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Disclosures / References

 Dr. Coplan is author of Making Sense of Autistic Spectrum Disorders: Create the brightest future for your child with the best treatment options (Bantam-Dell, 2010), and receives royalties on its sale



Chapter 12: Behavior management and psychopharmacology

This presentation will include a discussion of off-label drug use

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Outline / Basic Premises - 1

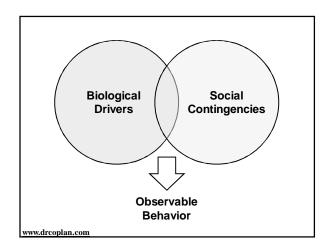
- Biologically driven behaviors / traits
 - Cognitive Rigidity
 - Dysregulation of Attention
 - Dysregulation of Arousal
 - Dysregulation of Sleep
 - Dysregulation of Sensory Processing
- Occur irrespective of environmental contingencies
- Do not serve a social function
- Specific behaviors / traits are tied to specific neurotransmitters / brain systems

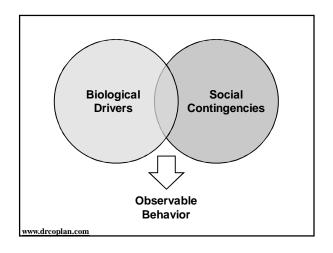
Outline / Basic Premises - 2

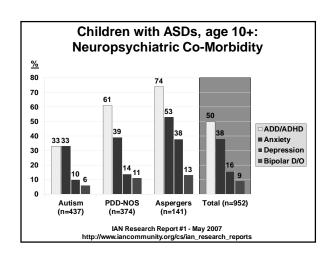
- · Socially driven behaviors
 - Occur in response to environmental contingencies
 - Serve a social function
 - Attention
 - · Access to desired objects or activities
 - · Escape from undesired activities
 - A-B-C Model
 - What is the Antecedent to the behavior?
 - · What is the Behavior itself?
 - What are the Consequences for the behavior?

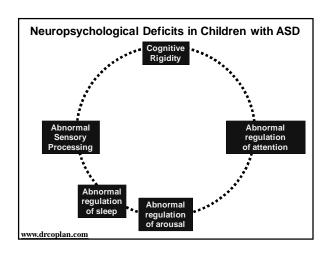
Outline / Basic Premises - 3

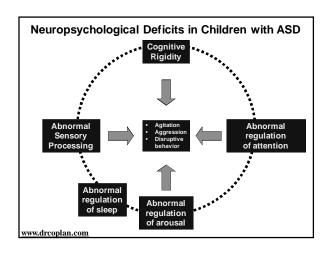
- Behavior analysis needs to take biological and environmental factors into account:
 - Underlying biological traits often provide the child with lots of opportunities to make unfortunate discoveries (viz: Tantrums or SIB are great ways to get attention or escape from tasks)
- Intervention often requires both pharmacologic and behavioral measures

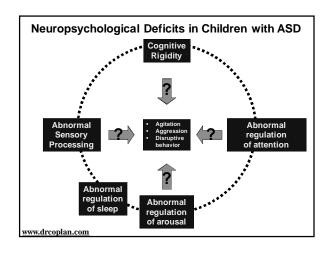


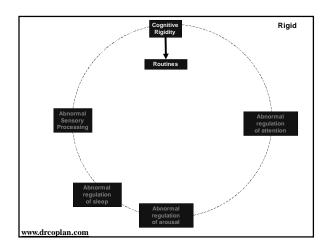


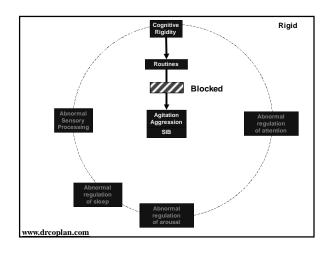


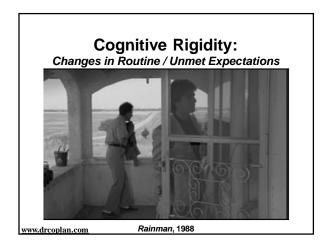




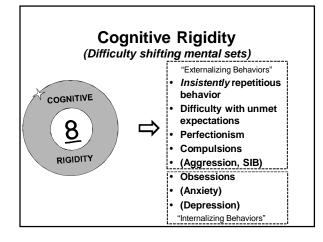


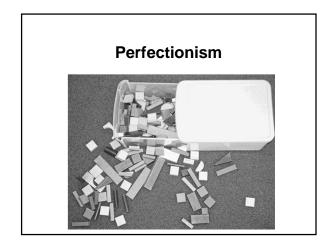


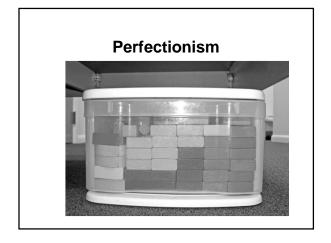


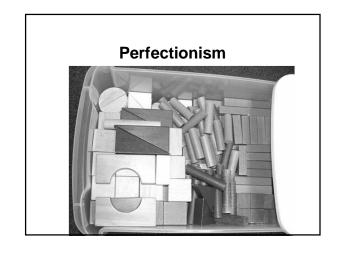


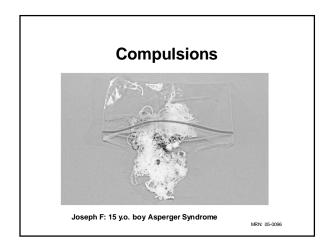


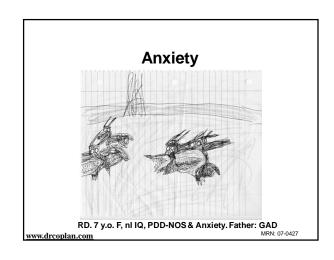


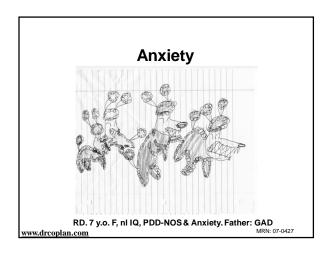


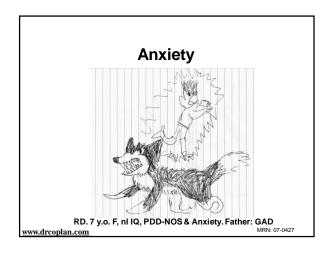


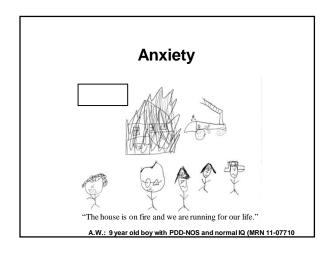


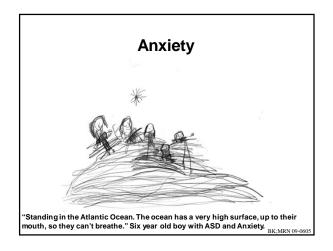


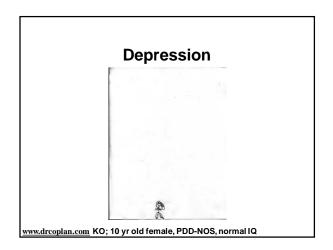


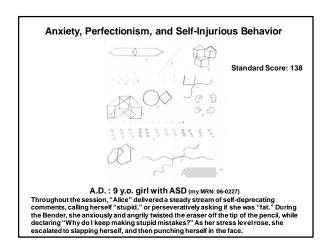


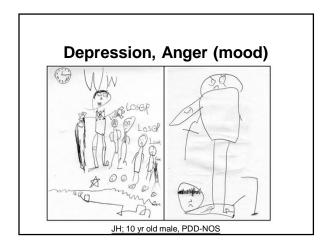


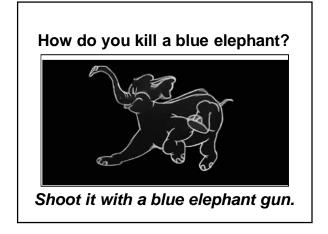


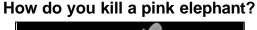










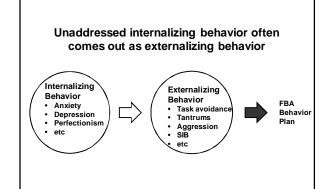


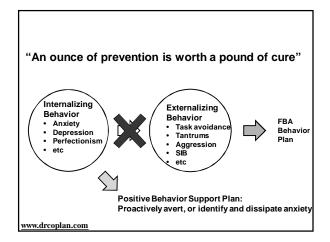


How do you kill a pink elephant?



Hold it by the trunk until it turns blue, then shoot it with a blue elephant gun.





The Story of Billy's Box - 1 (or, why it's important to ID internalizing behavior)

 8 y.o. boy with ASD and normal Nonverbal IQ

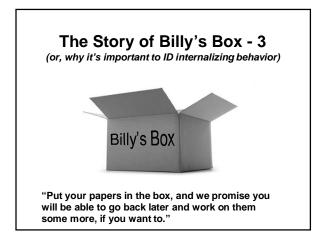
- · Severe tantrums at school
- Antecedents:
 - TRANSITIONS
- Function?
 - Not attention, escape, access
 - "Biological" (i.e. "just part of his ASD")?

The Story of Billy's Box - 2

(or, why it's important to ID internalizing behavior)

Q: "Billy – You're always getting in trouble at school. What's going on?"

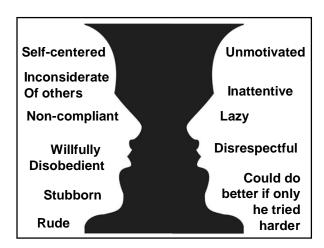
A: "I'm afraid that if I hand in my work, I'll never get a chance to go back and make it perfect."

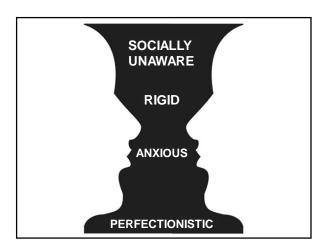


Positive Behavior Support Plan for Internalizing Behavior

- Staff Awareness
- Visual Schedules
 - What am I supposed to be doing do now?
 - What am I supposed to do next?
- Relaxation Techniques
 - Mental Imagery
 - Isometrics
 - Deep Breathing
 - "Break" cards
- Cognitive Behavioral Therapy (CBT)
- SSRIs

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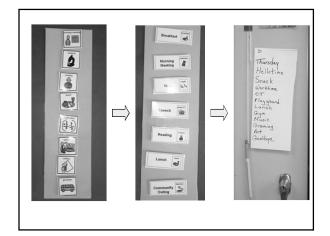




"We caution against the use of the word "stubborn" to characterize Ryan's classroom behavior. Ryan's task avoidance and non-adherence to teacher instruction reflect *cognitive rigidity and anxiety*, *rather than "stubborn" behavior*. Re-framing his actions will lead to more appropriate intervention, placing the focus on *anxiety management and cognitive flexibility*, rather than "compliance."

Positive Behavior Support Plan for Internalizing Behavior

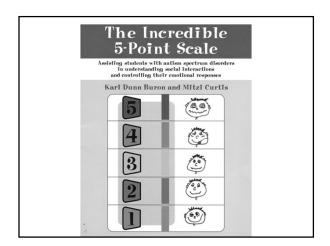
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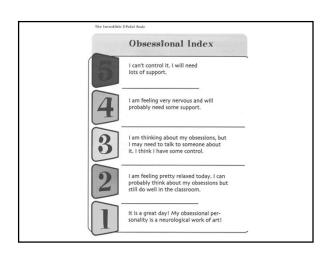


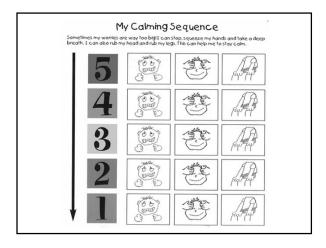
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Positive Behavior Support Plan for Internalizing Behavior

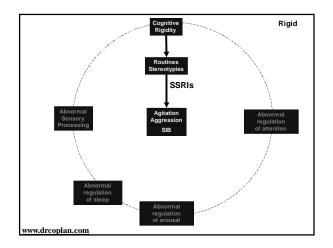
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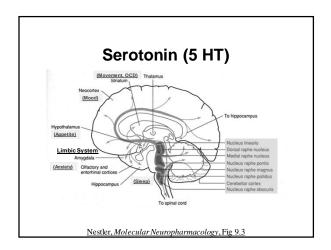
Selective Serotonin Reuptake Inhibitors (SSRIs)

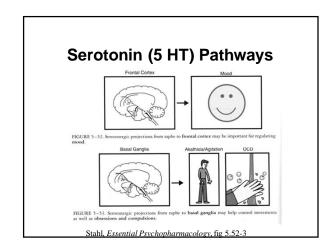
- Primary targets
 - Cognitive Rigidity
 - Anxiety
 - Obsessions (thoughts)
 - Compulsions (behavior)
 - Perfectionism
 - Depression
 - Stereotypies: Probably not
- "Downstream" benefit:

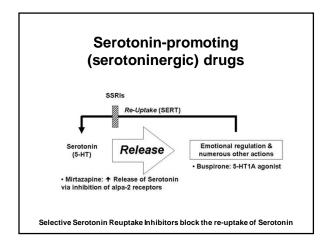
 - A Quality of Life

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SSRIs in ASDs • Side Effects - Activation • Hyperactivity • Irritability • Insomnia • Agitation - Uncommon or irrelevant • GI dysfunciton • Sexual dysfunction • "Black Box" warning (suicidal mentation)

Selective Serotonin Reuptake Inhibitors (SSRIs)

Generic Name	Brand Name		Comment	
Fluoxetine	Prozac	•	The first selective SRI	
Fluvoxamine	Luvox			
Sertraline	Zoloft	•	May be less activating	
Citalopram	Celexa	•	Prolonged QT interval	
Escitalopram	Lexapro		Prolonged QT interval	
And others				

Selective serotonin reuptake inhibitors (SSRIs) for autism spectrum disorder (ASD).

Williams, K., et al., Cochrane Database Syst Rev, 2010. 8: p. CD004677

Authors' conclusion:

"There is no evidence that SSRIs are effective as a treatment for children with autism. In fact, there is emerging evidence that they are not effective and can cause harm. As such SSRIs cannot be recommended as a treatment for children with autism at this time.'

Selective serotonin reuptake inhibitors (SSRIs) for autism spectrum disorder (ASD).

Williams, K., et al., Cochrane Database Syst Rev, 2010. 8: p. CD004677

- · Studies reviewed: 7 randomized controlled trials / 271 participants
 - Fluoxetine (2), fluvoxamine (2), fenfluramine (2), citalopram (1)
 Subjects: Children (5); Adults (2)

 - Varying inclusion criteria for Dx of ASD and IQ
 - 17 different outcome measures

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"Data were unsuitable for meta-analysis"

Selective serotonin reuptake inhibitors (SSRIs) for autism spectrum disorder (ASD).

Williams, K., et al., Cochrane Database Syst Rev, 2010. 8: p. CD004677

- Treatment-emergent symptoms
 - Citalopram: 1 child with new onset seizures (continued to have seizures after citalopram was stopped)
 - Fenfluramine: ↑ stereotypies; withdrawal, sadness; ♥appetite
 - "With monitoring, dose adjustment and time, all but one of these adverse effects were resolved"
 - Fluoxetine (Hollander 2005): 6 of 37 children had their dosage reduced due to agitation

 • 2 children in the placebo group also had their "dosage" reduced.
 - Difference between groups: Not significant

 Reviewers disregard the fact that by the end of the trial, "anxiety
 - and nervousness" was lower in the fluoxetine group compared to placebo: 15.9% vs. 33%
 - Fluvoxamine: No significant difference in side effects between SSRI and placebo

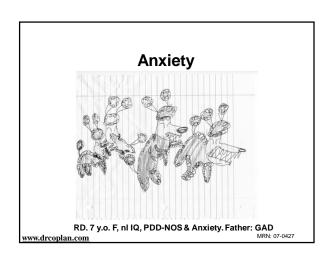
Pharmacotherapy for anxiety disorders in children and adolescents

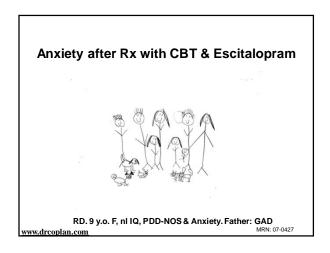
lpser JC, Stein DJ, Hawkridge S, Hoppe L. Cochrane Database of Systematic Reviews 2009, Issue 3.

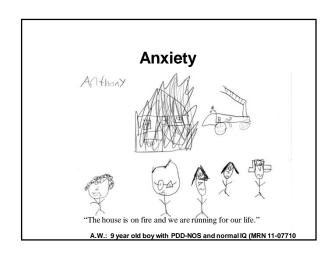
- Studies reviewed: 22 RCTs/ 2,519 participants
 - Short-term (average 11 wks)Mean age 12 yrs

 - o Drugs studied (versus placebo)
 - o SSRIs :15 (fluoxetine 6, fluvoxamine 2, paroxetine 3, sertraline 4)
 SNRIs: 5, (clomipramine 3), venlafaxine 2)
 Benzodiazepines: 2: (alprazolam 1, clonazepam 1)
 Tricyclic antidepressants: 1 (desipramine)
- Meta-analysis
 - o Response rate: Medication 59%; Placebo 31%

 - 7.3% of subjects treated with SSRIs withdrew bec/o side effects
 "The overwhelming majority of evidence of efficacy was for the SSRIs, with the most evidence in paediatric OCD"











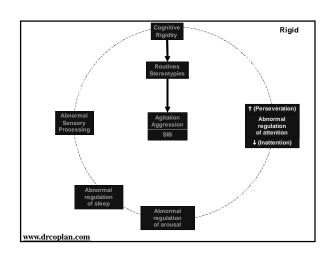
After one week on Sertraline

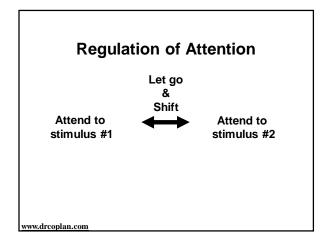
Sent: Thursday, May 31, 2012 To: James Coplan Subject: amazing shift in A.D. Importance: High

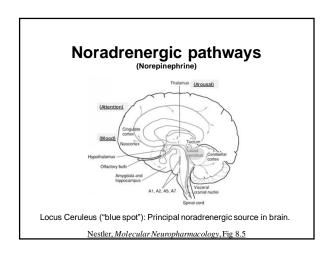
Dr. Coplan,

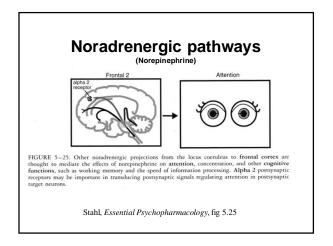
DI. Copfain,
I "know" that it takes several weeks for SSRI's to "kick in" but the child
I saw in my office today is simply a different child and the
improvements are being noted across settings by mutiple adults. There
was NO self abuse, NO negative self statements, an availability for
interventions, just a complete transformation. We "fixed" mistakes, "redid" errors, told jokes, and played together. The "core" Autistic symptoms are obviously still there - perseveration on bras, drawing, etc - but mood-wise there is no question that A. is already benefitting from the Sertraline... Impossible perhaps but really visibly clear...
Thank you very much.
S.S. Ph.D.

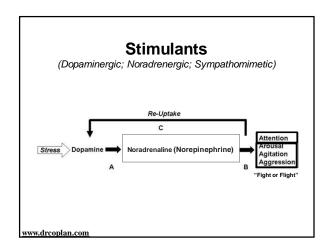






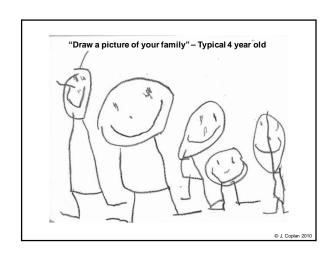


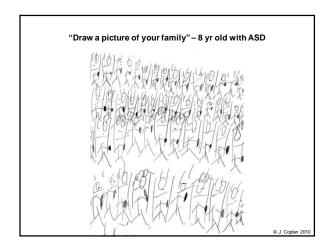


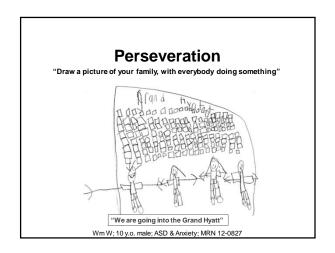


Abnormal Regulation of Attention - 1

- Perseveration
 - Inability to "Let go and shift"
 - Gets "stuck"
 - "Overattention Deficit Disorder"
- Compounds the effects of cognitive rigidity



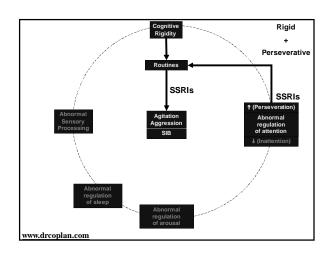




Abnormal Regulation of Attention (Perseveration)

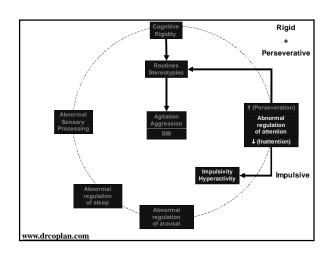
- Interventions
 - Verbal preparation for transitions
 - Visual Schedules
 - SSRIs (OCD: Proven; ASD: likely)

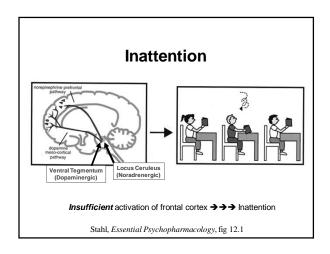
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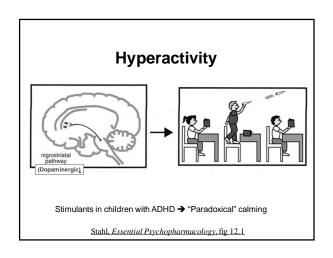


Abnormal Regulation of Attention - 2

- Inattention
 - Inability to focus
 - Impulsive
 - Distractible







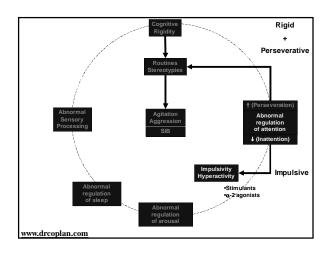
Inattention

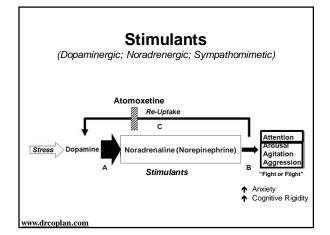
Classroom interventions

- Limited stimuli
- Short work periods
- Preferential seating

Medication

- Stimulants & Norepinephrine Reuptake Inhibitors (NRIs)
 - May ↑ anxiety / rigidity / agitation / Tics
- Alpha-2 agonists
 - Sleepiness
 - · Occasional weepiness
 - . Hypotension: Rare (start low, & go slow)





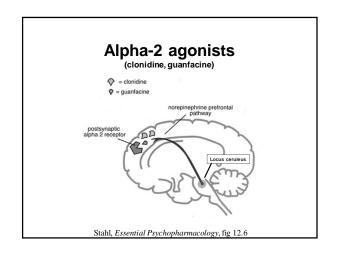
Stim	nulants,	NRI's
Generic Name(s)	Brand Name(s)	Comment
Amphetamine		FDA Schedule II
Dextroamphetamine	Dexedrine, Dextrostat	FDA Schedule II
Dextroamphetamine + amphetamine	Adderall	FDA Schedule II
Methylphenidate	Concerta, Ritalin, Metadate	FDA Schedule II
Dexmethylphenidate	Focalin	FDA Schedule II
Atomoxetine, Attentin	Strattera	Norepinephrine reuptake Inhibitor (NRI), not FDA Schedule II
Lisdextroamphetamine	Vyvanse	Pro-drug of D-amphetamine Not FDA-II

Alpha-2 Agonists

Generic Name	Brand Name(s)	Comment	
Clonidine	Catapres	More sedating than guanfacine	
Guanfacine	Tenex, Intuniv		

Frontal cortex / Locus Ceruleus: post-synaptic alpha-2 receptors
 Sympathetic outflow (autonomic nervous system): Pre-synaptic autoreceptors → ◆BP

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Alpha-2 Agonists

Benefits

- ♠ Attention Span
- No exacerbation of anxiety / rigidity

Side Effects

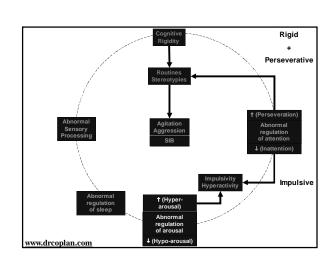
- Sleepiness: Common
- Emotional Lability (crying) - occasional
- Hypotension (low BP) rare

Clinical Pearl

- Beware of Cognitive Rigidity masquerading as ADHD
 - Perseveration on inner stimuli: "Inattentive"
 - Perfectionism:
 - "Problems w. task completion"
 - Anxiety:
 - "Rushes through work"
 - · "Out of seat behavior"

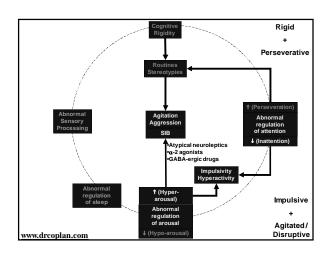
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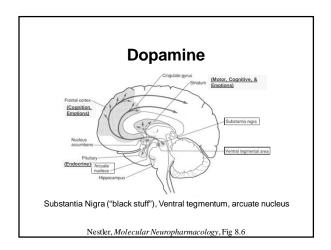
Regulation of Arousal Hypoarousal • Lethargic Calm & Fight or Flight Response "Red Alert" • Adrenaline • Heart Rate • Resp. Rate • Combative

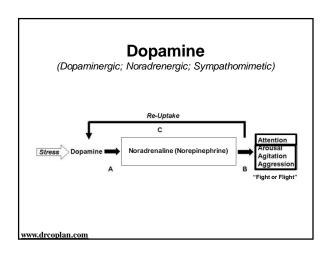


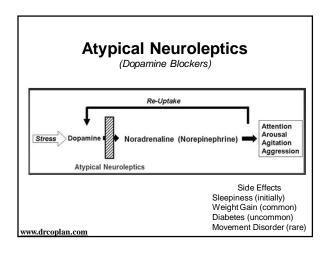
"He is so hard to calm down when he gets upset....His emotional thermostat doesn't work"

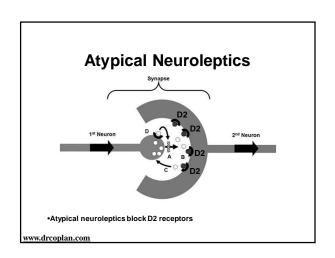
Parent of an 8 year old with ASD





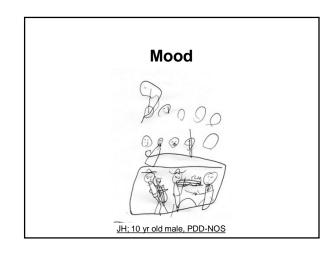


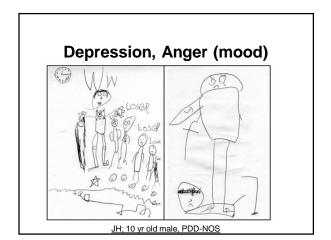


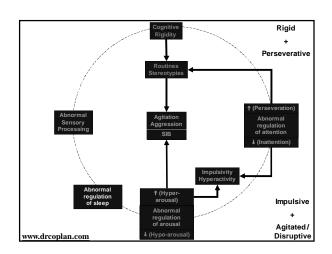


Atypical Neuroleptics

Generic Name	Brand Name	Comment
Aripiprazole	Abilify	 Relatively less risk of weight gain FDA approved for Rx of ASD
Clozapine	Clozaril	Bone marrow suppression
Olanzapine	Zyprexa	Greater risk of weight gain
Quetiapine	Seroquel	Greater sedation
Risperidone	Risperdal	Greater risk of weight gain FDA approved for Rx of ASD
Ziprazidone	Geodon	Relatively less risk of weight gain







Regulation of Sleep - 1

- Melatonin
 - Brain hormone
 - ★ Metabolic rate (Heart, Temp)
 - "You're sleepy now"
- Suppressed by light
 - 24 hr cycle
 - Seasonal cycle

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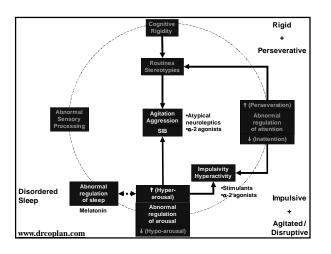
Regulation of Sleep - 2

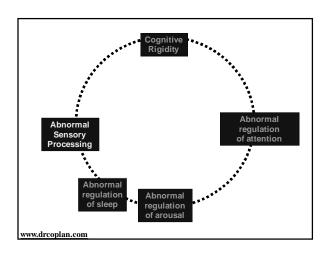
- Abnormal melatonin cycling
 - Primary disorders of sleep
 - Blindness
 - ASD
- Symptoms
 - Delayed onset of sleep
 - Shortened duration / frequent wakening

Regulation of Sleep - 3

- Shared genetic control
 - Regulation of sleep
 - Regulation of arousal
- Family history of sleep disorder

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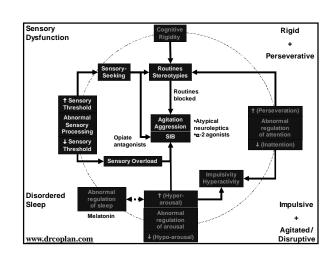


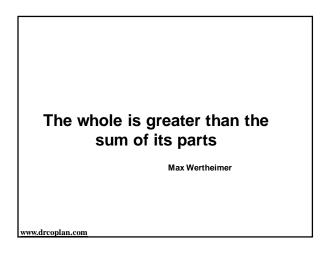


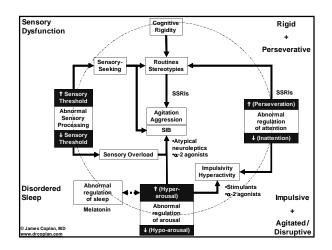
Sensory Processing

- Subjective Properties
 - Familiar / Unfamiliar
 - Pleasant / Unpleasant
 - Strong / Weak
 - Internal / External
- Sensory Input → Self-awareness
- Mirror Neurons → Empathy

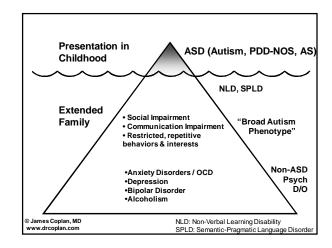
Mostofsky, S. and J. Ewen, Altered Connectivity and Action Model Formation in Autism Is Autism. Neuroscientist, 4/15/2011

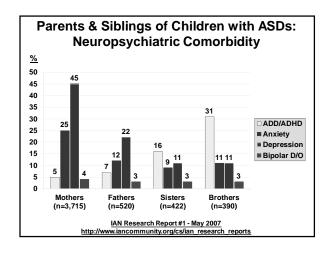


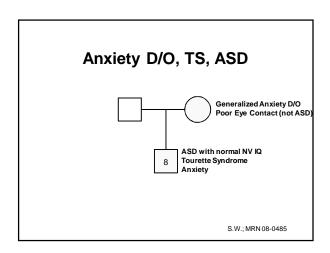


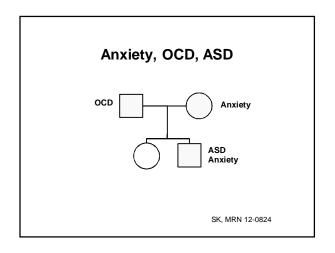


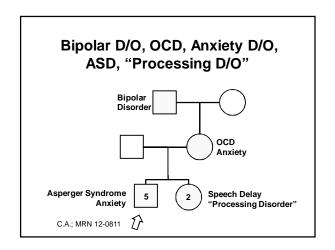
Prognosis & Family Genetics

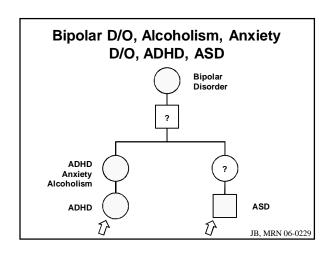


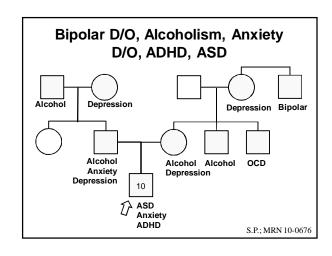


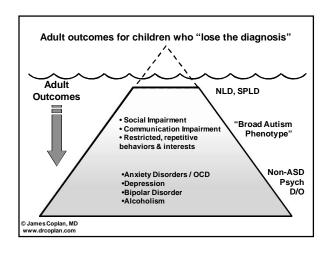


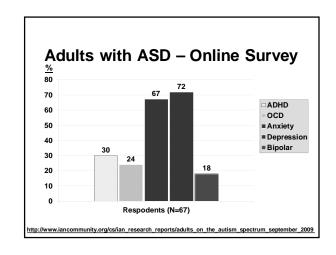


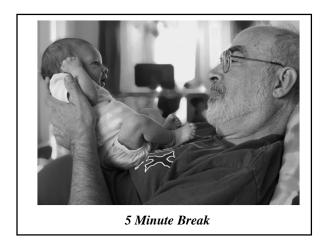


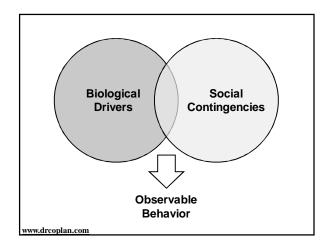












The ABC's of Behavior Analysis

- What is the Antecedent?
- What is the Behavior?
- What is the **C**onsequence?

Antecedents

- External:
 - Imposition of a task
 - Change in routine
 - Denial of access to object or activity
 - Other....
 - Or: No apparent external antecedent

Antecedents

- Internal:
 - Hunger
 - Thirst
 - Fatigue
 - Cognitive Rigidity
 - Dysregulation of Attention
 - Dysregulation of Mood / Arousal
 - Dysregulation of Sensory Processing
 - Other biological drivers

Behavior

- What, exactly, is the behavior?
 - "Topography"
 - Verbal
 - Physical
- Frequency
- Intensity
- Duration
- Timing

Behavior

- What is the child's developmental level?
- Is the behavior normal for the child's developmental level?
 - Tantrums / Noncompliance
 - 24 month old "striving for autonomy"
 - Cognitive Rigidity
 - 4 yr olds are "rule based" by nature
 - "Impulsivity" / "Inattention"
 - · Attention span is a function of devel. level

Behavior

- What is the child's ability to communicate?
 - Does "disruptive" behavior serve a communicative function?

Behavior

- Acute change or chronic?
- General health?
 - Vital signs, I&O, Level of consciousness
 - Pain?
- Anything new in child's life?
 - Recent change of meds

Consequences

- Reinforcers
 - Positive
 - Negative
- Aversives

Consequences 1: Reinforcers

- Reinforcers lead to an increase in frequency of the antecedent behavior
 - Positive Reinforcement (adds something)
 - Attention
 - · Access to desired object / activity
 - Negative Reinforcement (removes something)
 - Escape from Task
 - Avoidance of punishment
 - -"We drive within the speeding limit because we receive the negative reinforcement of not getting a speeding ticket"

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Food Selectivity

Negative and Positive Reinforcement of unwanted behavior

- Parent removes non-preferred food ([-] reinforcement)
- Parent provides child with his/her preferred food ([+] reinforcement)
- Alternatives
 - FirstThen (The Premack Principle)
 - Put refusal on extinction
 - The kitchen is closed between meals

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Consequences 2: Aversives

- Aversives lead to a decrease in the likelihood of recurrence of the antecedent behavior
- Logical Consequences
 - If child refuses to use toilet, child must carry backpack with spare clothes, when family is in public
- Over-correction
 - Must wash out soiled diaper
 - If the child spills milk on purpose: child must mop the entire kitchen floor

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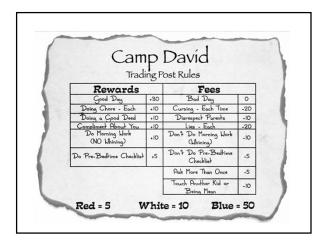
Disruptive Behavior: Function & Best Response

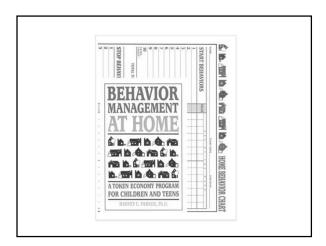
- Attention
 - 1-2-3 **⇒** "Time Out" (T.O.)
- Access
 - Never grant access to desired object in response to disruptive behavior
- Escape
 - Never permit the child to escape from a task via disruptive behavior.
 - Walk child through task first, then ⇒ T.O.
 - OR: Send child to T.O., and as soon as T.O. is complete, resume the task where you left off.

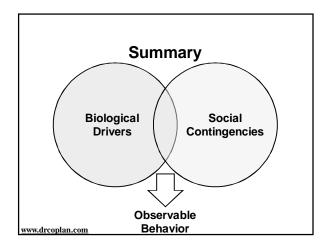
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Token Economy: The next step beyond Time Out

- · Concretely specified behaviors
- Earn and Lose Points
- Points
 → Access to preferred items
 - Preferred toys, Computer time, etc.
 - NO access to preferred item at other times
 - "Extra" treats not as effective
- Works with children who understand rule-based play (CandyLand, Uno, etc.)







An ounce of prevention....

- Identify internalizing behaviors before they lead to externalizing behaviors
 - Behavior Management Plan that proactively seeks to avert or dissipate anxiety

Summary

- · Why this child?
 - What is this child's developmental Level?
 - Is this stage-appropriate behavior?
 - Does the behavior serve a social function?
 - · Escape, access, attention
 - Is the classroom placement appropriate?
 - · Language level?
 - Does this behavior occur in other settings?
 - · Family factors?
 - Parents consistent at home?
 - Parental psychopathology? (Anxiety, Depression, Alcohol)

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Summary

- · Why this child?
 - Neuropsychological factors?
 - Cognitive Rigidity
 - · Dysregulation of attention
 - Dysregulation of arousal
 - Sensory Seeking / Sensory Overload
- · Behavioral Intervention Usually
- · Change in classroom setting sometimes
 - Shift from rote to inferential learning (2nd 3rd grade): challenge
- · Medication: Sometimes

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Summary

Directions for future research:

- · Better phenotyping of ASD
 - Clinical
 - Genetic
- · Better drug studies
 - Drug vs. Behavioral Therapy vs. Combination
 - Drug vs. Drug (not just drug vs. placebo)
 - Drug combinations (not just monotherapy)
 - Stimulant + SSRI, e.g.
 - Better outcome measures
 - Quality of Life
 - Long-term outcome
- Brain / Behavior / Drug imaging

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Summary

Services for adult "survivors" of childhood ASD

- Mental Health
 - Anxiety
 - Depression
 - Mood Disorders
 - Depression
- Vocational
- · Workplace Social Skills
- Parent / Caregiver support
- · Spouse / Partner support

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Disclosures / References

 Dr. Coplan is author of Making Sense of Autistic Spectrum Disorders: Create the brightest future for your child with the best treatment options (Bantam-Dell, 2010), and receives royalties on its sale



Chapter 12: Behavior management and psychopharmacology

This presentation will include a discussion of off-label drug use



