

Pay Attention: ADHD Through the Lifespan

Anthony L. Rostain, MD, MA
Professor of Psychiatry and Pediatrics
Perelman School of Medicine at the
University of Pennsylvania

Week 7: Complications of ADHD



Homework Review

Co-morbidity seen with ADHD

- | | |
|-------------------------|---|
| Externalizing disorders | <ul style="list-style-type: none">• Oppositional Defiant Disorder• Conduct Disorder |
| Internalizing disorders | <ul style="list-style-type: none">• Anxiety Disorders• Mood Disorders<ul style="list-style-type: none">– Bipolar Disorder– Major Depression– Dysthymia |
| Cognitive disorders | <ul style="list-style-type: none">• Learning Disorders• Language Disorders |

Co-morbidity seen with ADHD

Special populations

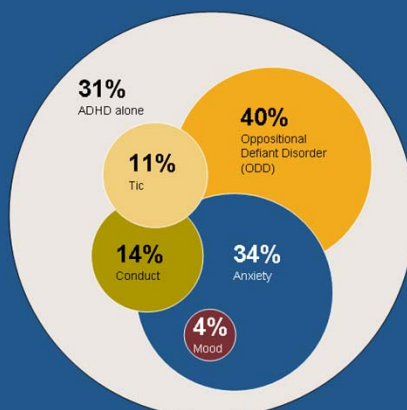
- Tourette Syndrome
- Obsessive Compulsive Disorder
- Autistic Spectrum Disorders
- Fetal Alcohol Syndrome

Other problems

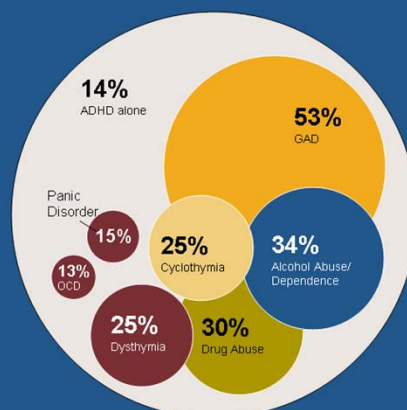
- Sleep Disorders
- Substance Use Disorders
- Post-Traumatic Stress Disorder

Patients With ADHD Frequently Have Coexisting Disorders

Children & Adolescents



Adults



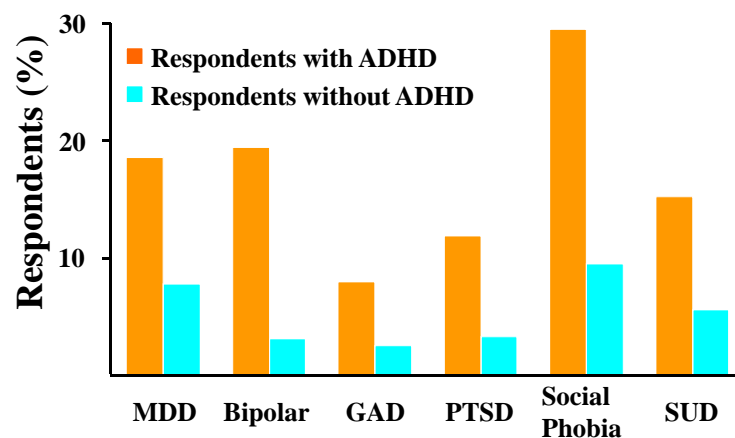
MTA cooperative: N=579.

Factors Related to Comorbidity

- Greater severity of ADHD symptoms
- Non-intact families of origin
- Low parental income
- Decreased parental life satisfaction
- Decreased parental interest in and monitoring of the child
- Affiliation of patient with deviant peers
- Parental ADHD, depression, antisocial personality disorder, & substance abuse

Hurtig, T. et al. (2007). *European Child and Adolescent Psychiatry*, 16, 362-369.
Also Barkley, R. (2006). *ADHD: A Handbook for Diagnosis & Treatment*. New York: Guilford.

Common Comorbid Psychiatric Disorders (Epidemiological Sample)



National Comorbidity Survey Replication (N = 3199)

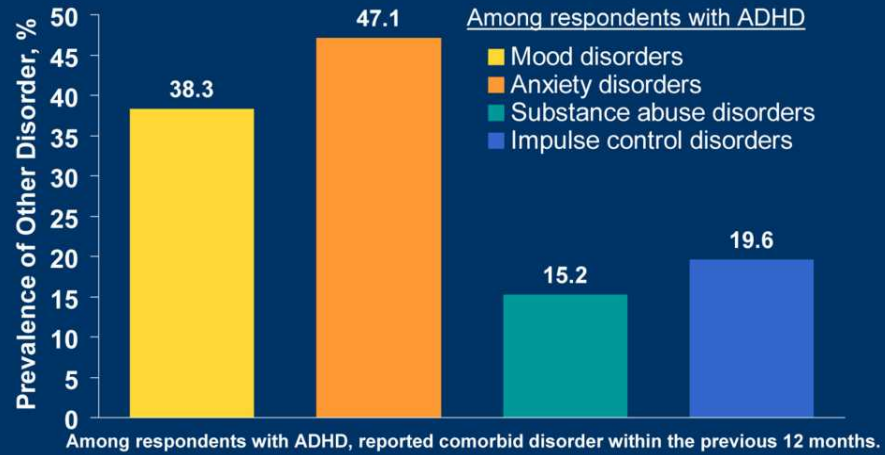
Among respondents aged 18-44 years with ADHD, comorbid disorder within previous 12 months
For all comparisons, $P < 0.05$

Kessler et al. *Am J Psychiatry*. 2006;163:716-723.

8

Comorbidity of Other DSM-IV-TR Disorders With ADHD in Adults

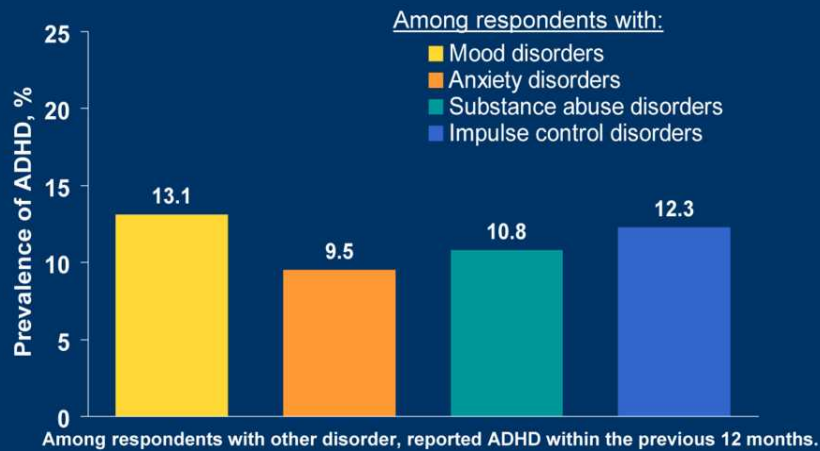
National Comorbidity Survey Replication (N=3199)



Kessler RC, et al. *Am J Psychiatry*. 2006; 163:716-723.

Comorbidity of ADHD With Other DSM-IV-TR Disorders in Adults

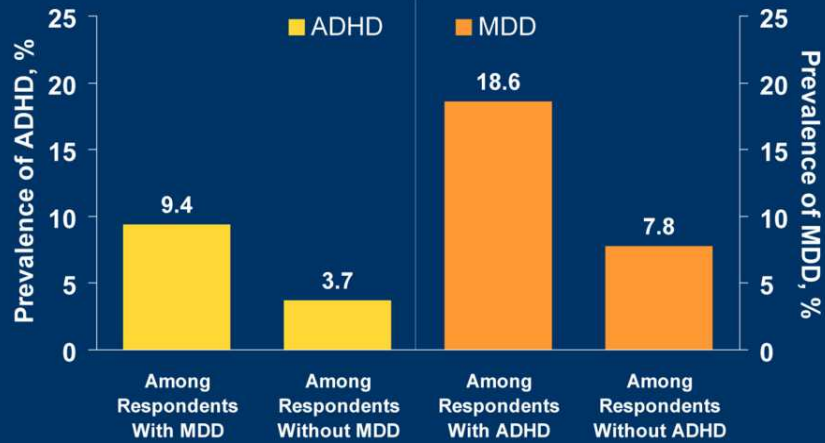
National Comorbidity Survey Replication (N=3199)



Kessler RC, et al. *Am J Psychiatry*. 2006; 163:716-723.

Comorbidity of Adult ADHD With MDD in Adults

National Comorbidity Survey Replication (N=3199)



Kessler RC, et al. *Am J Psychiatry*. 2006; 163:716-723.

ADHD and Comorbidity in Adults

Symptom domains

- Hyperactivity
- Inattention
- Impulsivity

+

Psychiatric comorbidities

- Anxiety and mood disorders
- Disruptive behavior disorders (conduct disorder and oppositional defiant disorder)

Lead to

Functional impairments

Self

- Low self-esteem
- Accidents and injuries
- Smoking
- Substance abuse
- Delinquency

School / Work

- Academic difficulties, underachievement
- Employment difficulties

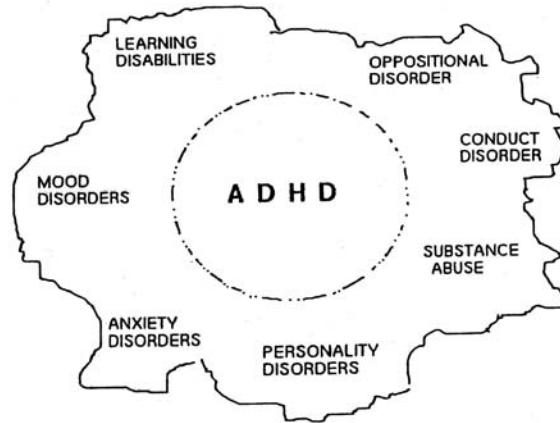
Home

- Family stress
- Parenting difficulties

Social

- Poor peer relationships
- Socialization deficit
- Relationship difficulties

Elusive Boundaries of ADHD



Mid-Lecture Questions

Disruptive Disorders (ODD, CD)

Oppositional Defiant Disorder – Diagnostic Criteria

A pattern of negative, hostile and defiant behavior lasting at least 6 months, during which four (or more) of the following are present:

- 1) often loses temper
- 2) often argues with adults
- 3) often actively defies or refuses to comply with adults' requests or rules
- 4) often deliberately annoys people
- 5) often blames others for his or her mistakes or misbehavior
- 6) is often touchy or easily annoyed by others
- 7) is often angry and resentful
- 8) is often spiteful or vindictive

Oppositional Defiant Disorder (40-80%)

- ADHD cases are 11x more likely to have ODD
- ADHD contributes to and likely causes ODD
 - This likely occurs through the impact of ADHD on emotional self-regulation (an executive function)
 - This can account for the well-established findings that ADHD medications reduce ODD as much as they do ADHD
- Some ODD is related to disrupted parenting
 - Inconsistent, indiscriminate, emotional, and episodically harsh and permissive (lax) consequences teaches social coercion as a means of social interaction
 - Poor parenting can arise from parental ADHD and other high risk parental disorders in ADHD families
- Early ODD predicts persistence of ADHD and increases risk for CD/MDD and anxiety disorders

Treatment Impact of ODD

- Both stimulants and atomoxetine reduce ODD when it is comorbid with ADHD; not when ODD is alone
 - Higher doses may be needed for comorbid cases
- Requires adjunctive parent training in behavior management methods; response is age-related:
 - 60-75% successful for children
 - 25-35% treatment response after 13+ yrs. of age
 - May need to treat parent's ADHD first to succeed
 - May need to add problem-solving communication training of teen and parents after age 14 years

Conduct Disorder – Diagnostic Criteria

A repetitive and persistent pattern of behavior in which the basic rights of others or major age-appropriate societal norms or rules are violated, as manifested by the presence of **three** (or more) of the following criteria in the past 12 months, with at least one criterion present in the past 6 months:

- **Aggression to people and animals**
 - 1) often bullies, threatens, or intimidates others
 - 2) often initiates physical fights
 - 3) has used a weapon that can cause serious physical harm to others (e.g., a bat, brick, bottle, knife, gun)
 - 4) has been physically cruel to people
 - 5) has been physically cruel to animal
 - 6) has stolen while confronting a victim (e.g., mugging, purse snatching, extortion, armed robbery)
 - 7) has forced someone into sexual activity

Conduct Disorder – Diagnostic Criteria

- **Destruction of property**
 - 8) has deliberately engaged in fire setting with the intention of causing serious damage
 - 9) has deliberately destroyed others' property (other than by fire setting)
- **Deceitfulness or theft**
 - 10) has broken into someone else's house, building, or car
 - 11) often lies to obtain goods or favors or to avoid obligations (i.e., "cons" others)
 - 12) has stolen items of nontrivial value without confronting a victim (e.g., shoplifting, but without breaking and entering; forgery)

Conduct Disorder – Diagnostic Criteria

- **Serious violations of rules**
 - 13) often stays out at night despite parental prohibitions, beginning before age 13 years
 - 14) has run away from home overnight at least twice while living in parental or parental surrogate home (or once without returning for a lengthy period)
 - 15) is often truant from school, beginning before age 13 years
- *Specify type based on age at onset:*
 - Childhood-Onset Type:** onset of at least one criterion characteristic of Conduct Disorder prior to age 10 years
 - Adolescent-Onset Type:** absence of any criteria characteristic of Conduct Disorder prior to age 10 years

ADHD & Conduct Disorder (20-56%)

- If starts early, represents a unique family subtype
 - More severe, more persistent antisocial behavior
 - Worse family psychopathology
 - Antisocial personality, substance use disorders, major depression
 - Parent hostility, depression, & low warmth and monitoring interact reciprocally with child conduct problems over time to adolescence**
 - Greater association with ADHD (especially inattention symptoms)
 - Less responsive to behavioral or family interventions
- Increased risk of psychopathy (20%)

*Barkley, R. A. et al. (2008). *ADHD in Adults: What the Science Says*. New York: Guilford.

** *Special issue on reciprocal influence across development, Journal of Abnormal Child Psychology* (2008), vol. #36 (July).

ADHD & Conduct Disorder (20-56%)

- Increased risk of psychopathy (20%)
- Father desertion, parent divorce more common
- Major depression more likely to precede/co-exist with CD
- If CD starts late (>12), related to social disadvantage, family disruption, & affiliation with deviant peers
- School drop out, drug use, and teen pregnancy are more likely in comorbid cases than in ADHD alone*

*Barkley, R. A. et al. (2008). *ADHD in Adults: What the Science Says*. New York: Guilford.

** Special issue on reciprocal influence across development, *Journal of Abnormal Child Psychology* (2008), vol. #36 (July).

Treatment Impact of CD

- Stimulants and atomoxetine reduce aggressive behavior and antisocial acts but stimulants may work more rapidly to gain case control
 - Higher doses often required in comorbid cases
 - Stimulant effectiveness may deteriorate with duration of treatment (3+ yrs) in this subset of ADHD cases (MTA study)
- Parent and family interventions often required
 - Problem-solving, communication training and parent BMT
 - Multi-systemic therapy where available
 - Treatment of parental depression and other psychiatric disorders
 - Family relocation to better neighborhoods advisable

*Waschbusch, D. A. et al. (2007). *Journal of Clinical Child and Adolescent Psychology*, 36(4), 629-644.

Treatment Impact of CD

- If psychopathy (callous-unemotional traits) is present there is limited or no response to behavior therapy alone – medication is necessary first, then follow up with behavioral treatments*
- Avoid group treatment due to deviancy training by aggressive peers
- Involvement of social service and juvenile justice agencies is highly likely
- Mood stabilizers, atypicals, or antihypertensives may be needed for highly aggressive/explosive cases or Bipolar Disorder

*Waschbusch, D. A. et al. (2007). *Journal of Clinical Child and Adolescent Psychology*, 36(4), 629-644.

Mid-Lecture Questions

Anxiety and Mood Disorders

Significance of Anxiety Disorders in Children and Adolescents

- 10% - 20% of school age children exhibit symptoms of social withdrawal, anxiety, isolation, hypersensitivity, depression & self-consciousness
- Many experience debilitating physical symptoms: headaches, stomachaches & irritable bowel syndrome
- Child anxiety disorders persist & even worsen over time

Categories of Anxiety Disorder

- Separation Anxiety Disorder
- Social Anxiety Disorder (Social Phobia)
- Selective Mutism
- Generalized Anxiety Disorder (GAD)
- Specific (Simple) Phobia
- Panic Disorder
- Obsessive-Compulsive Disorder (OCD)
- Posttraumatic Stress Disorder (PTSD)

Common Themes Across All Anxiety Disorders

- Fear signals are triggered in situations where there is no serious threat of harm (“overestimation of danger”)
- Minimal or moderate stress produces an exaggerated reaction (“over-reactivity”)
- Intense/out of proportion anxiety results in *distress* and/or functional *impairment*

ADHD and Anxiety Disorders (10-40%)

- Considered a stealth or hidden comorbidity in child ADHD cases if only parents are interviewed about child anxiety symptoms. High comorbidity with adult ADHD (30%+)
- Related in part to poor emotion regulation in ADHD
- Some legitimate anxiety disorders are likely and risk increases with age
- Most common are simple phobias or separation anxiety in early childhood; GAD becomes more common with age
- Risk is related to earlier inattention more than to impulsive-hyperactive*

* Reinke, W., & Ostrander, R. (2008). *Journal of Abnormal Child Psychology*, 36(7), 1109-1122.

**See Pfiffner, L. & McBurnett, K. (2006). *Journal of Abnormal Child Psychology*, 34, 725-735. Also Kopley, H., & Ostrander, R. (2007). *Journal of Attention Disorders*, 10, 317-323.

ADHD and Anxiety Disorders (10-40%)

- Greater association with disruptive and stressful events
- Often show lower levels of impulsiveness
- Anxiety disorders more likely in parents and family**
 - Child and parental anxiety are associated with low rates of positive parental behavior, over-protectiveness of the child, less autonomy for the child, lower child self-sufficiency, and parent modeling of anxiety
- Comorbid cases are more impaired than ADHD alone cases
- Need to look more carefully for child physical or sexual abuse, or bullying at school

* Reinke, W., & Ostrander, R. (2008). *Journal of Abnormal Child Psychology*, 36(7), 1109-1122.

**See Pfiffner, L. & McBurnett, K. (2006). *Journal of Abnormal Child Psychology*, 34, 725-735. Also Kopley, H., & Ostrander, R. (2007). *Journal of Attention Disorders*, 10, 317-323.

Treatment Impact of Anxiety Disorders

- More responsive to behavioral therapies
- May respond better to social skills training (and possibly cognitive therapies)
- Treatment can reduce school avoidance and social phobia
- Stimulants can exacerbate anxiety in some cases:
 - Studies are conflicting – 8 say it can but MTA study and Abikoff (2005) study did not find this relationship

Treatment Impact of Anxiety Disorders

- Atomoxetine (ATX) significantly reduces anxiety
 - Effect Size = .3-.5
- AACAP revised Practice Parameters recommend ATX as a possible first choice treatment in these comorbid cases
- Family counseling may be required to limit family setting induction of anxiety; focus parent BMT on increasing positive parenting behavior and reducing over-protectiveness and less so on parent discipline tactics
- Try cognitive behavioral training with ADHD adults to further cope with anxiety disorders

Definition & Criteria of Mood Disorders

Major Depressive Disorder: feelings of unhappiness inconsistent with one's circumstances

Dysthymic Disorder: chronic condition of unhappiness but does not reach clinical levels

- Depressed (sad) or expansive mood
- Decreased energy, stamina, activity level
- Loss of interest or pleasure in almost all activities (anhedonia)
 - Changes in appetite, weight, sleeping patterns
- Irritability, poor concentration, or difficulty making decisions
- Feelings of worthlessness & thoughts of death or suicide

Major Depression (0-45%)

- Likely genetic linkage to ADHD
- Genes create a vulnerability to MDD
- MDD expressed upon exposure to repeated social and emotional distress, physical trauma, etc.
- Also related to presence of CD in child or adult patient & family
- Often manifest low self-esteem in childhood in contrast to other ADHD cases
- MDD onset may not be until adolescence or later
 - In adults with ADHD, MDD is related to higher GAD and social phobia but lower SUDS and school disciplinary actions and grade repetitions in history***

*Elgar et al. (2007). Maternal and paternal depressive symptoms and child maladjustment: The mediating role of parental behavior. *Journal of Abnormal Child Psychology*, 35, 943-955.

*Gerdes, et al. (2007). Maternal depressive symptomatology and parenting behavior: Exploration of possible mediators. *Journal of Abnormal Child Psychology*, 35, 705-714.

***Fischer, A. et al. (2007). The role of comorbid major depressive disorder in the clinical presentation of adult ADHD. *Journal of Psychiatric Research*, 41, 991-996.

Major Depression (0-45%)

- Associated with increased suicidal ideation (4x) and attempts (2x) in ADHD cases during peak risk years in high school
- Parental depression is elevated in these child cases; depressed parents:
 - show decreased positive parenting and nurturance, greater irritability and expressed emotion, irritability and open hostility, erratic use of discipline tactics, child rejection, and poor child monitoring – these are associated with increased later risk for child behavior and internalizing problems*
- Evaluate carefully for presence of child physical or sexual abuse or victimization by bullying in child cases

*Elgar et al. (2007). Maternal and paternal depressive symptoms and child maladjustment: The mediating role of parental behavior. *Journal of Abnormal Child Psychology*, 35, 943-955.

*Gerdes, et al. (2007). Maternal depressive symptomatology and parenting behavior: Exploration of possible mediators. *Journal of Abnormal Child Psychology*, 35, 705-714.

***Fischer, A. et al. (2007). The role of comorbid major depressive disorder in the clinical presentation of adult ADHD. *Journal of Psychiatric Research*, 41, 991-996.

ADHD and Suicidality

- ADHD is associated with a greater risk for suicidal ideation & suicide attempts*
 - Ideation in high school (33 vs. 22%)
 - Attempts in high school (16 vs. 3%)
 - Attempts are worse (46% vs. 11% hospitalized)
 - Ideation after high school (25% vs. 12%), attempts 6 vs 3%); risks for ideation found even at age 27
 - Associated with comorbid MDD (4x), CD (somewhat), and more severe ADHD
 - Evaluate carefully for child physical or sexual abuse or victimization by bullying

*Barkley, R. A. & Fischer, M. (2005). *The ADHD Report*, 13 (6), 1-4.

*Barkley, R. A., Murphy, K. R., & Fischer, M. (2008). *ADHD in adults: What the science says*. New York: Guilford

Treatment Impact of MDD

- May require mixed ADHD/SSRI therapy
 - Stimulants and atomoxetine do not treat MDD
- May need cognitive-behavioral therapy
- Assess for parental induction of depression in children and exaggeration of child ODD symptoms given higher maternal depression
- Parent depression may require separate treatment
- In parent training use a “go slow” approach to punishment so as not to contribute to depressive cognitive schemas (self-statements) or to already excessive parental use of criticism and discipline
 - start with all reward programs initially until MDD symptoms lift then introduce mild, selective punishments.



ADHD and Bipolar Disorder

Overlapping symptoms

- Poor attention
- Hyperactivity
- Impulsivity

Bipolar symptoms

- Expansive mood
- Grandiosity
- Manic quality
- Rage episodes/severe irritability
- History of bipolar disorder in first-degree relatives

No evidence-based treatment exists for this patient population. Expert consensus suggests treating bipolar disorder first.

Sheffer et al. *Am J Psychiatry*. 2005;162:58-64. Wilens et al. *Biol Psychiatry*. 2003;54:9-16. Rapaport. *N Engl J Med*. 2005;352:165-173. Pliszka. *Pediatr Drugs*. 2003;5:741-750. Pavuluri et al. *Biol Psychiatry*. 2006;60:936-941.

Child Bipolar Disorder (C-BPD) (3-6%)

- Controversial - 0-27% incidence in ADHD cases
- Not documented in follow-up studies, however
- Some cases are misdiagnosed (BPD vs. ADHD/ODD)
 - Several ADHD symptoms appear on bipolar symptom list in DSM
 - Irritability could substitute for mania in children (could be ODD)
 - No requirement for cycling or periods of remission in children
- DSM-V will likely correct these issues
 - Diagnosis will require mixed moods (bipolarity), cycling between them, grandiosity, mania and other typical bipolar symptoms which can be present in children (see next slide)
- Family history of BPD is common in child BPD; not in ADHD
 - Childhood BPD has 7-8x family risk of BPD than does ADHD or adult onset BPD
- Probably a one-way comorbidity – 3-6% of ADHD cases have BPD while 80-97% of child BPD have ADHD

Differential diagnosis from ADHD

- Irritability: 95% (BPD) vs. 65% (Disruptives)
- Elation: 73% vs. 15%
- Grandiosity: 80% vs. 10%
- Excessive talking: 92% vs. 38%
- Racing thoughts: 30% vs. 5%
- Flight of ideas: 69% vs. 14%
- Decreased sleep: 42% vs. 12%
- Pressured speech/motor: 84% vs. 35%
- More active: 85% vs. 43%
- Uninhibited socializing: 32% vs. 3%
- Hypersexual: 53% vs. 3%

From Luby & Belden, 2006, *Development and Psychopathology*, 18, p. 971

C-BPD Diagnostic Keys

- Mood regulation is a major problem and moods are often severe (i.e., rage attacks, violence, destructive)
- Mood states are not related to immediate environmental events in a rational sense (irrational and inconsistent)
 - ADHD kids have rational but somewhat excessive emotions
- Grandiosity, elated mood, psychotic-like thinking (paranoia, delusions, auditory hallucinations, disjointed thought), and hyper-sexuality are involved in C-BPD but not in ADHD
- Overlapping ADHD symptoms are far more severe (distractible, impulsive, hyperactive, decreased sleep, excessive talking)
- Disruptive (aggressive) behavior rated as 3+SDs on rating scales like the CBCL (85 or higher) goes with CBPD, not with ADHD
- BPD is significantly more prevalent in biological relatives

Mid-Lecture Questions

Substance Use Disorders

Is ADHD a Risk Factor for SUD?

Many clinical studies report a connection, however

- Most are not population based (referral bias)
- Many are retrospective (recall bias)
- Some don't take co-morbidity into account

Which ADHD Children Are at Greatest Risk for SUD?

- Comorbid psychiatric disorders
- Family history of SUD (may contribute to both ADHD and SUD)
- Persistent ADHD
- Other biological markers?
- Social skills deficits

Which ADHD Children Are at Greatest Risk for SUD?

- Severity of childhood symptoms?
- Inattention (for tobacco)?
- Impulsivity or disinhibition (for other drugs)?
- Gender differences: findings contradictory so far
- Ethnic or racial group differences: inadequately studied so far

Which ADHD Children Are at Greatest Risk for SUD?

Biological associations:

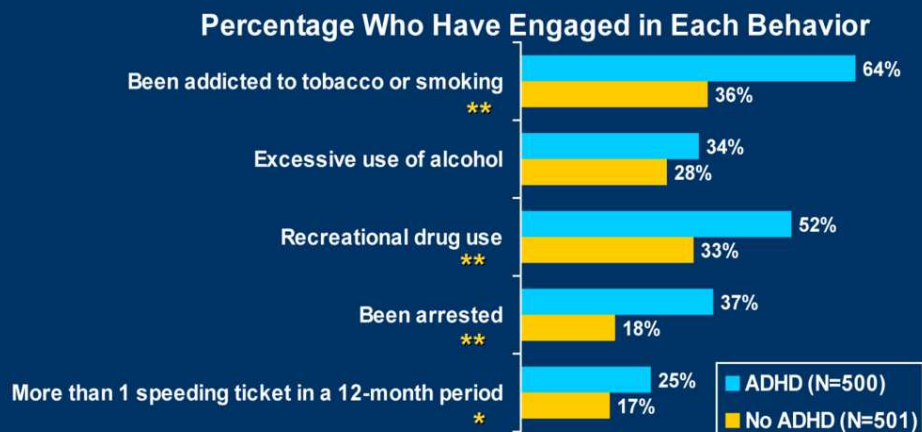
- Through prenatal exposure to alcohol, smoking, perhaps drugs
- Low birth weight
- Dopaminergic system: Self-medication? (especially tobacco)
- Perhaps an internalizing/inattentive/self-medicating late-onset subtype?
- Perhaps sensitization through use of stimulants

Which ADHD Children Are at Greatest Risk for SUD?

Psychosocial factors that might impact use/early use:

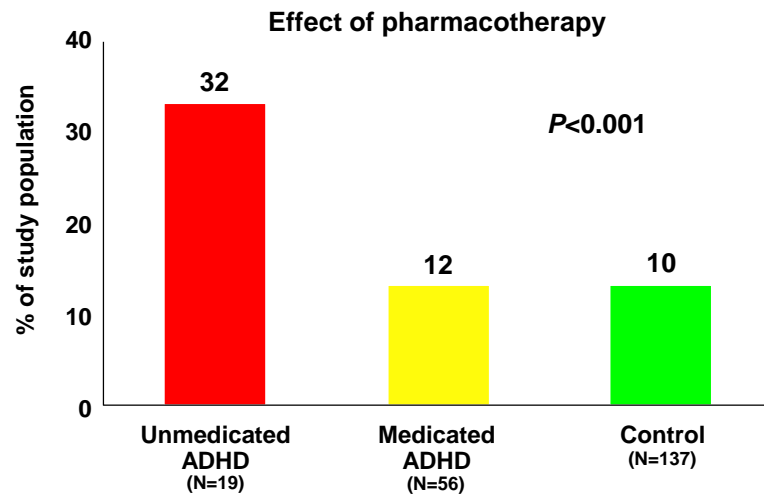
- Weak attachment to & conflict with parents, school secondary to behavior problems
- Disordered alcohol or drug expectancies
- Association with deviant peers
- Attribution (fulfilling expectations)?
- Parental modeling, monitoring, coping (ADHD parents or child-induced)

Prevalence of Addiction, Antisocial, or Destructive Behaviors



Biederman et al. *J Clin Psychiatry* 2006;67:524.

Pharmacotherapy Significantly Reduces Substance Abuse in Adults with ADHD



Biederman J, et al. *Pediatrics*. 1999;104:e20-e25.

Quiz Questions