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# ADHD: A Neurodevelopmental Disorder Through the Ages



### ADHD - Points to be addressed

How the diagnosis is made.

Controversies

Why diagnosis is important.

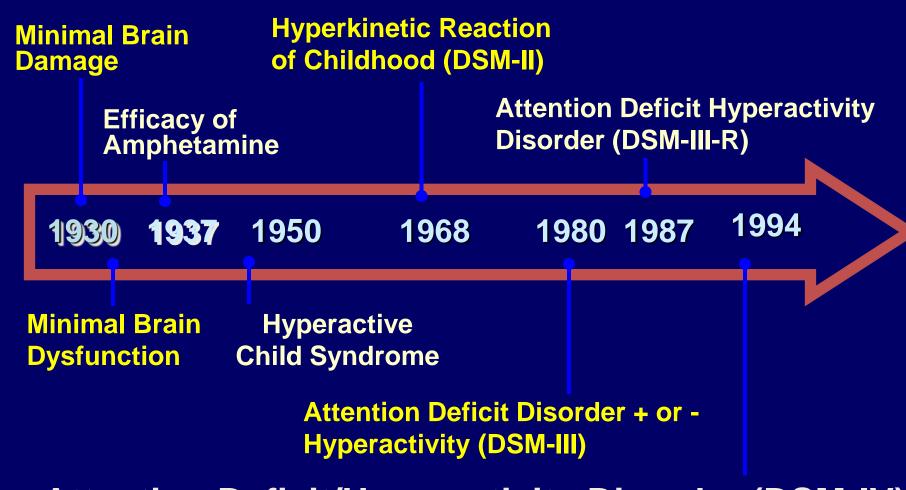
Historical aspects.

Age related manifestations.

What happens through life.

Treatment options.

### ADHD Historical Timeline



Attention Deficit/Hyperactivity Disorder (DSM-IV)

### How do we diagnose ADHD?

- In children, ADHD is diagnosed based on reports of behavior by caretakers, and other adults, especially teachers.
- The behaviors are extremes of common, ordinary, behaviors.
- Controversies arise from such behavioral approaches.

### Diagnoses do NOT Include Variations of Normal Development

#### Some Examples:

- Tantrums in a 2 year old
- Distress at separation in early childhood
- Fear of animals at age 4
- Sibling rivalry
- Feeling down after a loss
- Resenting authority
- Lying to avoid being punished

## When a child has a psychiatric disorder

- Important functions are delayed or impaired
- The dysfunctions are not under easy willful control (inflexible)
- The dysfunction are not reversed by simple environmental change
- There is suffering or Impairment

## Controversies about diagnosing children

- >We are medicalizing variations in normal development.
- ➤ Being young means not going with the flow being different is normal.
- Diagnosing children stigmatizes them (no evidence for this).
- >There are legitimate concerns, BUT

## Important benefit of psychiatric classification-1

✓ We can help children and their families.

✓ There are treatments that work.

✓ We know that child and adolescent psychiatric disorders are not innocuous. They incur risk for future dysfunction in a proportion (not all).

### Important Benefits of Psychiatric Classification-2

Communication: enables a common language.

Clinical Care: guides treatment choices. Prognosis: tells us what we may expect over time (recovery/other problems).

Knowledge: unless we classify conditions, we cannot study them. We remain ignorant about what is best for the child.

### Important Benefits of Psychiatric Classification-3

Knowledge:

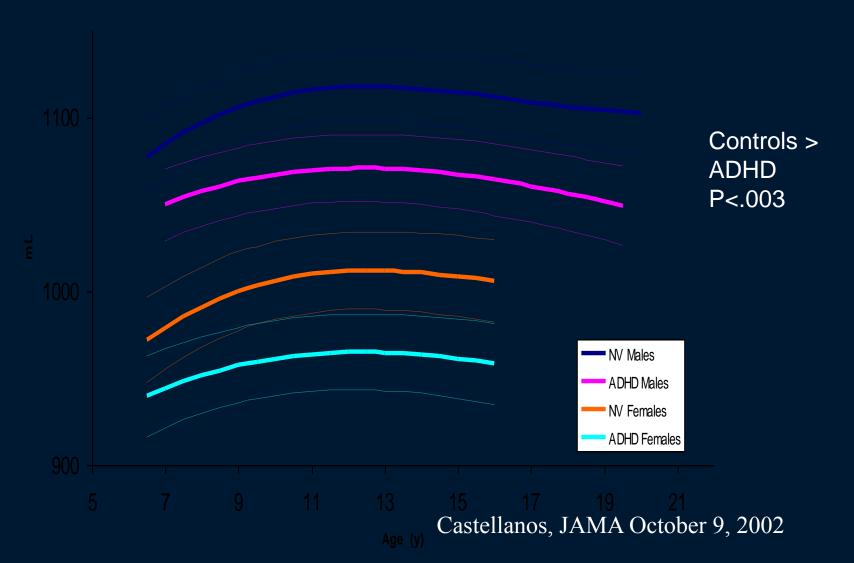
 Studies of brain development have led to new insights about ADHD.

Systematic studies that would not have been possible without the diagnosis have shown that **ADHD** is a "brain disorder" or a "neurodevelopmental disorder."

#### 1) ADHD has a strong genetic component

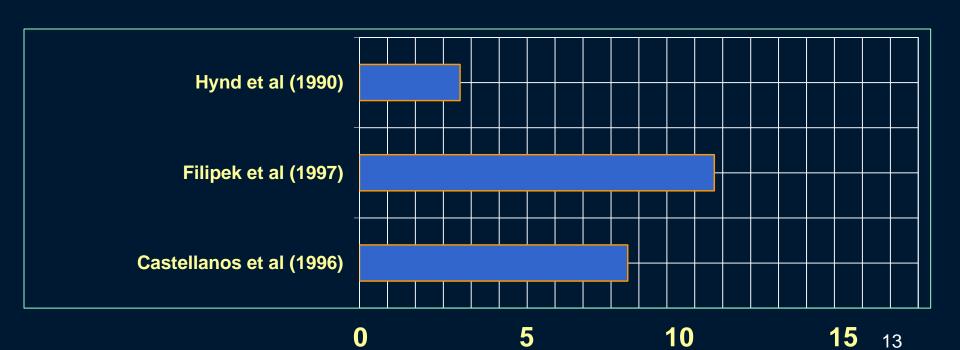
- up to 92% concordance in monozygotic twins
- heritability of 0.75
- molecular genetic studies have implicated specific genes
- 2), children and adults with ADHD have thinner cortical volumes than normal children.

#### Total Cerebral Vol. Growth Curves



#### **ADHD - Anatomic MRI Studies**

Frontal Lobes Percent decrease in size in Individuals with ADHD Compared to controls (dozens of additional studies)



## Is Cortical Thickness Clinically Relevant?

Longitudinal study at NIMH found that:

Children with ADHD who had thinner prefrontal cortex than normal children were more likely to retain ADHD at follow-up+ than children whose prefrontal cortex\* was no different from controls.

- +5.7 year follow-up to age 13
- \* No effect of total cortex volume.

### Importance of the Disorder

- ✓ Elevated prevalence in the population (abt 5%)
- ✓ Most common disorder in child psychiatric clinics
- ✓ Incurs impairment in multiple domains of function at ALL AGES
- Can have deleterious long-term consequences

## Functional impairment with ADHD at all Ages

- ✓ Interferes with learning
- ✓ Problematic relationships with adults and peers
- √ Rejected by peers
- ✓ Stress on the environment
  - School or Work Place
  - > Family

#### **ADHD**

 Inattention, hyperactivity, impulsivity that are inconsistent with developmental level and lead to significant problems for the person.

- The overt manifestations of ADHD vary with developmental level
  - Preschool
  - School age (6 12)
  - Adolescence
  - Adulthood

#### Inattention

- ✓ Careless mistakes
- ✓ Difficulty sustaining attention
- √ Seems not to listen
- √ Fails to finish tasks
- ✓ Difficulty organizing
- ✓ Avoids tasks requiring sustained attention
- ✓ Loses things
- ✓ Easily Distracted
- √ Forgetful

#### **Hyperactivity**

- ✓ Unable to stay seated
- ✓ Moving excessively (restlessness)
- ✓ Difficulty engaging in leisure activities quietly
- √"On the go"
- √ Talking excessively

### **Impulsivity**

- √ Blurting out answers
- ✓ Difficulty awaiting turn
- ✓Interrupting/intruding upon others
- **√Impatient**

### Well-Documented Domains of Impairment in Individuals with ADHD (at all ages)

- ✓ Social Relationships
- **✓** Family Function
- ✓ School or Work Performance, and/or Adjustment

# Impairment – All Ages (Social Relationships)

- √ Significantly impaired relationships
- ✓ Often loud and intrusive
- √Others quickly form negative impressions, leading to rejection.
- ✓ Negative social relationships affect all important functions (work, marriage, parenting)

### Impairment (Family Function)

- √ Families have high levels of conflict
- √ Family members are stressed
- ✓ Parents are often overwhelmed and demoralized

## Impairment in Children (Academic Performance)

- ✓ Significantly more school failure
- ✓ Many require special tutoring
- ✓ Placement in special classes or having to repeat a grade is common
- ✓ Rates of learning disorders range from 10% to 20%

## Impairment with ADHD in Children (School Adjustment)

✓ Teachers see as working less hard, learning less, behaving less appropriately

✓ Disrupts the class; parents often have to visit the school about child's behavior

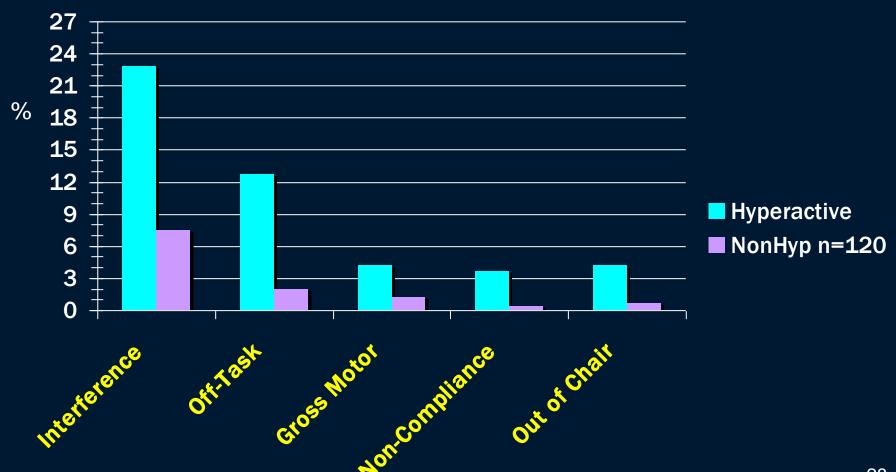
Difficulty completing homework

## Teacher reports of ADHD-like behavior have been controversial

 Are teachers intolerant? It's the teacher's problem, not the child's.

• Understandable, but not likely.....

# "Blind" Observers' Classroom Ratings of Hyperactive Children and Classmates



### Long-Term Course

 A very legitimate concern has been the long-term adjustment of young children diagnosed as having ADHD.

## Longitudinal Study of Boys with ADHD from Age 8 to 41 Years

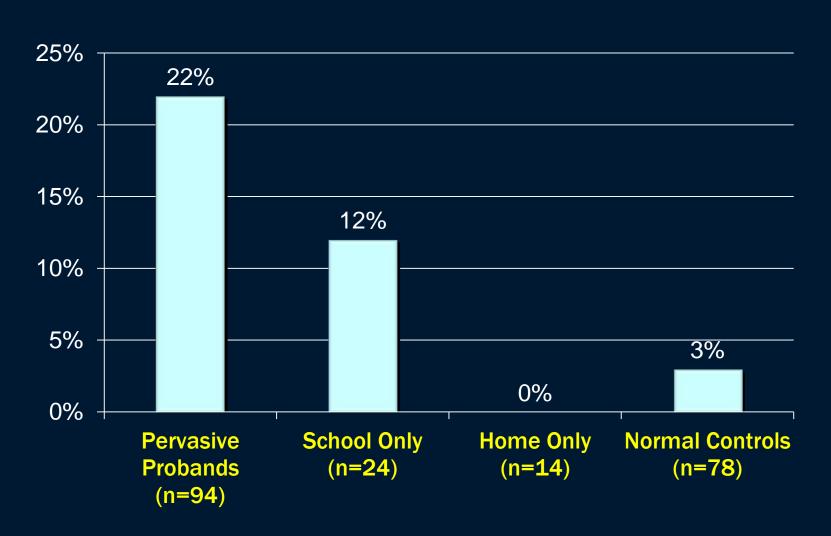
- We diagnosed Combined ADHD in 207
   Caucasian boys, 6 to 12 years (mean, 8).
- They have been followed up 3 times:
  - At age 18 10 years after the original diagnosis
  - At Age 25 17 years after the original diagnosis
  - At age 41 33 years after the original diagnosis (the longest prospective study).

## Not All Children Referred Had Cross-Situational ADHD

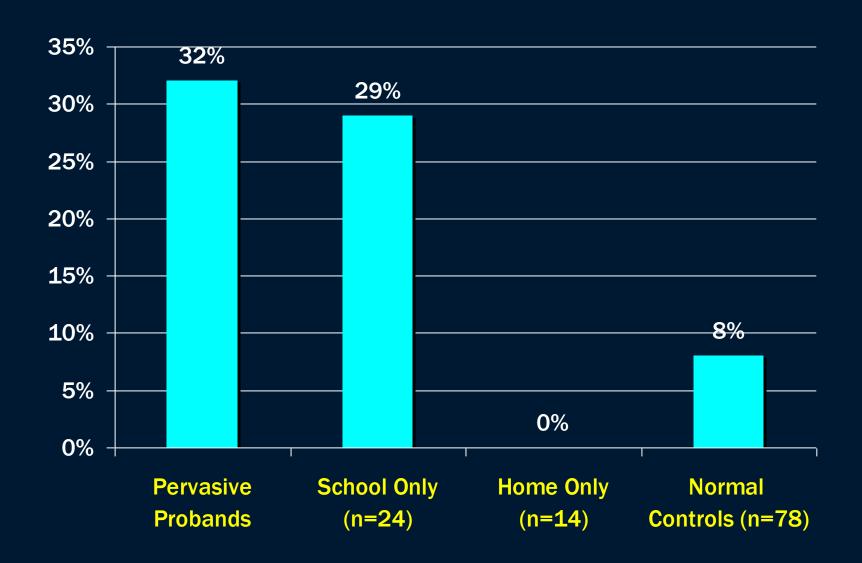
 A number of children were reported to have ADHD only by their parents, and others only by their teachers.

- Did this matter? Yes:
- Outcome was a function of the disorder's pervasiveness.

#### ADHD at Follow-Up - 10 Years Later



### Conduct Disorder at Follow-Up – 10 Years Later



 Two brief descriptions for a flavor of the children we diagnosed and followed up.

### Rob, 6, First Grade

*History:* There have been complaints about Rob's behavior since nursery school (where he fell, "had a concussion because he would not stay still".) The teacher could not control him.

#### **At Referral**

In School: Rob is "uncontrollable", "will not sit still for a minute", and is "disruptive". Teachers have him in isolation and don't allow him into the lunch area.

<u>At Home</u>: Rob is "very active", "constantly moving and talking". "He tries to behave but he says he can't help it."

**During testing:** Rob was in constant motion and had difficulty sustaining attention.

### Francis, 8, Third Grade

*History:* F. "has always been a hyperactive kid, even as an infant". Parents, school, and pediatrician complained about it. In nursery school, he was inattentive and overactive.

#### **At Referral**

**In School:** "He lacks self-control, has a short attention span, is disorganized, forgetful, impulsive and constantly moving; other children are annoyed by his impulsivity".

At home: "He can't seem to sit still, is extremely active, and constantly running and jumping." Doesn't follow directions, must be told several times to do the same thing, he's difficult to discipline.

**During testing:** Restless and somewhat hyperactive.

# Major Findings 10 and 17 Years Later (at ages 18 and 25)

## Compared to Non-ADHD Controls, ADHD Probands

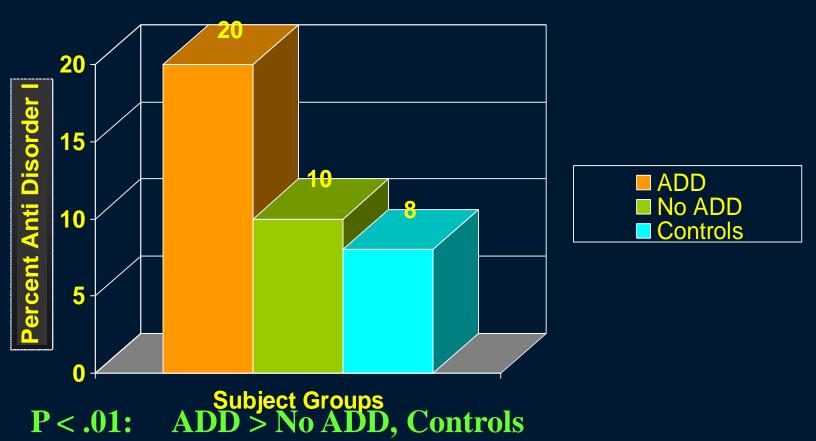
- Had poorer academic performance and completed less schooling (by age 41, 32% had not completed HS, vs. 5% of controls).
- Had poorer social functioning.
- Had lower occupational rankings.

# Only <u>Three disorders</u> were significantly more prevalent in the ADHD group:

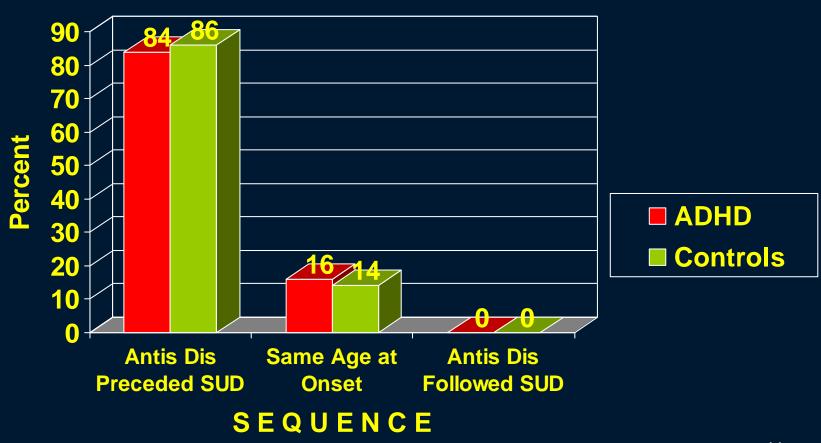
- ADHD
- Antisocial Personality Disorder
- Substance Use Disorders

#### Are these related? YES

### 10 Year later: Antisocial Disorder depended on the Persistence of ADD



### Substance disorders depended on the development of antisocial disorders

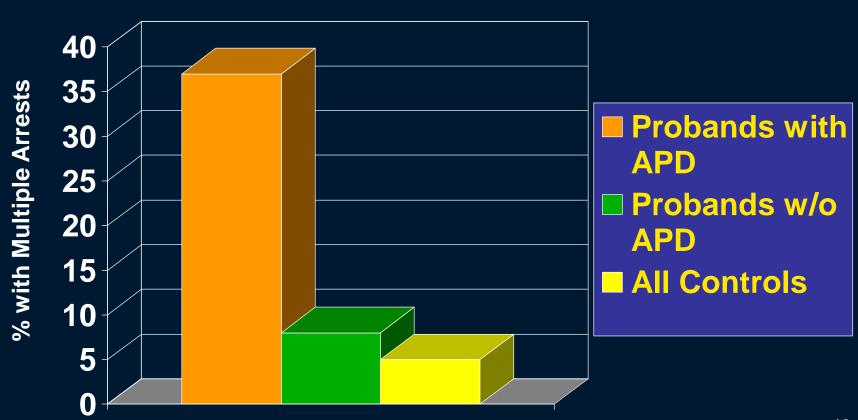


# Is elevated SUD due to greater drug exposure in children with ADHD?

#### NO

77% of ADHD individuals and 75% of Controls had tried drugs.

# Relationship between Antisocial Personality Disorder (APD) and Multiple Arrests



## Developmental Cascade of Psychiatric Disorders

- 1. Childhood ADHD, on to
- 2. Adolescent Antisocial Disorder, on to
- 3. Substance Use Disorder, on to
- 4. Criminality into adulthood

#### How About At Age 41, 33 Years Later?

√ 198 of the 207 boys with ADHD were located and contacted. Of these, 15 (8%) were identified as *Deceased*.

√ 173 of the 178 Male Controls were located and contacted. Of these, 5 (3%) were identified as Deceased.

8% vs. 3%, Chi-Square = 3.97, p = .05

#### Rates (%) of Ongoing Diagnoses -

| DSM-IV Diagnosis                   | ADHD<br>(n = 135) | Controls $(n = 136)$ |
|------------------------------------|-------------------|----------------------|
| ADHD**                             | 16%               | 4%                   |
| Antisocial Personality Disorder*** | 16%               | 0                    |
| Substance Use Disorder             | 22%               | 17%                  |
| Alcohol                            | 10%               | 15%                  |
| Drugs (Cannabis, Cocaine, etc.)**  | 14%               | 5%                   |
| Nicotine Dependence***             | 30%               | 9%                   |

\*p < .05 \*\*p < .01 \*\*\*p < .001

#### N (%) Psychiatrically Hospitalized

| Ever Hospitalized            |                              |     |  |  |
|------------------------------|------------------------------|-----|--|--|
| Probands<br>(n=135)<br>n (%) | Controls<br>(n=136)<br>n (%) | p≤  |  |  |
| 20 (15%)                     | 7 (5%)                       | .01 |  |  |

# Mean Number of Psychiatric Hospitalizations (among those hospitalized)

| ADHD Group |       | Controls  |       |     |
|------------|-------|-----------|-------|-----|
| Mean (SD)  | Range | Mean (SD) | Range | p≤  |
| 3.4 (4.3)  | 1-24  | 1.6 (.9)  | 1-3   | .03 |

# Substance Use Disorders Had Very Negative Consequences.

- They were strong predictors of 1) psychiatric hospitalizations, and
- 2) major depression.

# We know that children with ADHD are at risk for other disorders during adolescence.

How about during adulthood (from age 21 on)?

#### Rates of New Disorders Since Age 21\*

(Mean Age 41)

| DSM-IV Diagnosis            | Probands<br>(n = 135) | Controls<br>(n = 136) | p <                      |  |  |
|-----------------------------|-----------------------|-----------------------|--------------------------|--|--|
| Adjustment Disorder         | 4%                    | 6%                    | NS                       |  |  |
| Substance Use Disorders     |                       |                       |                          |  |  |
| Alcohol                     | 6%                    | 10%                   | NS                       |  |  |
| Non-alcohol                 | 4%                    | 6%                    | NS                       |  |  |
| Any Alcohol or Non-Alcohol  | 4%                    | 10%                   | .03                      |  |  |
| Nicotine                    | 8%                    | 6%                    | NS                       |  |  |
| Mood Disorders              | 30%                   | 22%                   | NS                       |  |  |
| <b>Anxiety Disorders</b>    | 11%                   | 8%                    | NS                       |  |  |
| Other                       | 1%                    | 0%                    | NS                       |  |  |
| Any Disorder Excluding ADHD | 7%                    | 14%                   | <b>.05</b> <sub>51</sub> |  |  |

<sup>\*</sup>Unpublished data

### Contrary to expectation, during adulthood:

The subjects with a childhood history of ADHD did not develop new psychopathology more often than controls.

# The persistence of childhood ADHD into late adolescence was the main cause of negative outcomes.

The period of increased risk for new psychopathology was limited to adolescence.

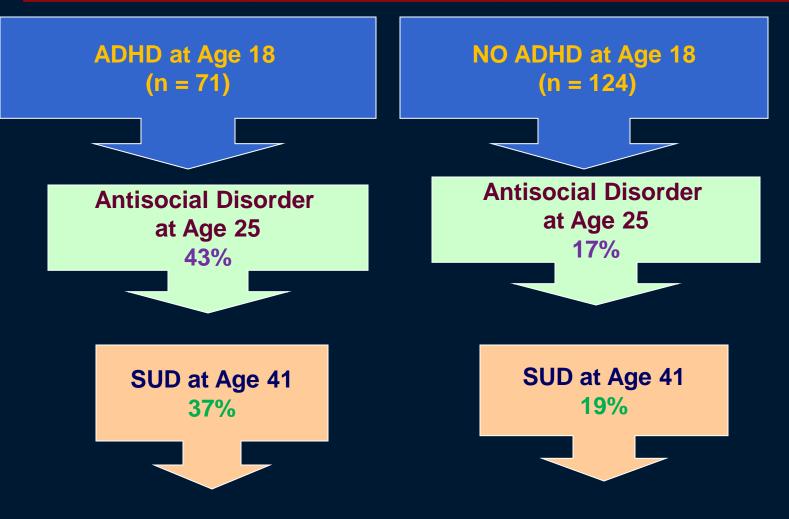
This does not mean that, in adulthood, ADHD children were not worse off than controls. They were.

But their elevated dysfunction in adulthood reflects persistence of malfunction that had its onset in adolescence.

Our findings stress the *importance* of continued monitoring and treatment of children with ADHD, even when conduct disorder is absent when they are first seen.

#### Chronology of ADHD, Antisocial Disorder, and SUD

(Original N = 207 with ongoing ADHD))



- Psychostimulants
  - Amphetamines, levo- and dextro-amphetamine (Benzedrine, Dexedrine, Desoxin)
  - Methylphenidate

#### **History of:**

how they were discovered (1920's) their further development

- short and long-acting
- oral and patch delivery

- > Non-stimulant Medications
  - >Atomoxetine (Strattera)
  - ➤ Bupropion –(Wellbutrin)

They do not have nearly the same efficacy as stimulants and should not be first line treatments (possible exception – SUD).

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#### Studies of Psychosocial Treatments

- Two long-term controlled studies compared multimodal treatment to stimulant medication –
  - 1) MTA\* with duration of 14 months
  - 2) New York/Montreal study of 24 months

<sup>\*</sup> MTA, Multimodal Treatment of ADHD

## MTA Study - 14 Month Outcomes on ADHD Symptoms

 In children with ADHD, age 7-10 years:

Medication was superior to the intensive multimodal behavioral treatment

(14 months with parents, teachers and children)

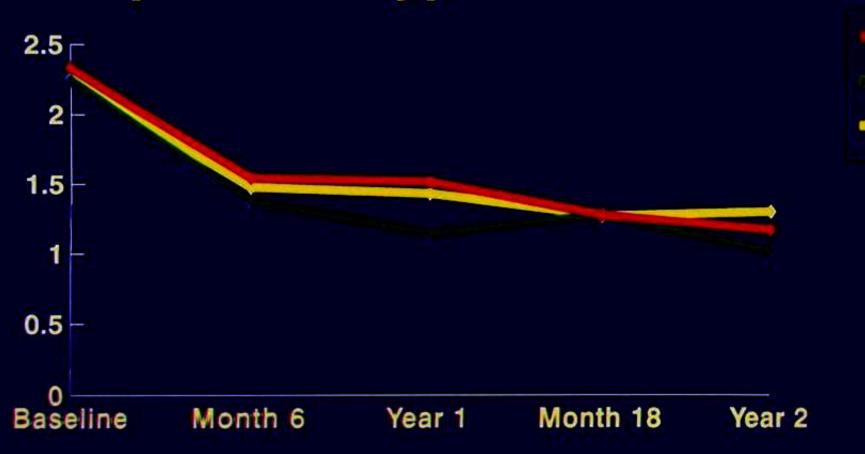
#### **NY/Montreal Study**

## Children, average 8 years, were ALL treated with a stimulant for 2 years:

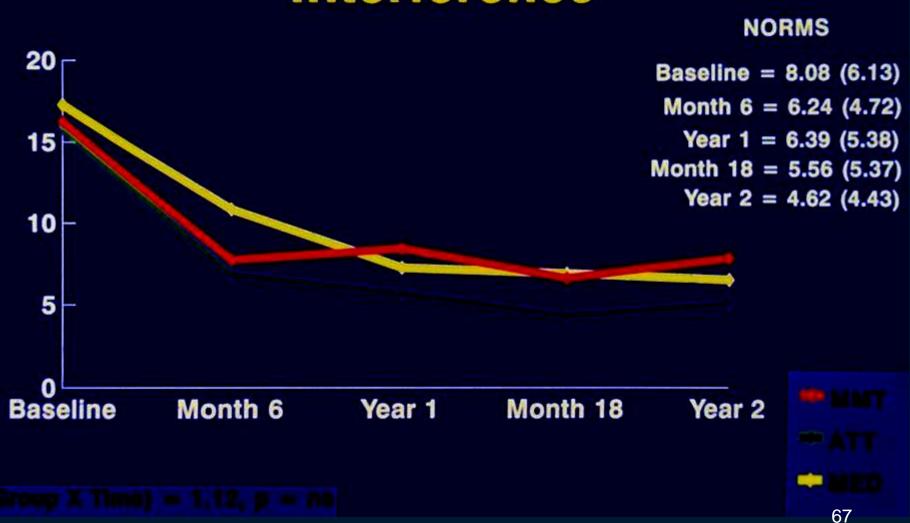
- One third got nothing else.
- One third <u>also</u> received a very active <u>multimodal treatment</u>.
- One third <u>also</u> received a "control", or mock, multimodal treatment.

All for 2 years

#### Conners Parent Scale: Impulsive-Hyperactive Factor



## Observed Levels of Classroom Interference



#### **Treatment Considerations**

**Key Implications for Parents and Practitioners:** 

Continued treatment, with adequate doses of medication is essential

Combined treatments may be desired by parents, and may help them cope, but they do not affect the child's ADHD symptoms