Round 26: Autism

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Autism

- Spectrum of disorders that are characterized by:
 - Impairment in social interaction
 - Communication
 - Restricted repetitive and stereotyped patterns of behavior, interests, and activities

- Some Autistic children have difficulty learning, have signs of lower than normal intelligence
- Other children have normal to high intelligence, learn quickly, especially in specific interest categories
 - Yet, have trouble communicating and applying what they know in everyday life
 - Adjusting to social situations

What are the signs of Autism?



Autism Characteristics

Social Communication/Interaction

- •Fails to respond to his or her name
- •Resists cuddling and holding, and seems to prefer playing alone, retreating into his or her own world
- •Has poor eye contact and lacks facial expression
- •Doesn't speak or has delayed speech, or loses previous ability to say words or sentences
- •Can't start a conversation or keep one going, or only starts one to make requests or label items
- •Speaks with an abnormal tone or rhythm and may use a singsong voice or robot-like speech
- •Repeats words or phrases verbatim, but doesn't understand how to use them
- •Doesn't appear to understand simple questions or directions
- •Doesn't express emotions or feelings and appears unaware of others' feelings
- Doesn't point at or bring objects to share interest
- •Inappropriately approaches a social interaction by being passive, aggressive or disruptive
- •Has difficulty recognizing nonverbal cues, such as interpreting other people's facial expressions, body postures or tone of voice

Autism Characteristics

- Patterns of behavior
 - Limited, repetitive patterns of behavior, interests or activities, including:
 - •Performs repetitive movements, such as rocking, spinning or hand flapping
 - •Performs activities that could cause self-harm, such as biting or head-banging
 - •Develops specific routines or rituals and becomes disturbed at the slightest change
 - •Has problems with coordination or has odd movement patterns, such as clumsiness or walking on toes, and has odd, stiff or exaggerated body language
 - •Is fascinated by details of an object, such as the spinning wheels of a toy car, but doesn't understand the overall purpose or function of the object
 - •Is unusually sensitive to light, sound or touch, yet may be indifferent to pain or temperature
 - Doesn't engage in imitative or make-believe play
 - •Fixates on an object or activity with abnormal intensity or focus
 - •Has specific food preferences, such as eating only a few foods, or refusing foods with a certain texture

Autism Prevalence & Risk Factors

- 1 in 54 children will be diagnosed with Autism Spectrum Disorder
 - 1 in 34 boys
 - 1 in 144 girls
- Can be diagnosed as early as 2 years old, usually by 4
- Likely the combination of Genetic and Environmental factors
 - 80% Genetic possible link to chromosome 16
 - 20% Environmental
 - 1% Maternal factors
 - Extreme pre-term babies born before 26 weeks of gestation
 - Advanced Parental age
 - Exposure to pesticides, toxins, air pollution
 - Previous child with autism, 2-18% chance of having a second child with Autism

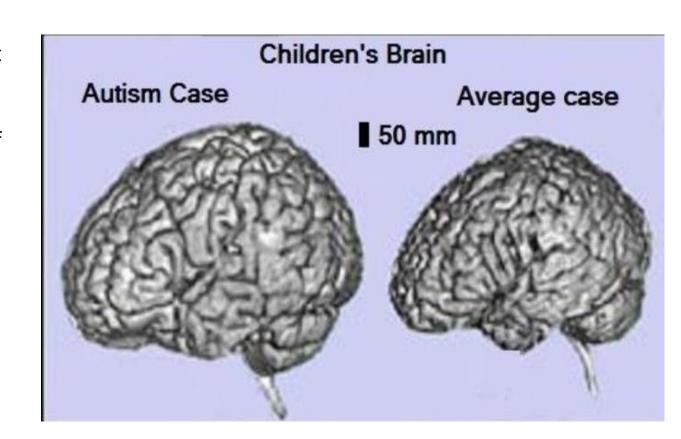
Estimated Autism Prevalence 2020



Autism Behavioral Development

- Often children show symptoms of autism within the first year.
 - Some children show signs of autism spectrum disorder in early infancy
 - Reduced eye contact
 - Lack of response to their name or indifference to caregivers.
- Some children appear to develop normally in the first year, and then go through a period of regression between 18 and 24 months
 - Suddenly become withdrawn or aggressive
 - Lose language skills they've already acquired
 - Signs usually are seen by age 2 years

- Most common finding is accelerated brain development early in childhood (2-4 y/o)
 - Significantly faster expansion of the surface area of their cortex from 6 to 12 months of age.
 - In year 2, brain volume increases much faster in autistic children than in their non-autistic peers
 - Amygdala is enlarged in children up to 5 years
 - Extent of enlargement correlates to symptom severity
 - Adolescents & adults amygdala is smaller



- One theory suggests autism is linked to too many synapses insufficient synaptic pruning
 - Insufficient pruning may be linked to excess of protein mTOR, which disrupts autophagy
 - Medication Rapamycin inhibit mTOR & restores normal synaptic pruning and appears to alleviate symptoms in mouse-model of autism



- Possible phenotypes
 - Deficits is Social Language Processing & Social Attention
 - Abnormalities in inferior frontal gyrus (IFG, Broca's area), superior temporal sulcus (STS), and Wernicke's area
 - Impairments of Social Behaviors
 - Abnormalities in the frontal lobe, superior temporal cortex, parietal cortex, and amygdala
 - Restrictive, repetitive behaviors of ASD
 - Abnormalities in the orbitofrontal cortex and caudate nucleus
 - Also been found in OCD, GAD, and Schizophrenia

- Poor Face/Emotion Processing
 - Hypo-activation in Bilateral, and specifically left, fusiform face area and occipital face area
- Poor Verbal Processing/Communication
 - More symmetrical brain
 - Reduced leftward language lateralization
 - Higher rate of being left-handed
 - Hypoactivation in
 - Left inferior, middle frontal gyrus, and left angular gyrus
 - Anterior Cingulate Cortex (ACC)
 - Also related to poor response monitoring
- Dysfunction in
 - Sensory processing -> Somatosensory areas
 - Motor control (i.e., Restrictive, repetitive Behaviors) -> Motor cortex, Supplementary motor area
 - Inhibitory control & Cognitive Flexibility -> Frontal regions

Connectivity Differences

- Poor connectivity between hemispheres reduced volume of corpus collosum
- Long-range under-connectivity
 - Difficulty in tasks that require integration of information from different parts of the brain (e.g., social function and complex motor tasks)
 - Problem is that visual information can be identified but not put into context emotionally appraised
- Short-range over-connectivity
 - Excellent performance in very specific tasks that utilize discrete, local brain regions (e.g., paying attention to specific details or narrow, surface level concepts)
- Different patterns of sulci & gyri
 - More folding in left parietal and temporal lobes and in right frontal and temporal lobes
 - Reduced gyrification
 - Children -> Right inferior frontal and medial parieto-occipital cortices
 - 8-40 years -> Left supramarginal gyrus

Mu Rhythms

- Alpha frequency (9-14 Hz)
- Observed over C3 & C4
- Is suppressed when actively executing an action, passively thinking about a movement, or watching someone else produce a movement
- Believed to represent mirror neuron activity -> understanding of others' actions/ Theory of Mind
- Experimentally inhibit Mu Rhythm Suppression with TMS
 - Participants perform worse on emotion recognition task

- 15 year old male
- Previously diagnosed with ADHD & Dyslexia
- Parents brought the boy in for possible Dyslexia.

Symptoms

- Reading, Spelling
- Comprehension, Word Recognition
- Attention span
- Basic life skills
- Simple tasks, basic motor skills, using utensils
- Lying
- Sensory sensitivity to textures
- Introverted & Socially Awkward

Pre/Perinatal

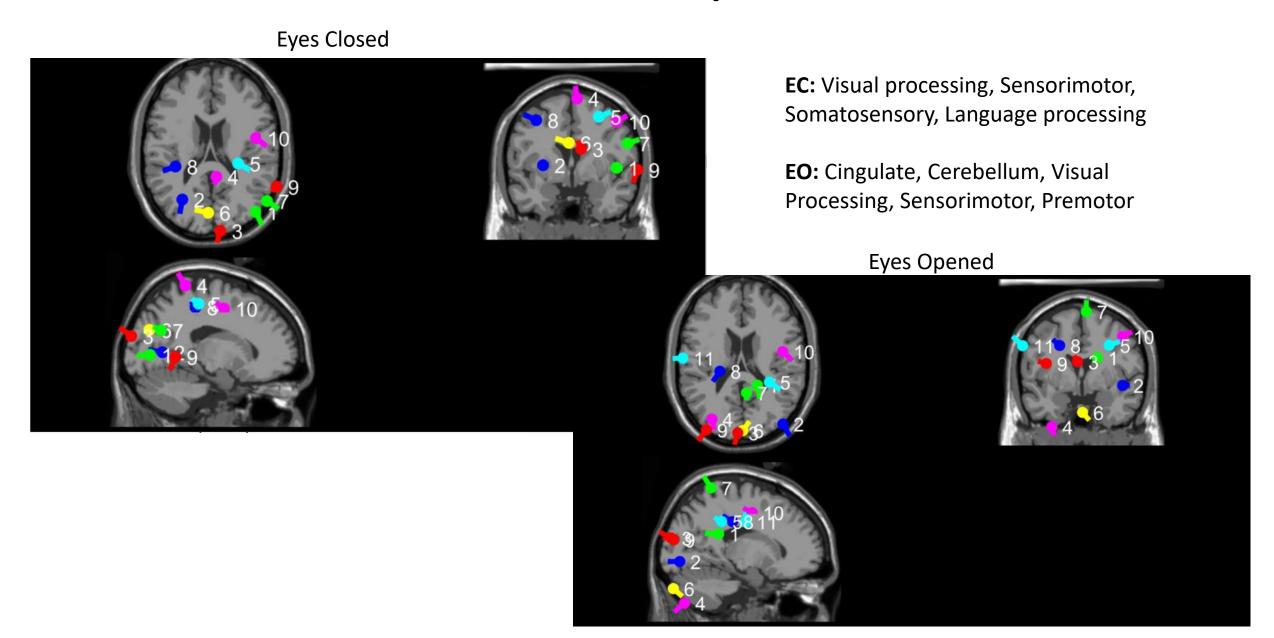
- Mother took prescription drugs
- Mother experienced preeclampsia, high blood pressure, bedrest, emergency c-section
- Pt was jaundice, had an infection, spent 3 days in NICU

Developmental History

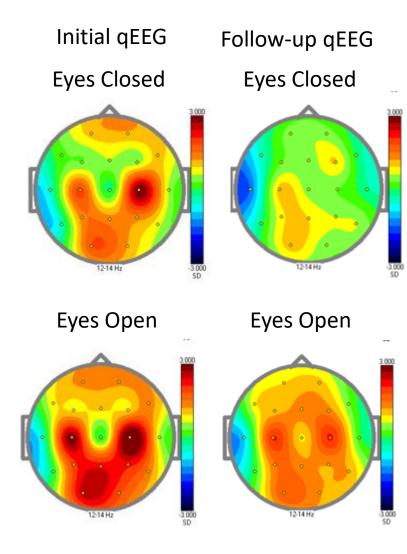
- Delayed toilet training, completed by 7 years
- Problems with fine motor movement, sensory issues, narrowed interest
- Fascinated with cars/wheels, lines up cars and doesn't let anyone touch them
- Knows a lot regarding cars and Nascar drivers

- **Neuropsychological testing** indicated the patient fell significantly below expectations on the follow:
 - Attention/concentration for attention to detail with a speeded response
 - Graphomotor speed/learning
 - Sustained auditory attention
 - Visual sustained attention ratings were invalid
 - Learning and memory for visual memory/learning
 - Executive functions for mental switching and visual problem solving
 - Verbal reasoning for expressive language and language memory
 - Perceptual reasoning for facial recognition and social processing
- Scores on achievement tests fell below expectation for:
 - Reading
 - Writing
 - Mathematics
- Relative differences were also noted for:
 - Working memory
 - Verbal and perceptual intellectual reasoning
 - Visuo-motor integration
 - Visual, rote memorization

*IBH Diagnosis => Autism, specific learning delays in math, reading & writing



Inhibit	2-7
Inhibit	8-14
Inhibit	16-28
Reward	6-15
A1	NZ



Change in Connectivity (Eyes Closed)

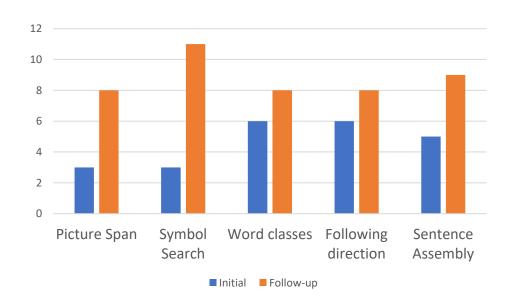
Post

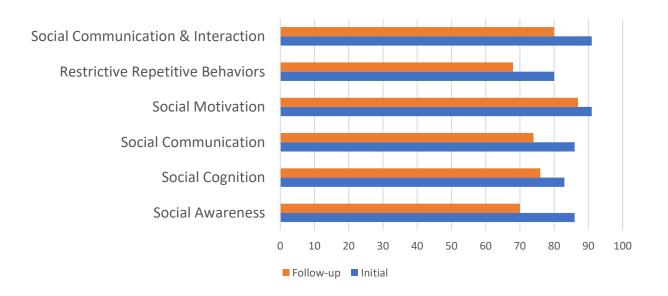
Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter
254-305s	8-13Hz	0.015652341	49.22628449	0.02628525	62.73602	99.0001457

Previous

Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter		
59-117s	4-14Hz	0.016184928	58.78434724	0.023319799	67.70655	129.899902		

24%





- Picture span -> working memory
- Symbol search -> processing speed domain
- Word class, following directions, sentence assembly -> Language

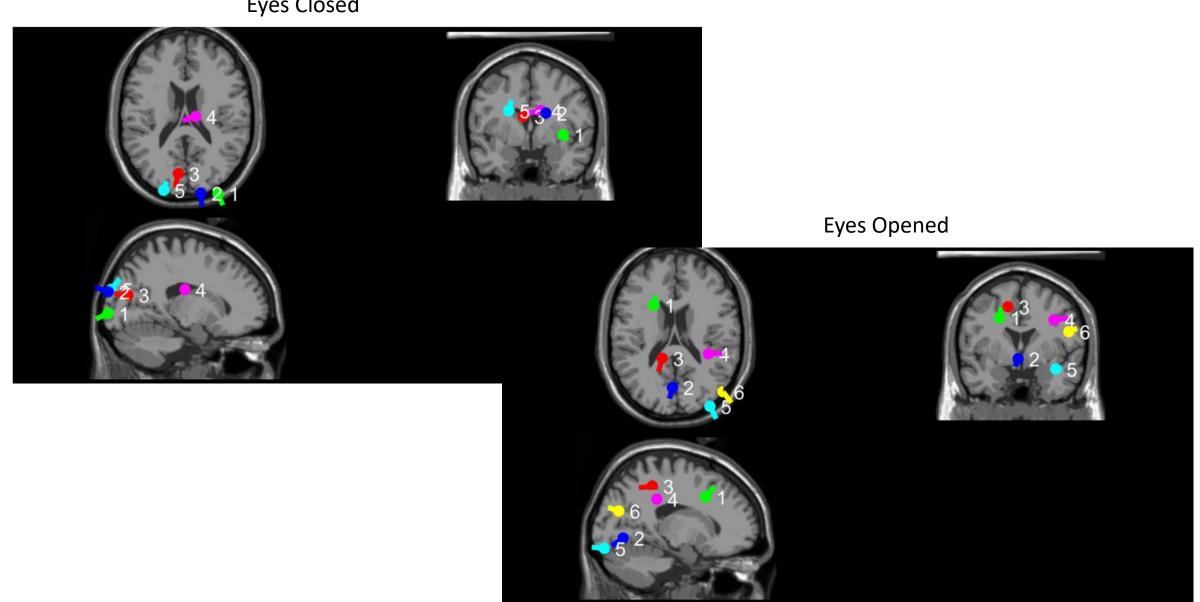
- 6 year old male
- Previously diagnosed with autism and sensory processing disorder
- Evaluation for attention deficits
 - High energy level
 - Short attention span
 - Inability to sit still/Fidgeting/Waiting his turn
 - Talkative/interrupting
 - Frequently loses items
 - Other Symptoms
 - Poor emotion regulation
 - Poor fine motor skills (e.g., opening snack, tying shoes, pants with zippers/buttons)
 - Delayed developmental milestones
 - Turning over, crawling, walking, potty training
 - Sensory issues
 - Textures (foods, clothes)
 - Loud sounds
 - Struggles with social cues

- Neuropsychological testing -> significantly below expectations:
 - Sensory-motor functions for visual motor precision
 - Attention/concentration for visual working memory
 - Verbal reasoning for understanding spoken paragraphs
 - Social perception for neutral facial expressions
- Achievement test scores for oral reading and reading fluency fell greater than two years behind
- Relative differences were also noted for:
 - Finger tapping sequences and imitating hand positions
 - Recalling sentences
 - Graphomotor speed/learning
 - Inhibition
 - Written expression

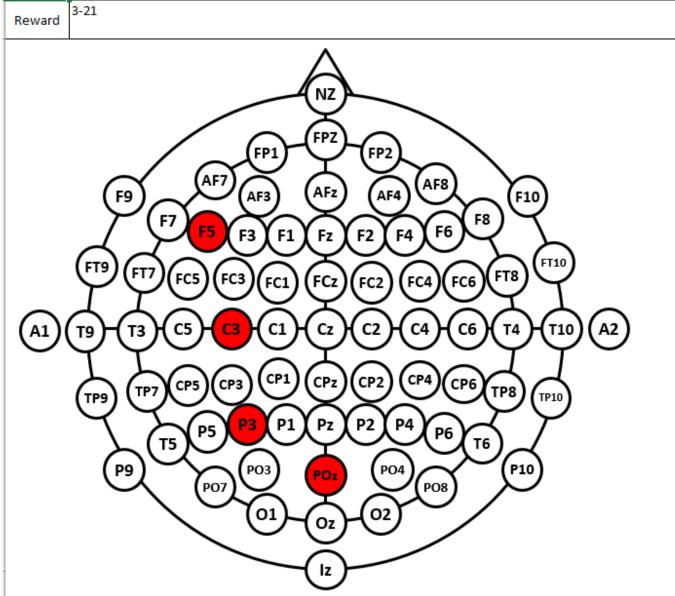
IBH Diagnoses -> Autism, Specific Learning delay reading,
Generalized Anxiety Disorder

- The parent report & clinical evaluations consistent with the following problem areas:
 - Self-care, behavioral regulation, social skills, emotional regulation
 - Autism

Eyes Closed



Inhib	it 4-6
Inhib	it 10-12
Inhib	it 20-30
Rewa	rd ³⁻²¹



Parent-report

- Improvements in black and white thinking, emotion regulation, & phonics/reading/writing
- Better attention span
- Fewer meltdowns

Neuropsychological Tests

- Improvements in visuomotor precision
- Decreases in aggression, conduct problems, & Depression

Post

Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter
27-55s	4-21Hz	0.022023777	48.44819485	0.027721917	57.97599	100.216607

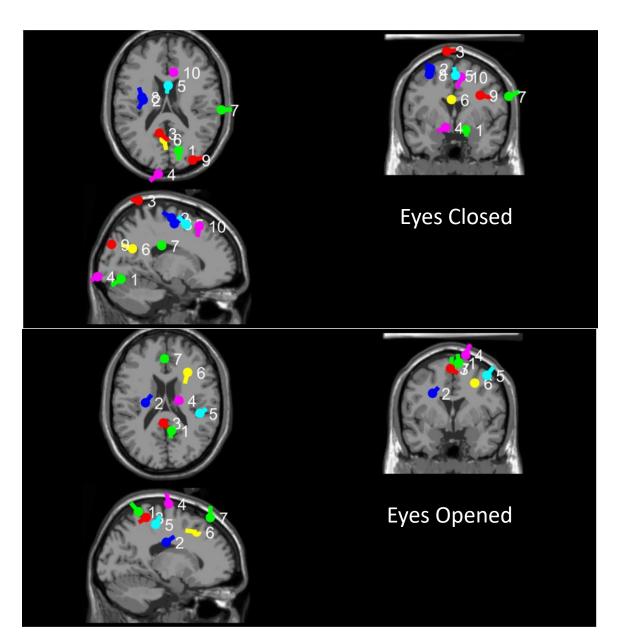
Previous

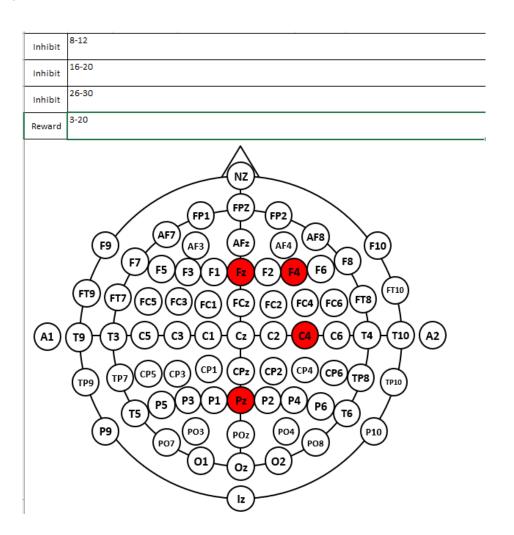
Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter
63-78s	3-21Hz	0.016994242	54.58975181	0.025637671	59.36918	133.644323

- 11 year old male
- Evaluate for ADHD
- Symptoms
 - Struggles with spelling, writing but excels in math
 - Attention span/listening
 - Recall
 - Excessive talking, distractibility
 - High need for organization (i.e., Closet, color coding schoolwork)
 - Exhibits compulsions or "must do" behaviors related to cleanliness
- Developmental History
 - Mother had gestational diabetes
 - Required oxygen for 45 mins after delivery

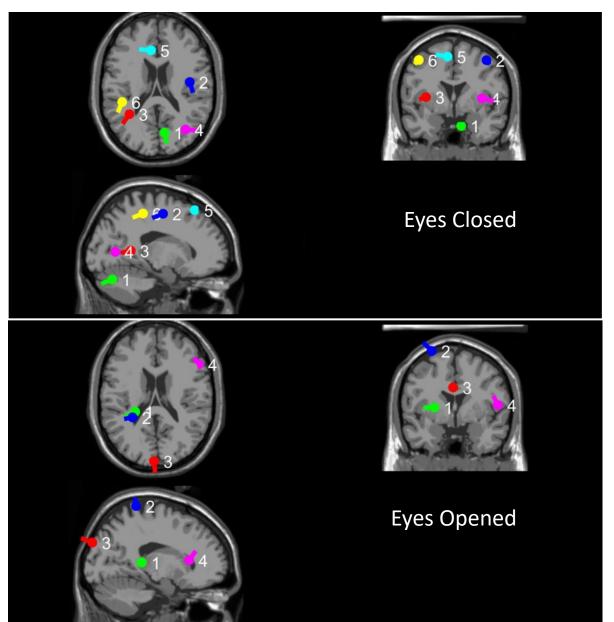
- Neuropsychological assessment
 - Significantly below expectations
 - Sensory-motor functions for imitating hand positions and manual motor sequencing
 - Perceptual Processing/Reasoning for visuomotor integration
 - Oromotor sequencing
 - Attention/concentration for sustained focus
 - Verbal Reasoning/Processing for abstract reasoning
 - Phonological processing
 - Quantitative analysis and social processing
 - Achievement tests fell below expectation for
 - Reading, Written Language and Mathematics, falling at least two years delayed.
 - Word Attack, Oral Reading, Passage Comprehension, Math Fluency, and Spelling each fell at least 3years below expectations.
 - Scales endorsed by the parents included those evaluating autism symptoms, externalizing problems, executive dysfunction and conduct issues.

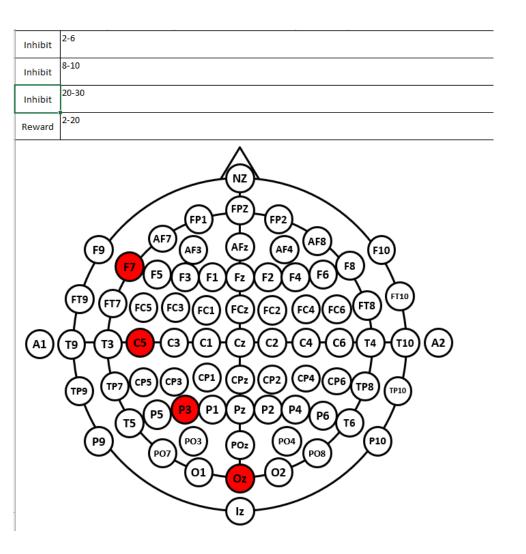
*IBH Diagnosis => Pervasive Developmental Disorder, ADHD, Specific learning delays in math, reading & writing





Neuropsychological tests showed improvements on a number of attention measures, but selfreports indicated no-change





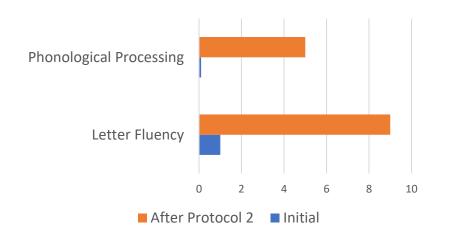
- Self-report indicated:
 - Fluency -> slightly improved
 - Ability to read out loud -> much improved
 - Attention/Focus -> much improved
 - Teachers indicate much improved behavior

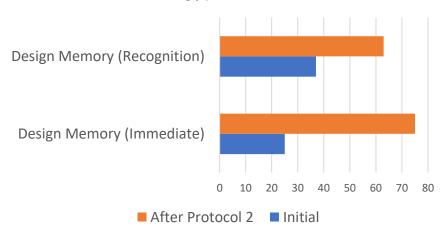
Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter
63-94s	3-20Hz	0.009927007	89.39164156	0.018501664	118.6873	200.550825

Previous

Time Range	Freq Range	Clust Coeff	Path Length	Global Eff	Radius	Diameter
47-94s	2-20Hz	0.007062285	150.9438165	0.010735914	130.8422	371.774539

46%





Thank You