No	Low	Moderate/Heavy			
			(n = 117)	(n = 323)	(n = 66)			
	Child							
	Age		6.9	6.9	6.9	NS		
	Gerand (W M)		49.6	517	53.0	NS		
	dinth weight		3124.0	3024.5	2699.8	.000		
	Gestational age		38.6	38,9	37.7	.002		
	Curreré lead (µg/dL)		4.8	47	6.0	.007		
	Performance IQ		85.5	85.2	85.0	NS		
	Mother							
	Age		23.2	26.0	28.1	.000		
	Education		11.4	11.7	11.2	.030		
	Married (%)		32.8	27.0	21.5	NS		
	Cigarenes (number/d)		4.0	8.9	14.0	.000		
	Cocaine use (%)		18.8	452	69.7	.000		
	Current alcohol (oz AA/d)		0.04	0.4	0.5	.000		
	Current drugs (% use)		0.0	15	4.5	.061		
	Family							
	Cureody (biological mother		87.2	854	62.1	.000		
	Custody changes (% yes)		16.5	21.1	35.4	.011		
	Father lives with child (%)		28.4	215	15.4	.108		
	Faiher drinks (%)		45.2	80.3	89.2	.000		
	Faiher uses drugs (%)		21.7	37.7	44.6	.002		
	SES		30.7	29,9	25.1	.001		
	HOME inventory		32.8	31.8	28.9	.012		
	Violence exposure		14.1	13.3	13.7	NS		
	Maiernal depression		16.1	17.2	17.2	NS		
	SCL-GSI		0.5	05	0.5	NS		

P values from corresponding t or χ^2 analyses. NS indicates not significant; AA, absolute alcohol; SCL-GSL Symptom CheckEst-Global Severity Index.

The effect was observed at average levels of exposure of as low as 1 drink per week.

Children with any prenatal alcohol exposure were 3.2 times as likely to have Delinquent behavior scores in the clinical range compared with non-exposed children.

Prenatal alcohol exposure remained a significant predictor of behavior after adjusting for covariates.

Sood et al, 2001

Parameter	Р	P Value		
	No	Low	Moderate/ Heavy	
Externalizing	<mark>8.5</mark>	11.1	13.2	.002
Aggression	6.9	9.0	10.5	.003
Delinquent	1.5	2.1	2.7	.005
Internalizing	5.1	6.4	6.8	.105
Anxious/depressed	2.5	3.2	3.3	NS
Somatic complaints	1.1	1.2	1.5	NS
Withdrawn	1.7	2.2	2.1	NS
Neither externalizing nor internalizing				
Social problems	1.9	2.4	2.3	NS
Attention problems	3.2	3.7	4.4	NS
Thought problems	0.7	0.8	0.9	NS
Total score	23.7	29.0	32.2	.025

NS indicates not significant.

Mana CDCL David Communities 2 Communities 1 France TADIE o

- Though not selffulfilling. Substantial variability.
- Sood et al., 2001

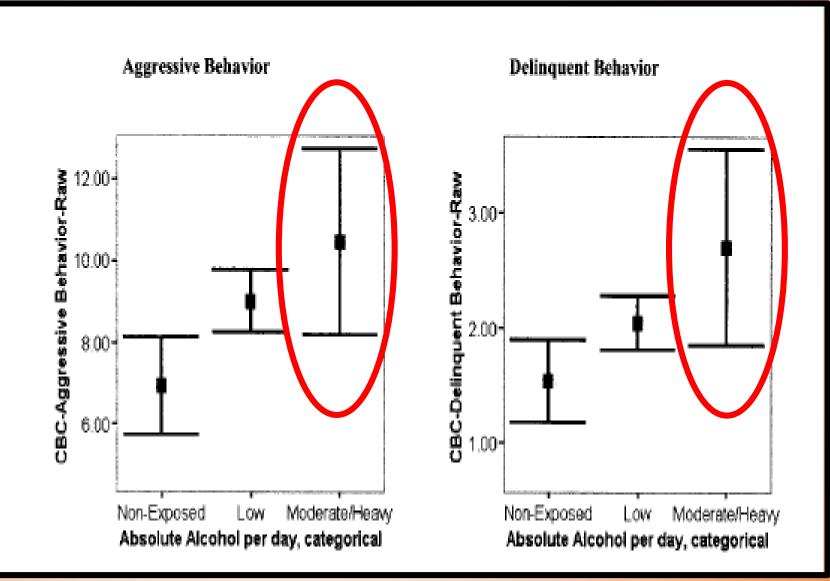
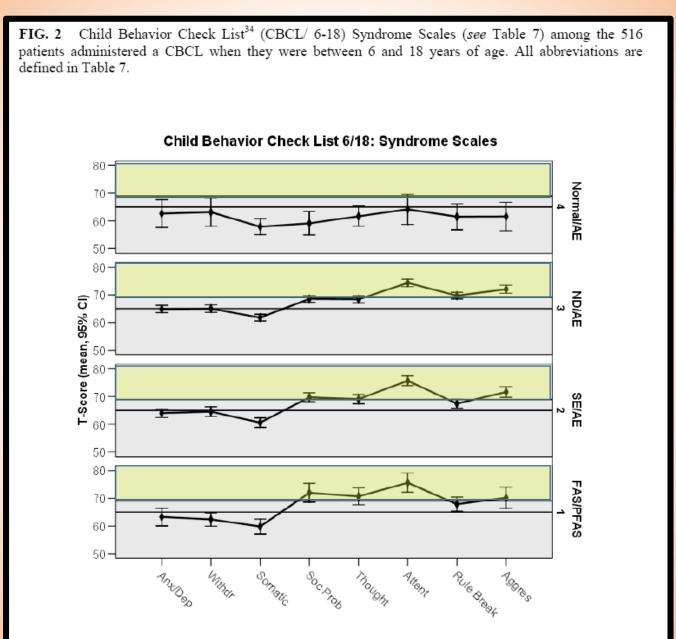




TABLE 7 Child Behavior Check List (CBCL/ 6-18) outcomes (see Figure 2) among the 516 patients administered a CBCL when they were between 6 and 18 years of age.

Washington: Astley 2010

	FASD Diagnostic Subgroups									Statistics		
Characteristic.	1. 59 FAS/ 95 PFAS N = 154			2. SE/AE		3. ND/AE		4. Normal CNS/AE		Total	ANOVA	
Characteristic										Total	Overall	Post Hoc
			N = 394		N = 722		N = 130		N = 1400		F (p) ^A	Duncan ^B
Problems: T-score ^C	Ν	Mean (SD)	Ν	Mean (SD)	Ν	Mean (SD)	Ν	Mean (SD)	Ν	Mean (SD)		
Internalizing	51	63.4(10.1)	154	64.5(10.9)	270	65.6(10.9)	25	60.8(14.1)	500	64.8(11.0)	1.9 (.14)	
Externalizing	<mark>51</mark>	<mark>69.1(9.9)</mark>	<mark>154</mark>	69.6(10.9)	270	70.8(10.3)	<mark>25</mark>	60.3(13.2)	<mark>500</mark>	69.8(10.8)	7.6 (.000)	123,4
Total	51	71.4(8.9)	154	71.3(9.3)	270	72.1(9.0)	25	61.9(12.7)	500	71.3(9.5)	9.1 (.000)	123,4
Syndrome Scales: T-score ^D												
Anxious/Depressed	51	63.0(11.3)	153	64.0(9.9)	269	64.9(10.9)	25	62.6(12.1)	498	64.3(10.7)	0.8 (.53)	
Withdrawn/Depressed	50	62.4(8.6)	153	64.6(11.2)	269	65.0(11.1)	25	63.1(12.4)	497	64.5(10.9)	0.9 (.42)	
Somatic Complaints	51	60.0(9.3)	153	60.6(10.8)	269	61.8(10.0)	25	57.9(7.0)	498	61.0(10.1)	1.6 (.19)	
Social Problems	50	72.0(12.0)	<mark>153</mark>	69.7(10.2)	<mark>269</mark>	68.5(10.2)	25	59.1(10.3)	<mark>497</mark>	68.8(10.7)	<mark>9.3 (.00</mark>)	123,4
Thought Problems	50	70.7(10.7)	153	69.1(10.6)	270	68.4(10.2)	25	61.6(8.8)	498	68.5(10.4	4.6 (.003)	123,4
Attention Problems	<mark>51</mark>	75.5(11.9)	<mark>153</mark>	75.7(11.0)	270	74.3(11.4)	<mark>25</mark>	<mark>64.2(13.1)</mark>	<mark>497</mark>	74.4(11.6)	7.6 (.000)	123,4
Rule-Breaking Behavior	<mark>51</mark>	<mark>67.9(8.9)</mark>	<mark>153</mark>	67.5(10.2)	<mark>269</mark>	<mark>69.7(10.0)</mark>	25	<mark>61.5(11.4)</mark>	<mark>498</mark>	68.4(10.2)	6.0 (.001)	123,4
Aggressive Behavior	50	70.2(13.1)	<mark>153</mark>	71.7(12.1)	<mark>269</mark>	72.0(12.2)	25	<u>81.6(12.5)</u>	<mark>497</mark>	71.2(12.4)	5.7 (.001)	123,4



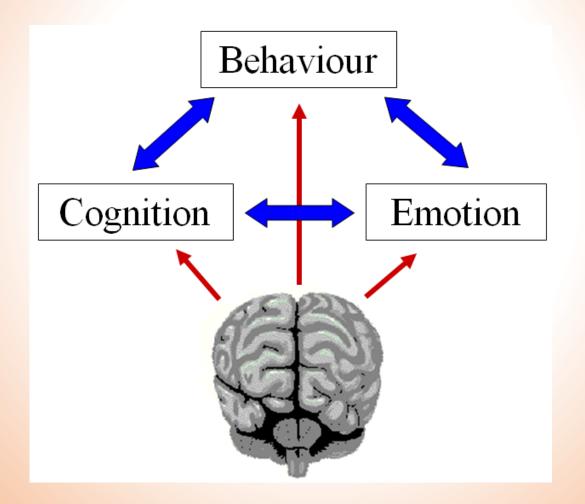
Astley et al 2010

TABLE 11 Mental health disorders reported in the medical records of the 1,064 patients 5 or more years of age at the time of the FASD diagnostic evaluation across the four study groups.

		FASD Diagnostic Subgroups										
Characteristic	1. 59 FAS/ 95 PFAS N = 154		2. SE/AE N = 394		3. ND/AE N = 722		4. Normal CNS/AE N = 130		Total N = 1400		Chi-square	
											Chi (p)	
Mental Health Disorders: N (valid%)												
One or more disorders	73	71.6	180	84.1	293	74.0	10	28.6	546	74.5	56 (.00)	
ADD/ADHD	<mark>53</mark>	<mark>59.6</mark>	<mark>161</mark>	<mark>59.9</mark>	233	<mark>55.2</mark>	0	0	<mark>447</mark>	<mark>53.9</mark>	148 (.00)	
Adjustment Disorder	4	2.6	8	2.0	29	4.0	3	2.3	44	3.1	3.9 (.27)	
Antipersonality Disorder	0	0	0	0	1	0.1	0	0	1	0.1		
Anxiety Disorder	2	1.3	10	2.5	8	1.1	0	0	20	1.4	5.8 (.12)	
Reactive Attachment Disorder	6	3.9	19	4.8	27	3.7	2	1.5	54	3.9	2.9 (.41)	
Bipolar/Manic Depression	4	2.6	10	2.5	13	1.8	3	2.3	30	2.1	0.8 (.85)	
Conduct Disorder	2	1.3	16	4.1	24	3.3	1	0.8	43	3.1	5.3 (.15)	
Depression	7	4.5	23	5.8	32	4.4	2	1.5	64	4.6	4.2 (.24)	
Dysthymic Disorder	3	1.9	7	1.8	23	3.2	2	1.5	35	2.5	3.0 (.39)	
Obsessive Compulsive Disorder	1	0.6	6	<mark>1.5</mark>	2	0.3	0	0	9	<mark>0.6</mark>	6.5 (.09)	
Oppositional Defiant Disorder	8	<mark>5.2</mark>	<mark>39</mark>	<mark>9.9</mark>	<mark>72</mark>	<mark>10.0</mark>	1	<mark>0.8</mark>	<mark>120</mark>	<mark>8.6</mark>	<mark>15.0 (.00)</mark>	
Post Traumatic Stress Disorder	10	6.5	32	8.1	49	6.8	4	3.1	95	6.8	3.9 (.27)	
Suicidal	2	1.3	3	0.8	5	0.7	0	0	10	0.7	1.7 (.64)	

Abbreviations:: Chi: chi-square test statistic across the 4 study groups. FAS: fetal alcohol syndrome. P: p-value. PFAS: partial FAS. ND/AE: Neurodevelopmental disorder/alcohol exposed. Normal CNS/AE: No central nervous system abnormalities/alcohol exposed. SE/AE: Static encephalopathy/alcohol exposed.

Neuropsychology of FASDs





Neurocognitive (The Holy Grail)

- Executive Function
 - Abstraction/judgment problems
 - Lack of control over emotions
 - Impulsivity
 - Inappropriate/immature social behaviors
 - Difficulty learning from consequences
- Attention (hallmark)
- Visual Spatial/Visual Motor
- Learning and Memory
- Motor



- Achievement (Arithmetic/Inability to manage money)
- Adaptive (Greater than expected day to day deficits given their IQ)

How Researchers Look at the Brain

Neuropsychological Testing

- Tests of information processing
- <u>Structural MRI findings</u>
- smaller total brain volume
- smaller volume of both the white and grey matter in specific cortical regions.
- consistent findings of alterations in the shape and volume of the corpus callosum, as well as smaller volume in the basal ganglia and hippocampi.

Functional MRI

• reduced functional connectivity between cortical and deep grey matter structures.

Proton magnetic resonance spectroscopy

• altered neurometabolic profiles in the frontal and parietal cortex, thalamus and dentate nuclei.

Diffusion tensor imaging

• lower fractional anisotropy in the corpus callosum.

Donald et al., 2015

Executive Function Deficits in FASDs and ADHD

Kingdon et al, 2016

FASD compared to no diagnosis

Dysmorphic FASD

- large deficits in planning, fluency, setshifting, and working memory
- Moderate deficits in vigilance and inhibition

Non-dysmorphic FASD

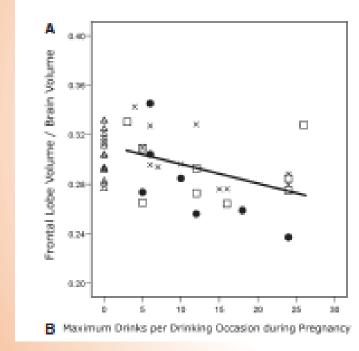
- large deficits in planning, fluency and set shifting
- moderate deficits in working memory
- Nonsignificant difference vigilance and inhibition

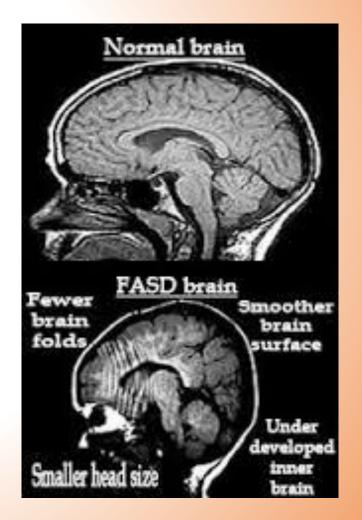
great deal of neuropsychological variability within ADHD groups

- fewer than half of children with ADHD exhibiting significant impairment on any specific executive function task
- FASD does not produce an identical cognitive profile to ADHD.

FASD compared to ADHD

- moderate to small deficits in fluency, planning, and set-shifting
- Small deficits on working memory
- no differences were found on measures of attentional vigilance or response inhibition





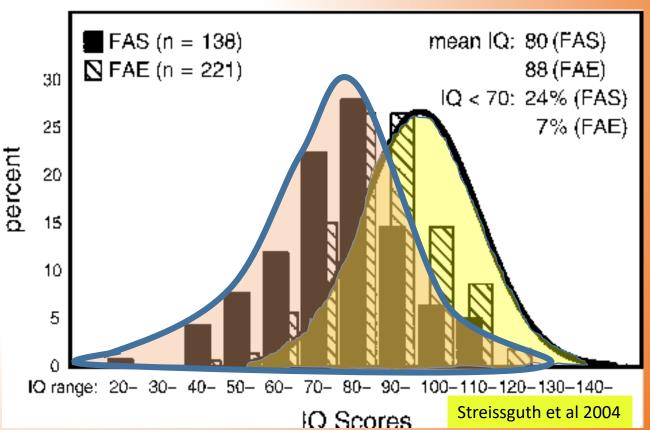
Large IQ impairments in

- FASD versus healthy controls 20 IQ points
- FASD versus children with ADHD 16 IQ points
- Children with FASD evidenced greater executive functioning impairments than healthy controls and ADHD groups, even among studies where groups were matched on intellectual functioning

Also

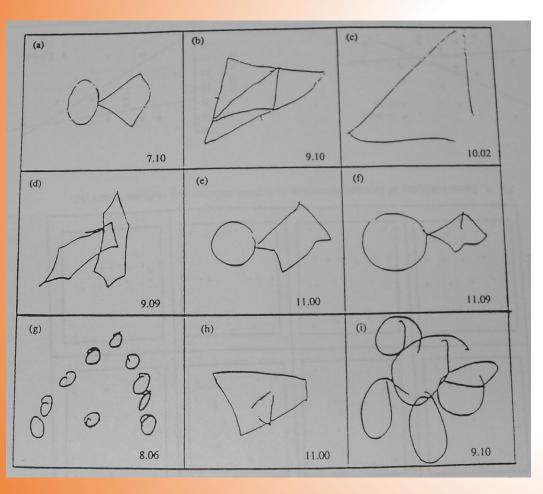
Kingdon et al, 2016

• Full-scale intelligence scores for FASD groups included in the current meta-analysis ranged from 64 to 99, with an average of 84 for both dysmorphic and nondysmorphic groups.



Drawing (VMI) data

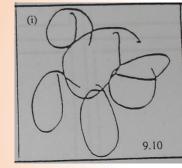
Uecker and Nadel, 1996



- Attention
- Vigilance
- Impulsivity
- Maturation
- Form
- Visual Motor
- Planning
- Organization



(c)

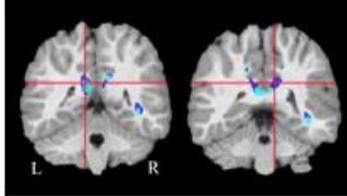


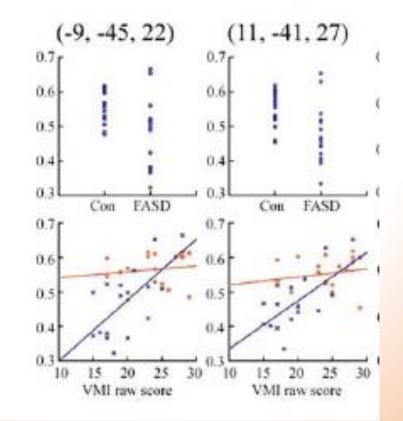
9.09

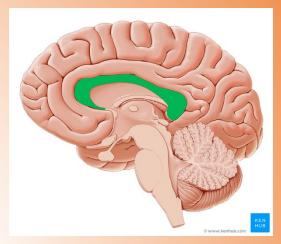
(d)

White matter abnormalities in brain areas associated with visual motor integration. Sowell, et al, 2008

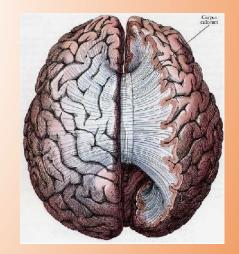
Lateral splenium





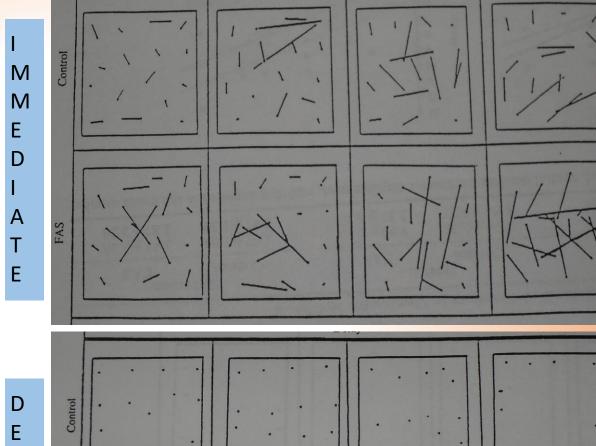




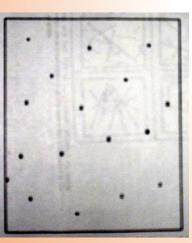


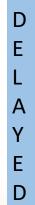
16 Object Spatial Localization/Distortion

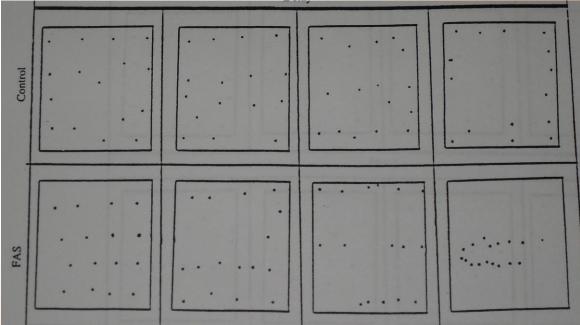
Uecker and Nadel, 1996





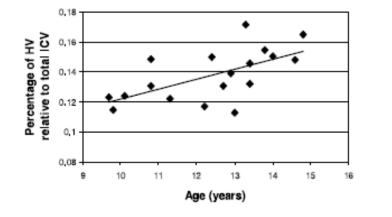


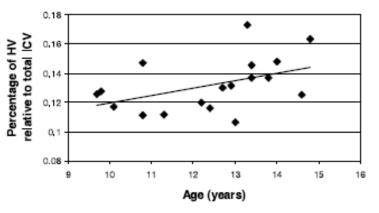


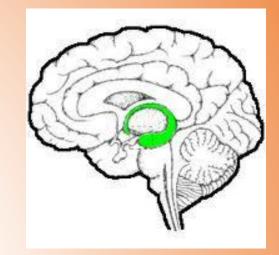


A. Left Hippocampal Volume by Age in Controls

B. Right Hippocampal Volume by Age in Controls







C. Left Hippocampal Volume by Age in FASD

Percentage of HV relative to total ICV

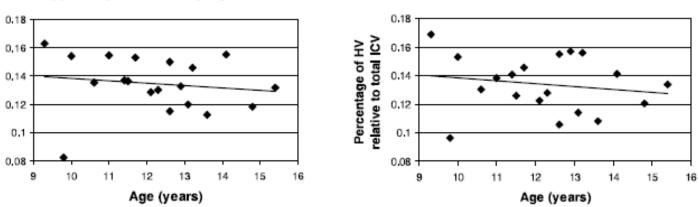


Fig. 3. Left and right hippocampal volume (presented as proportion of total intracranial volume) by age scatterplots for Controls (top graphs A and B) and FASD (bottom graphs C and D).

Willoughby et al, (2008). Effects of prenatal alcohol exposure on hippocampal volume, verbal learning, and verbal and spatial recall in late childhood.

D. Right Hippocampal Volume by Age in FASD

Verbal Learning and Memory

- The California Verbal Learning Test-Children's Version
- Children with FASD
 - learn the CVLT-C word lists more slowly than non-exposed children
 - recall fewer words
 - are less accurate at discriminating between target and distractor words on recognition trials
 - use less semantic clustering
 - Semantic Clustering reflects organization, specifically providing some insight on grouping, chunking, or categorizing abilities.

- Whole Brain
- Frontal Lobe
- Parietal region
- Corpus Callosum
- Cerebellum
- Basal Ganglia
- Hippocampus

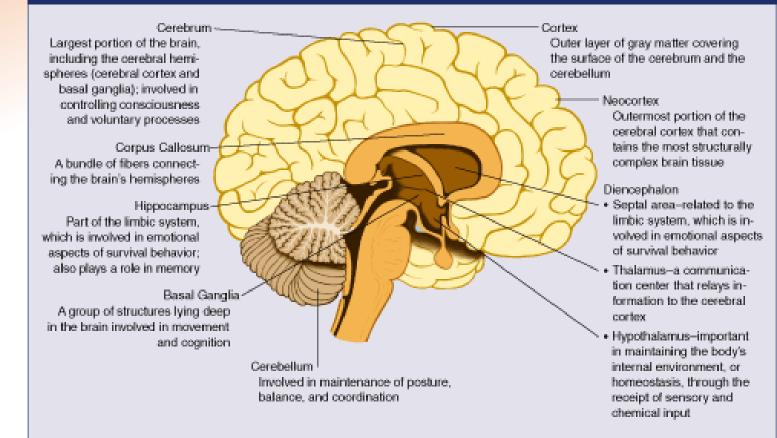
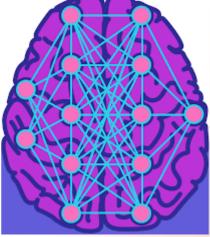


Figure from:http://people.uwec.edu/piercech/fas/fas...htm

Brain areas variable between individuals.

Impact on Neural Networks

 Dramatic changes in intra-neocortical connections between the <u>frontal, somatosensory and visual cortex</u> in mice born to mothers who consumed ethanol during pregnancy. The changes were especially severe in the frontal cortex, which regulates motor skill learning, decision-making, planning, judgment, attention, risk-taking, executive function and sociality.



H. El Shawa, C. W. Abbott, K. J. Huffman. Prenatal Ethanol Exposure Disrupts Intraneocortical Circuitry, Cortical Gene Expression, and Behavior in a Mouse Model of FASD. *Journal of Neuroscience*, 2013

Academics

• The alcohol-exposed children performed significantly worse than their peers in all academic areas, with particular weaknesses found in math performance.

Math reasoning < numerical operations < spelling and word reading.

- Over half of the alcohol-exposed group (58.2%) demonstrated low achievement on 1 or more academic domains.
- Brain imaging revealed several brain surface area clusters linked to math and spelling performance (parietal, occipital and temporal lobe areas)
- 44 boys, 23 girls eight to 16 years of age: 67 children with heavy prenatal alcohol exposure () and 61 children who were not prenatally exposed to alcohol (33 boys, 28 girls).

Leila Glass, Eileen M. Moore, Natacha Akshoomoff, Kenneth Lyons Jones, Edward P. Riley, Sarah N. Mattson. Academic Difficulties in Children with Prenatal Alcohol Exposure: Presence, Profile, and Neural Correlates. Alcoholism: Clinical and Experimental Research, 2017; DOI: <u>10.1111/acer.13366</u>

Adaptive Function Primary Disabilities (CNS mediated neurocognitive abilities and neuromaturation)

Secondary Disabilities (AKA: Day to Day Life)



DIATHESIS STRESS

- This model suggests individuals who are vulnerable (have PRIMARY DISABILITIES)
- And are exposed to stress, will develop problems (SECONDARY DISABILITIES).



• PROTECTIVE FACTORS

- Stable, nurturant, good quality home during critical / important parts of life (experience-dependent/expectant) or majority of life
- Relationship / Bonding
- Remain safe from violence
- Apply for and have appropriate aide (social services)
- Early identification

Secondary Disabilities

Rate increases drastically in adolescence

- School disruption (61%)
- Trouble with the Law (60%)
- Confinement experience (50%)
- Inappropriate sexual behaviors (49%)
- Drug/alcohol problems (35%)
- Mental Health diagnoses affecting adaptive function (94%)
 - At least one mental health episode
- Dysfunctional family interactions



Streissguth et al. Longitudinal study at the University of Washington

- In Sweden, a national register–based study of 79 adults with an FAS diagnosis, at a mean age of 32.
 - Compared to 3160 comparison individuals matched on age, gender, and place of birth.
 - Outcomes for men and women were similar
 - Rangmar, J., Hjern, A., Vinnerljung, B., Strömland, K., Aronson, M., & Fahlke, C. (2015). Psychosocial outcomes of fetal alcohol syndrome in adulthood. *Pediatrics*, 135(1), e52-e58.

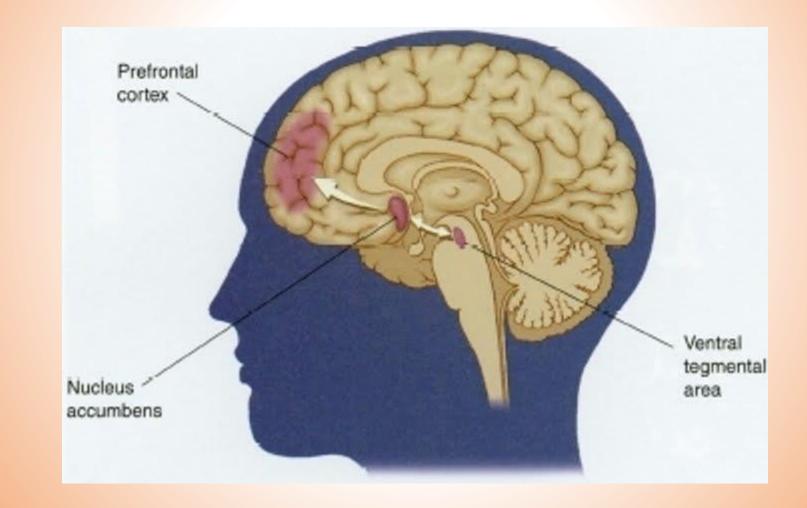
	FAS	Non-FAS				
State Care as children	81%	4%				
Special Education	25%	2%				
Unemployed	51%	15%				
Disability Pension	31%	3%				
Psychiatric Hospitalization:						
Alcohol Abuse	9%	2%				
Psychiatric Disorders	33%	5%				
Prescribed Psychotropic Drugs	57%	27%				
(Gender)	63% men	37% women				

Won't or Can't?

- At the interpersonal level, people often misinterpret the primary disabilities of individuals with FASD as willful behavior (e.g., "won't" vs. "can't").
 - May lead to inappropriate interventions or punishments
 - Result is that the individual with the FASD feels frustrated and discouraged
 - May use maladaptive behaviors to cope with the stress and frustration of not being understood or supported
- Importance of neuropsychology
 - Can frame and assist in understanding social, emotional and behavioral challenges from the information processing perspective

WHY?: Impact on the brain's reward circuitry

- After the prenatal brain is exposed to alcohol, the endocannibinoids have a different effect on certain dopamine neurons which are involved in addicted behaviors
- The dopamine neurons in the brain become more sensitive to a drug of abuse's effect. So, later in life, a person needs much less drug use to become addicted.
- But not just addiction. Decision making in general.
- Ventral Tegmentum: Reward evaluation. The value of the immediate reward may be overrepresented. Implications for a number of behaviors.
 - University at Buffalo. "Why does prenatal alcohol exposure increase the likelihood of addiction?." ScienceDaily. ScienceDaily, 7 July 2017. www.sciencedaily.com/releases/2017/07/170707211125.htm.
 - Kathryn Hausknecht, Ying-Ling Shen, Rui-Xiang Wang, Samir Haj-Dahmane, Roh-Yu Shen. Prenatal Ethanol Exposure Persistently Alters Endocannabinoid Signaling and Endocannabinoid-Mediated Excitatory Synaptic Plasticity in Ventral Tegmental Area Dopamine Neurons. The Journal of Neuroscience, 2017; 37 (24): 5798 DOI: 10.1523/JNEUROSCI.3894-16.2017





Alcohol and Pregnancy. No safe amount. No safe time. No safe alcohol. Period.

Interventions





No FASD specific interventions, though we have a foot in the door.

Traditional interaction approaches used by parents, teachers and other adults might not be effective for children with an FASD. Guiding principles include:

Concentrate on the child's strengths and talents

Accept the child's limitations/Positive Parenting/Differential reinforcement of positive behavior

Be consistent with everything (discipline, school, behaviors)

Use concrete language and examples

Use stable routines that do not change daily

Keep it simple

Be specific—say exactly what one intends (and follow through)

Structure your child's world to provide a foundation for daily living

Use positive reinforcement often (praise, incentives)

Supervise: friends, visits, and routines

Repeat, repeat, repeat

Summary of FASD Intervention Areas

- Social
- Family/School Communication Strategies: Positive Behavioral Support
- Math
- Executive Function
- Adaptive, Vocational and Life Skill

Bruin Buddies:

96 children 6 – 12 years completed the study. Vineland Social Composite z \leq 1.0; verbal IQ \geq 70

- Parent assisted children's friendship training
 - Target areas: understanding social cues, indiscriminant social behavior, communicating in social contexts
 - Skills taught: a) social network formation with the aid of b)informational interchange with peers leading to a common ground activity, c) entry into a group of children already in play, d) in home play dates, and conflict avoidance and negotiation.
 - Results was improvement at home as a result of treatment, though not in the classroom. More work was suggested.

O'Connor et al., (2006) J Consulting and Clinical Psychology

Georgia: Socioeconomic Habilitation using the Math Interactive Learning Experience

56 children 3 – 10 years of age. Consistent caregiver for 6 months before and after the study.

- Objective: to improve behavioral and mathematic functioning of alcohol affected children.
 - 6 weeks of math tutoring
 - Active learning approach using "plan-do-review" methodology
 - Caregivers received instruction in supporting math learning and weekly home assignments to compliment individualized tutoring sessions
 - Education to the teachers about alcohol related neurodevelopmental problems
 - Participants did make gains in math and behavior as assessed via pre- and post- testing with CBCL.

Behavior Consultation Intervention (WA)

52 children 5 – 11 years of age.

- Objective: To develop a program, Families Moving Forward, for families with children with FASDs
 - Positive behavior support techniques
 - (In WA state, 82% of FASD children had disruptive behaviors)
 - That the families tend to be highly stressed is not helpful.
 - Quality of caregiving a specific target
 - Parents of FASDs very diverse group of birth, kinship, foster and adoptive
- The program: 9 11 months, with at least 16 every-other weekend 90 min sessions. Program was designed with flexibility in mind.

Parent-Child Interaction Therapy

46 children 3 – 7 years of age.

- Objective: reduce behavior and parenting stress.
- PCIT provides a live coached practice of behavioral parenting skills
 - Enhance parent-child relationship; increase appropriate social skills; reduce inappropriate behaviors; promote positive discipline.
 - 14 weeks 90-min sessions
- Approx 50% attrition rate
- There was improvement, but no significant difference between PCIT and Parent Support and Management group.

Positive Behavior Support

- Emerged from a combination of applied behavioral analysis, the inclusion movement, and person-centered planning.
- The PBS approach primarily focuses on adapting the environment to enhance the individual's quality of life. The reduction of problem behaviors is a secondary goal.
 - 1) lifespan perspective,
 - 2) emphasis on prevention,
 - 3) a comprehensive lifestyle change and quality of life,
 - 5) systems change and multicomponent intervention,
 - 5) stakeholder participation (i.e., collaborative vs. expert-driven stance)
- Petrenko, Christie LM, Naira Tahir, Erin C. Mahoney, and Nancy P. Chin. "A qualitative assessment of program characteristics for preventing secondary conditions in individuals with fetal alcohol spectrum disorders." *Journal of population therapeutics and clinical pharmacology= Journal de la therapeutique des populations et de la pharamcologie clinique* 21, no. 2 (2014): e246.

 Recommended services for young adults with FASD focus on <u>life skills</u> <u>development and vocational services</u>. Support groups and counseling have also been found to promote positive adjustment and mental health. In particular, the benefits of <u>group services and activities</u> <u>include promoting social skills</u>, positive self-esteem, and a sense of <u>belonging</u>. Knowing the diagnosis is cited as important in providing interventions. A case manager specializing in or having a depth of knowledge about FASD has been found to be helpful.



Petrenko et al. 2014

Neurocognitive habilitation for children with FASDs (Children's Research Triangle)

78 children, 40 received the intervention. Specific ages not given.

- Recipients: Children with FASDs who had been adopted or who were in foster care.
 - These children do not have the protective factors of
 - Being raised in a stable nurturing home
 - Diagnosis before age 6; receiving early intervention
 - No sexual or physical abuse history
 - Not changing households every few years
- Core components: education and support to family
 - Also executive function
 - Alert program (car engine as a metaphor): 12 weekly 75-min sessions
 - Tools for memory, cause and effect reasoning, sequencing, planning, problem solving
- Results were promising.

How does your engine run? (Executive Function)

- The Alert Program for Self-Regulation
 - Beneficial neuroanatomical changes following treatment for their core deficit in executive functioning.
 - Increases in cortical gray matter in children with FASD, in regions involved with
 - response inhibition,
 - outcome monitoring,
 - and emotion regulation



Soh, D. W., Skocic, J., Nash, K., Stevens, S., Turner, G. R., & Rovet, J. (2015). Self-regulation therapy increases frontal gray matter in children with fetal alcohol spectrum disorder: evaluation by voxel-based morphometry. *Frontiers in Human Neuroscience*, *9*, 108. http://doi.org/10.3389/fnhum.2015.00108

Higher Level Organization

Children with FASD demonstrate a capacity to learn new skills, though do so through explicit instruction rather than through observation and abstracting rules, skills and ongoing knowledge as we typically expect.

- Organizational strategies across the board, especially as related to skills that mature. These are higher level executive functions. Development of these cant be taken for granted.
- Executive Functions can make it or break it!!
 - Contribute to decision making and problem solving.

Alcohol and Pregnancy. No safe amount. No safe time. No safe alcohol. Period.

Medication for ADHD symptoms in FASDs?

- Although it has been estimated that 50 90% of individuals with FASD have ADHD,
- Stimulants have had differential outcomes.
 - May reduce activity, but not improve attention
 - Results may be unpredictable, and even lead to poorer outcomes.
 - May increase spontaneous motor behavior later in life.
 - Clinical trial for Strattera underway.
 - In general, not a great response to methylphenidate



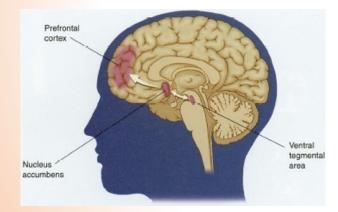
Animal Assisted Therapy (AAT)

One commonly documented strength among individuals with FASDs is that they tend to work well with animals. AAT is delivered by a professional service provider, such as a recreation therapist, and is designed to promote improvements in physical, emotional, social, or cognitive functioning. Animals used in AAT can include cats, rabbits, birds, fish, and perhaps most commonly, horses and dogs.

From AAP internet site: <u>https://www.aap.org/en-us/advocacy-and-policy/aap-health-initiatives/fetal-alcohol-spectrum-disorders-toolkit/Pages/Management.aspx</u>

Enhancing knowledge Pre-Pregnancy

- Transgenerational effects
- Prompts the question: Is FAS heritable?
- (for example, impact on reward circuitry)





 Charles W. Abbott, David J. Rohac, Riley T. Bottom, Sahil Patadia, Kelly J. Huffman. Prenatal Ethanol Exposure and Neocortical Development: A Transgenerational Model of FASD. Cerebral Cortex, 2017; 1 DOI: <u>10.1093/cercor/bhx168</u>

Enhancing knowledge Pre-Pregnancy

- Pregnant rats received the equivalent of one glass of wine, four days in a row, at gestational days 17-20, the equivalent of the second trimester in humans. Juvenile male and female offspring were then tested for water or alcohol consumption. Adolescent males were tested for sensitivity to alcohol by injecting them with a high-alcohol dose, which made them unresponsive (drunk on their back), and measuring the time it took them to recover their senses (back on their four paws). The results suggest that if a mother drinks during pregnancy, even just a little bit, she increases the risk that her progeny will become alcoholic.
- "Our findings show that in the rat, when a mother consumes the equivalent of one glass
 of wine four times during the pregnancy, her offspring and grand-offspring, up to the
 third generation, show increased alcohol preference and less sensitivity to alcohol," said
 Cameron. "Thus, the offspring are more likely to develop alcoholism.
- Michael E. Nizhnikov, Daniel O. Popoola, Nicole M. Cameron. Transgenerational Transmission of the Effect of Gestational Ethanol Exposure on Ethanol Use-Related Behavior. Alcoholism: Clinical and Experimental Research, 2016; DOI: 10.1111/acer.12978

The Future

- Neuroplastic Interventions: white matter development could be more resistant to the long-lasting effects of prenatal alcohol exposure and/or be more amenable to plasticity relating to environmental factors
 - Exercise
 - Shown to impact brain structure involved in learning and memory
 - Has been used as an intervention for ADHD, anxiety, PTSD, depression and other conditions
 - Mindfulness/Meditation
 - attention
 - Impulse Control
 - Awareness of Emotions
 - Social Compassion
 - ALERT program
 - Attention
 - Impulse Control
 - Awareness of Emotions
 - Social Compassion



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Web Resources (Handout)

- FAS Community Resource Center: <u>http://www.come-over.to/FASCRC/</u>
- National Organization on Fetal Alcohol Syndrome: <u>http://www.nofas.org/</u>
- American Academy of Pediatrics: https://www.aap.org/en-us/advocacy-and-policy/aap-health-initiatives/fetal-alcohol-spectrum-disorders-toolkit/Pages/Management.aspx



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