

Publicaciones Relevantes

Sobre el TDAH

2008 - 2012



Hospital Universitario
Ramón y Cajal

Comunidad de Madrid



Fundación para una infancia y adolescencia saludables

Journal of the American Academy of
CHILD & ADOLESCENT PSYCHIATRY



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Advancing the science of pediatric mental health and promoting the care of youth and their families

Andrés Martín, Editor
Madrid, Enero 28, 2012



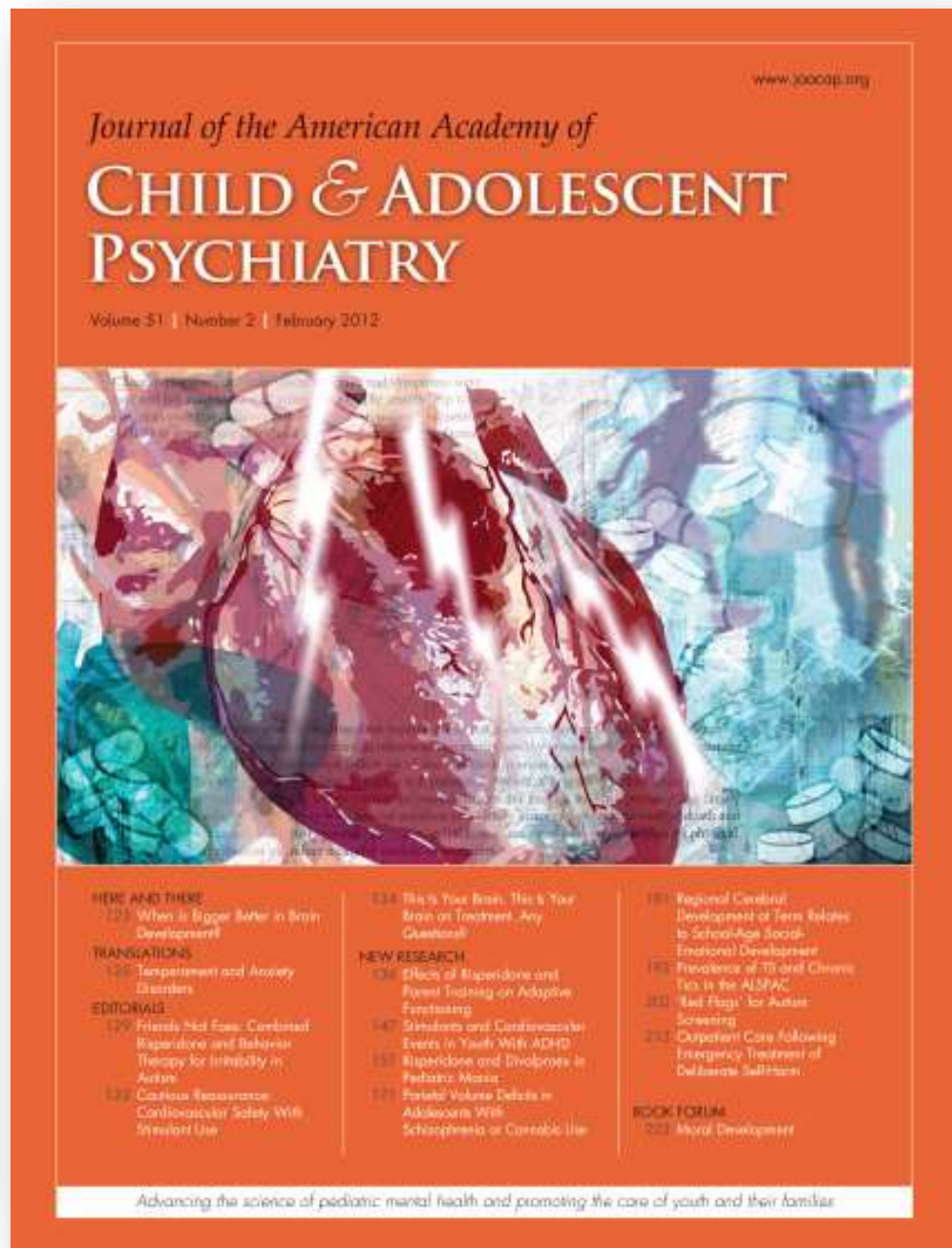
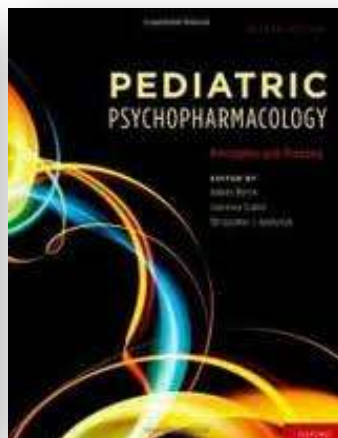
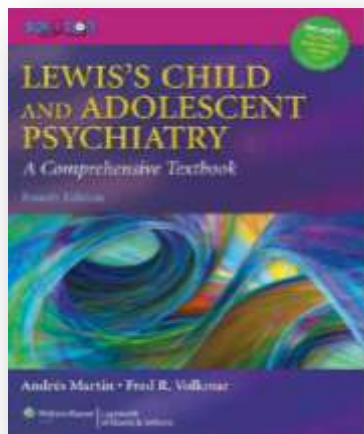
Yale Child Study Center

FROM GENERATION
TO GENERATION



Disclosures

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- Royalties from books
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- Travel, lodging, honorarium from CONFIAS



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Journal of the American Academy of

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EDICIÓN ESPAÑOLA

Volumen 4 | Número 1 | Junio 2010 | Páginas 1-51

PSYCHIATRY

EDICIÓN ESPAÑOLA



COMENTARIO A LA VERSIÓN ESPAÑOLA

De Estados Unidos a España
y de regreso: viajes amigos
para nuevos proyectos

EDITORIAL

Continúa el status quo: revisión
de los criterios diagnósticos
del TDAH

NUÉVA INVESTIGACIÓN

Trastornos psiquiátricos en
niños extremadamente
premáturos: resultados
longitudinales a la edad

de 11 años en el estudio
EPICure

Consecuencias de la
ampliación del criterio de edad
de inicio del TDAH hasta
los 12 años: Resultados de
un estudio prospectivo
de una cohorte de nacimiento

Diferencias en los síntomas
y diagnósticos del trastorno
por déficit de atención/
hiperactividad según el sexo
y la edad: consecuencias
para el DSM-V y la ODS-11

Más allá del modelo de la
doble vía: prueba indicativa
de la disociación de los
efectos del procesamiento
temporal, la inhibición y la
aversión a la espera en el
trastorno por déficit de
atención/hiperactividad

Esta publicación ha sido patrocinada por Janssen-Cilag

Advancing the science of pediatric mental health and promoting the care of youth and their families

COMENTARIO A LA VERSIÓN ESPAÑOLA

De Estados Unidos a España y de regreso: viejos amigos para nuevos proyectos

Apreciado compañero,

Llega a tus manos, con un cierto inevitable re-

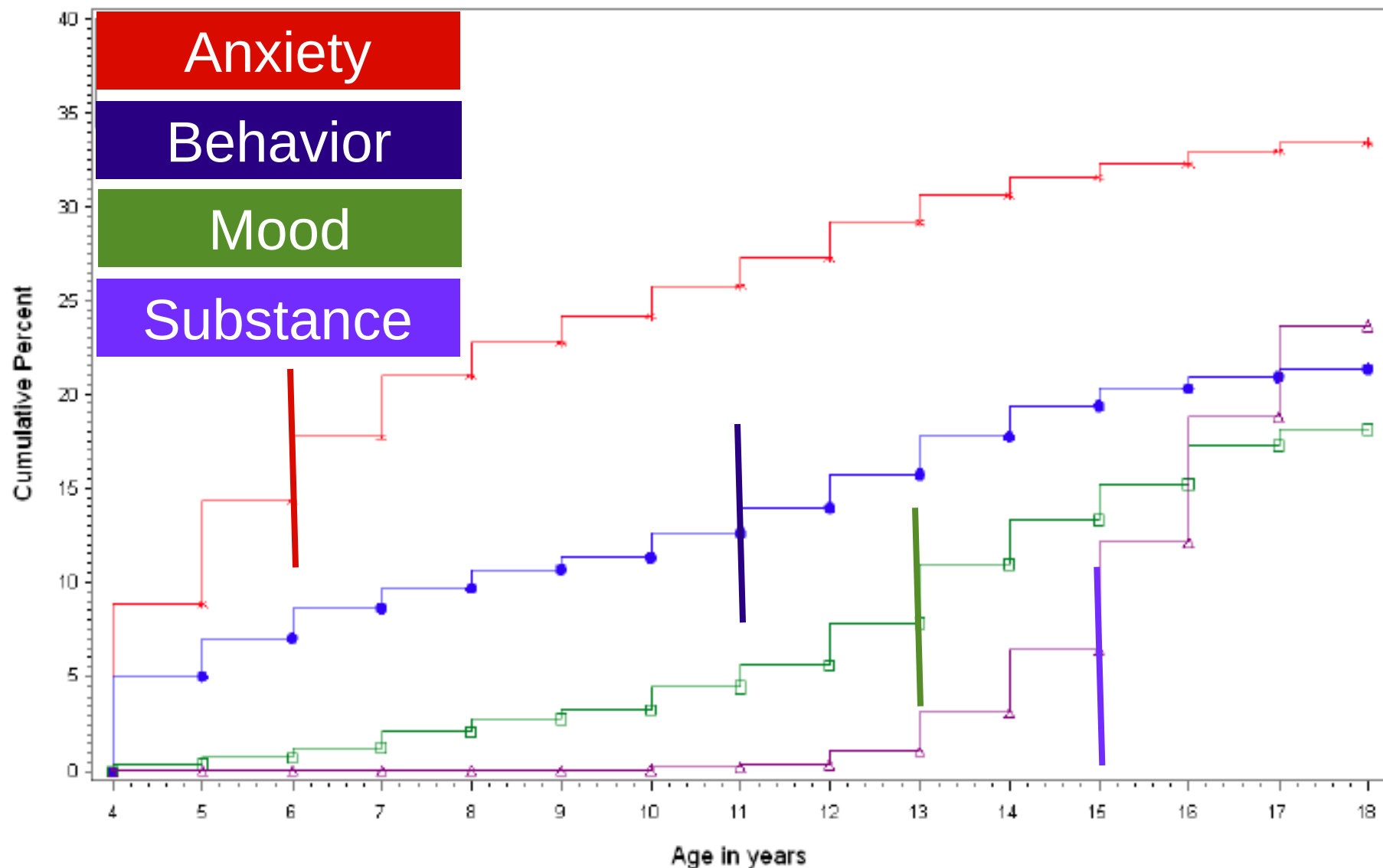
ja portada se colorea de novísimos “*covert arts*”, también de hispánico trazo (Socorro Rivera, que, como Andrés, proviene de la Ciudad de México),

PSYCHIATRY
EDICIÓN ESPAÑOLA

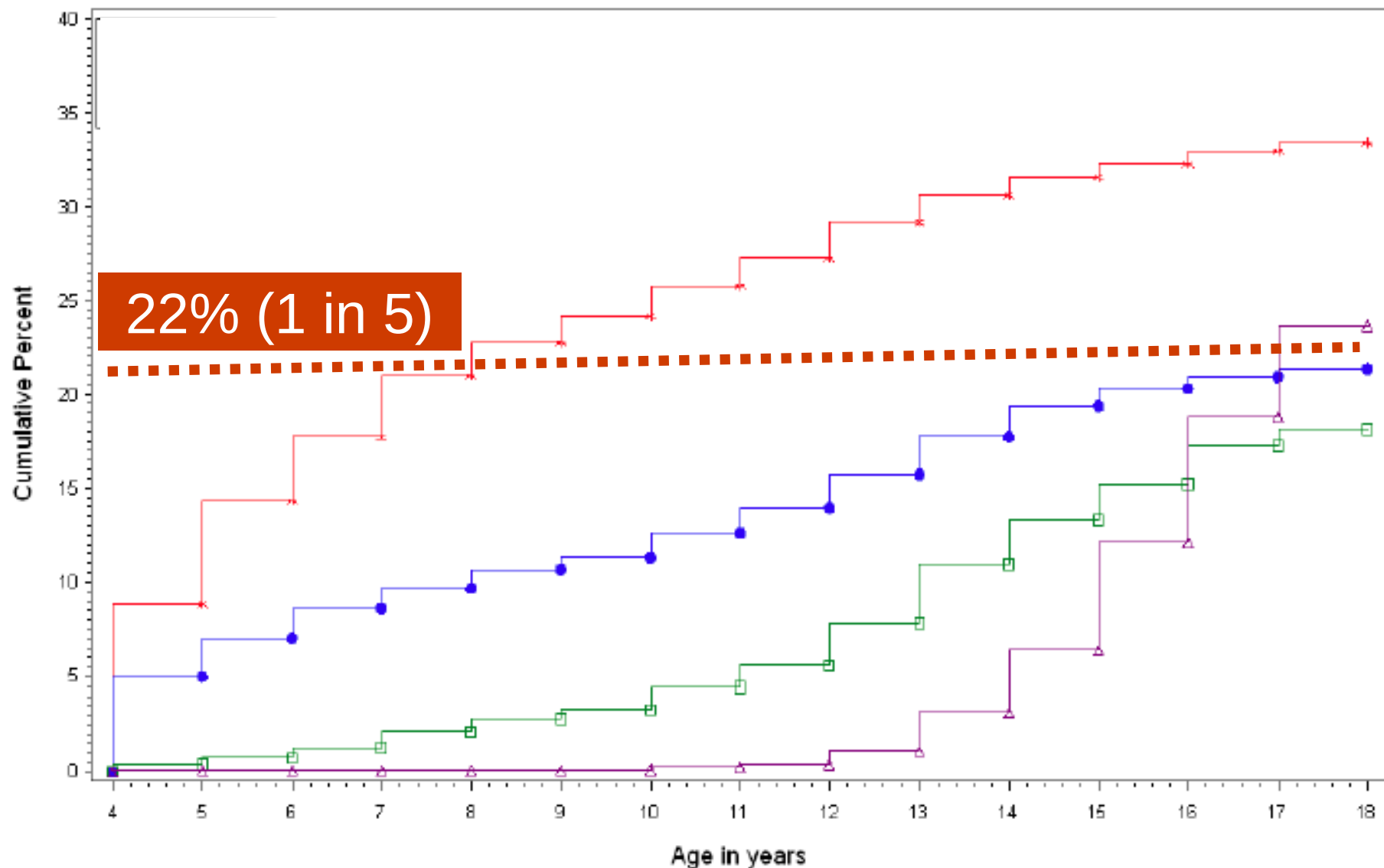


Con gracias a
Xavier Gastaminza
& Óscar Herreros!

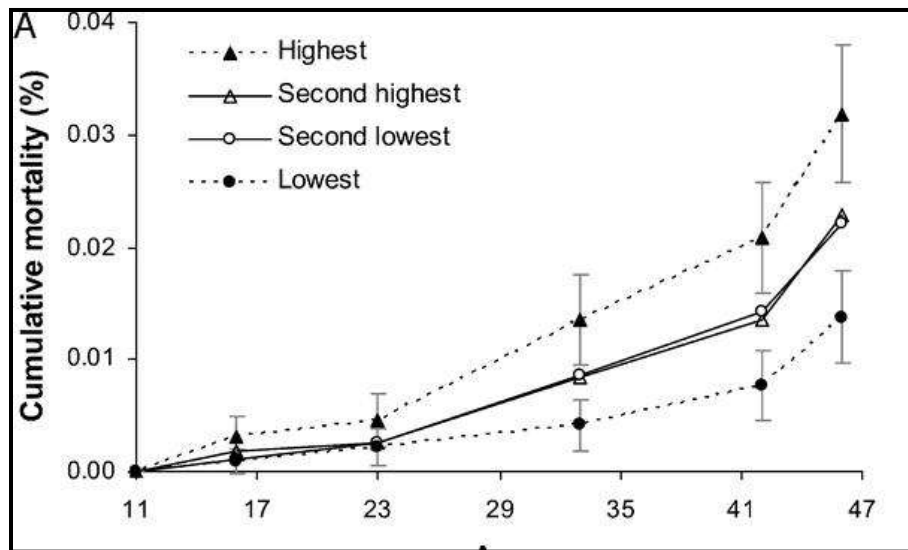




Lifetime prevalence of mental disorders in US adolescents: results from the National Comorbidity Study-Adolescent Supplement. Merikangas *et al* JAACAP 10.2010



Lifetime prevalence of mental disorders in US adolescents: results from the National Comorbidity Study-Adolescent Supplement. Merikangas *et al* JAACAP 10.2010



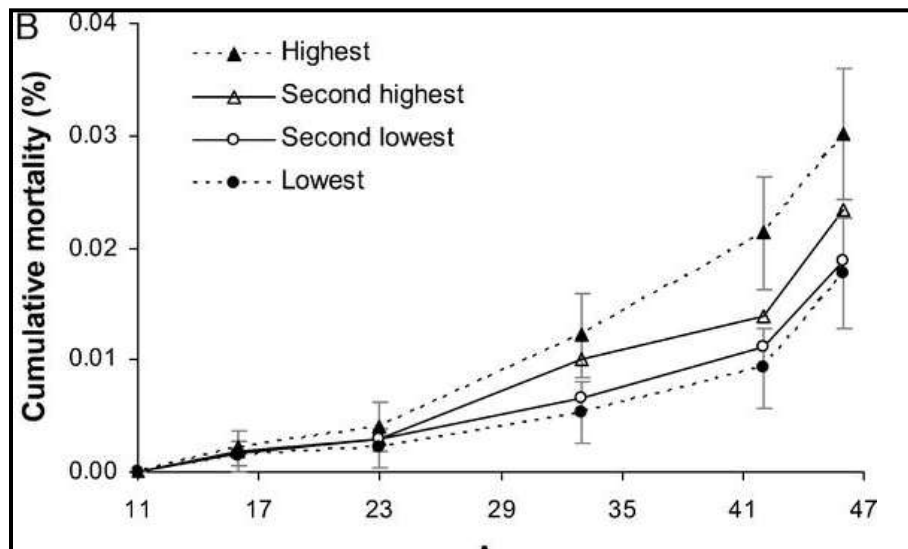
Externalizing
OR = 1.27 (1.13-1.44)

British Child Development Study
1958 Birth Cohort

N = 11,142

Deaths = 243

Follow up = 35 years



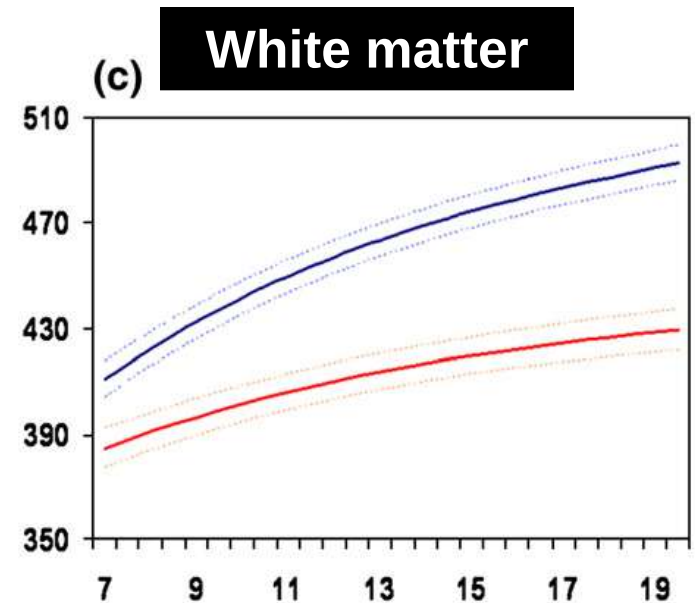
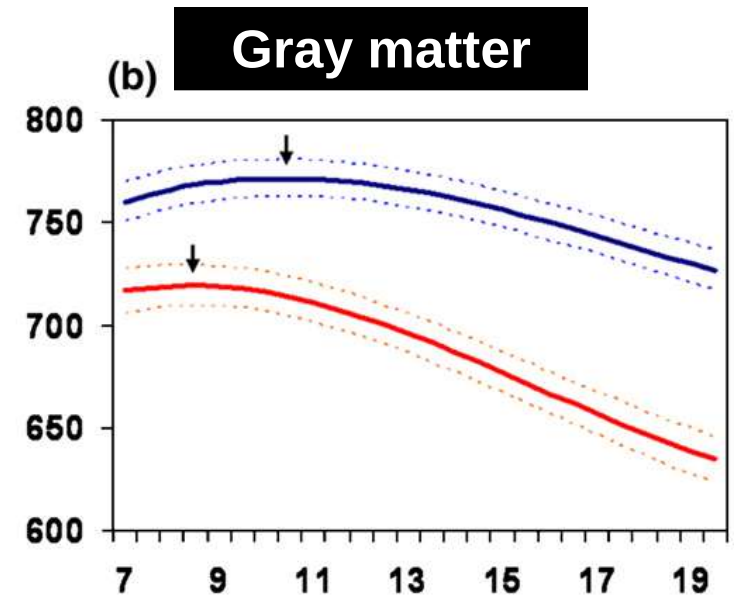
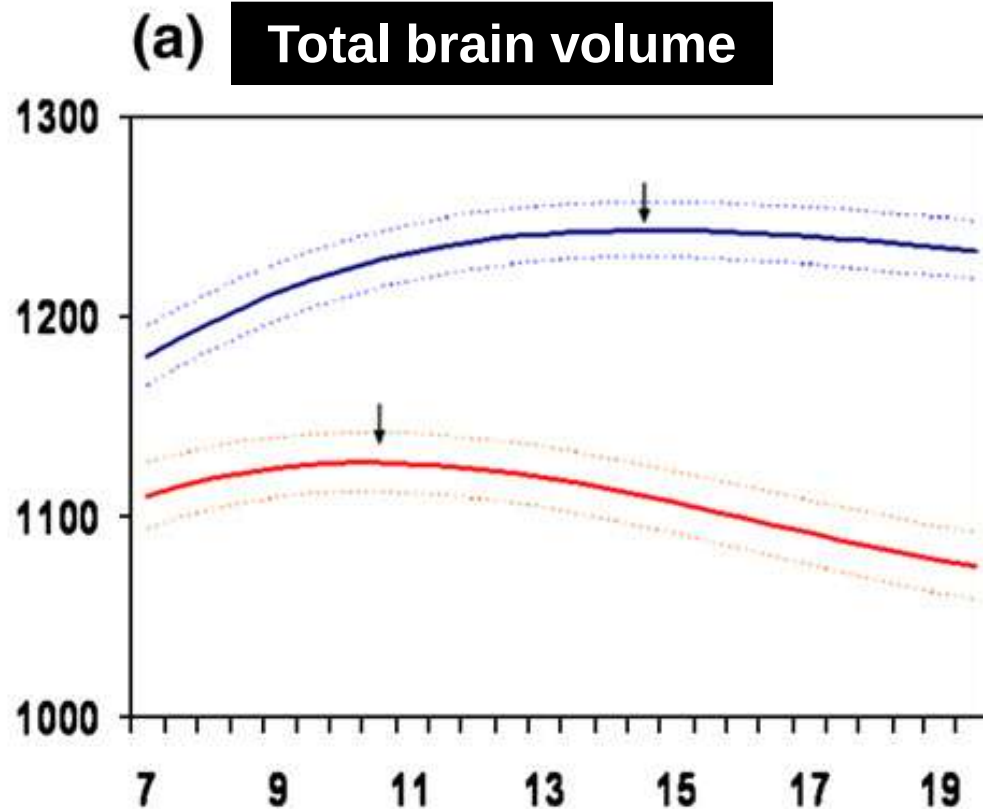
Internalizing
OR = 1.21 (1.06-1.37)



Childhood Problem Behaviors and Death by Midlife:
 The British National Child Development Study. Jokela *et al* JAACAP 1.2009

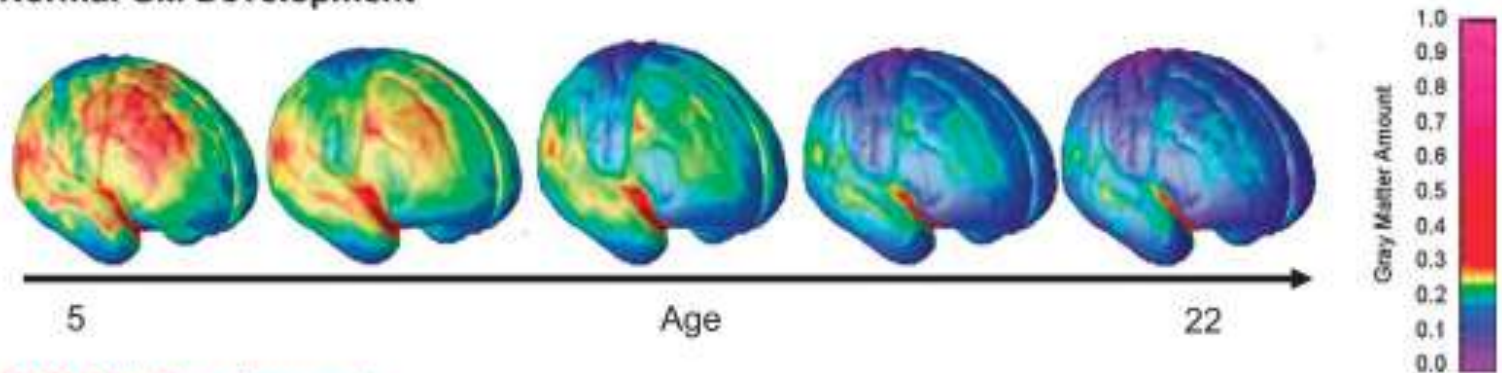


Sexual dimorphism

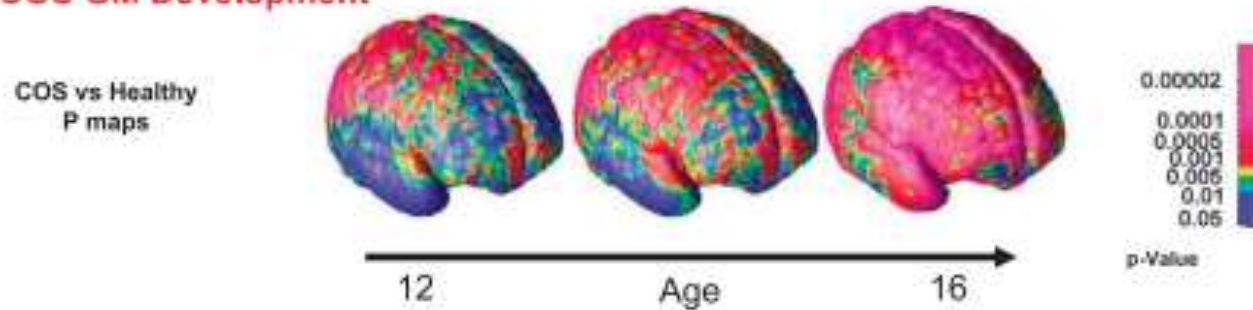


Sexual dimorphism of brain development. Lenroot *et al Neuroimage* 2007
(Anatomical MRI of typically developing children. Giedd *et al JAACAP* 5.2009)

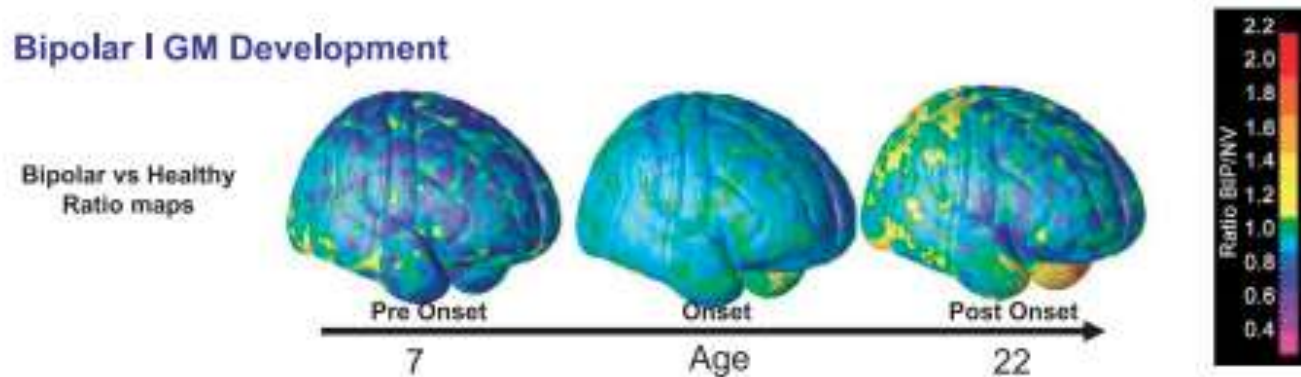
Normal GM Development

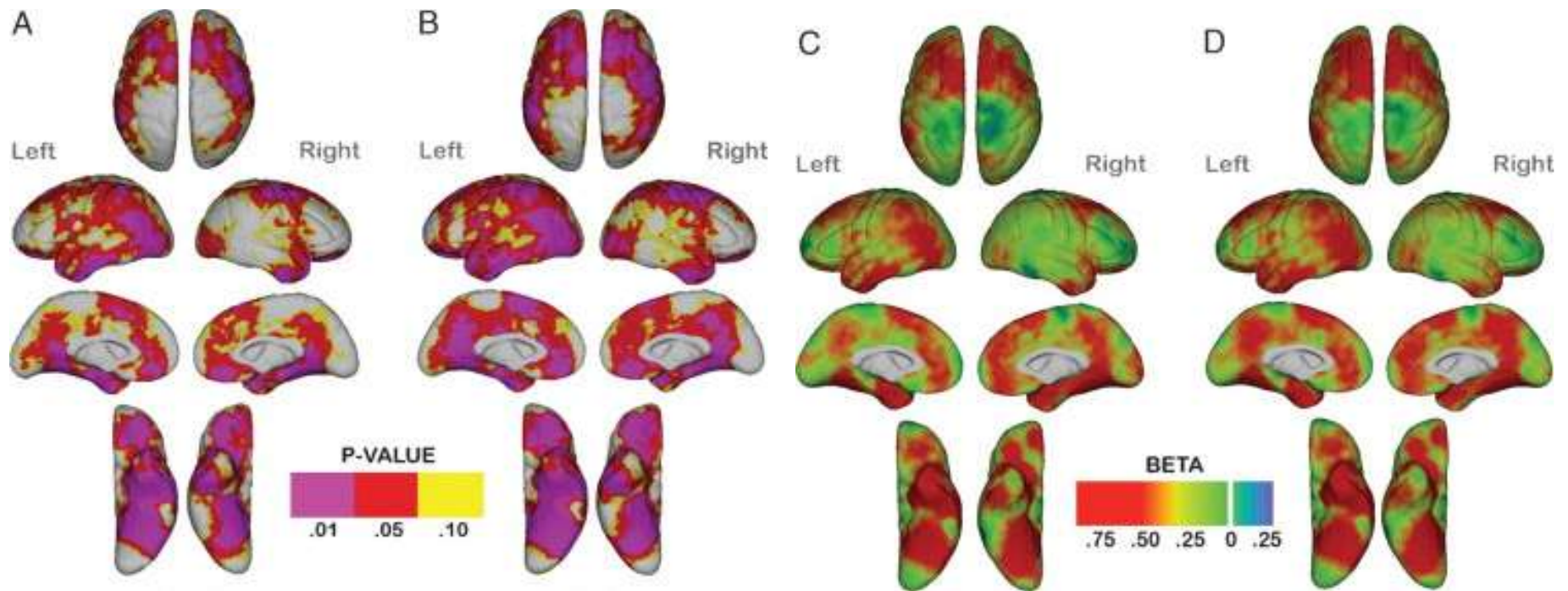


COS GM Development

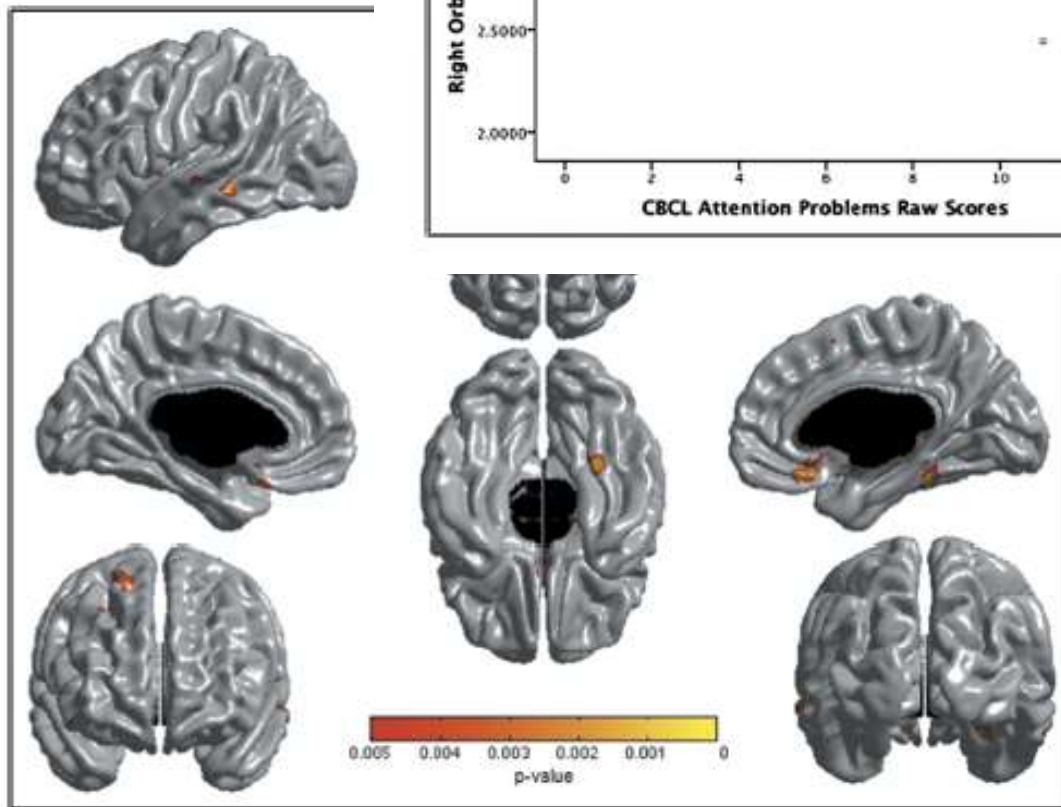
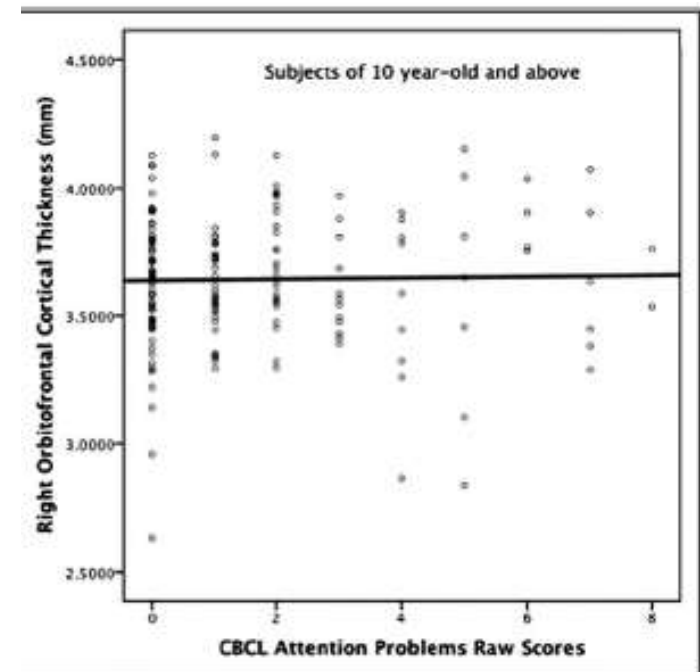
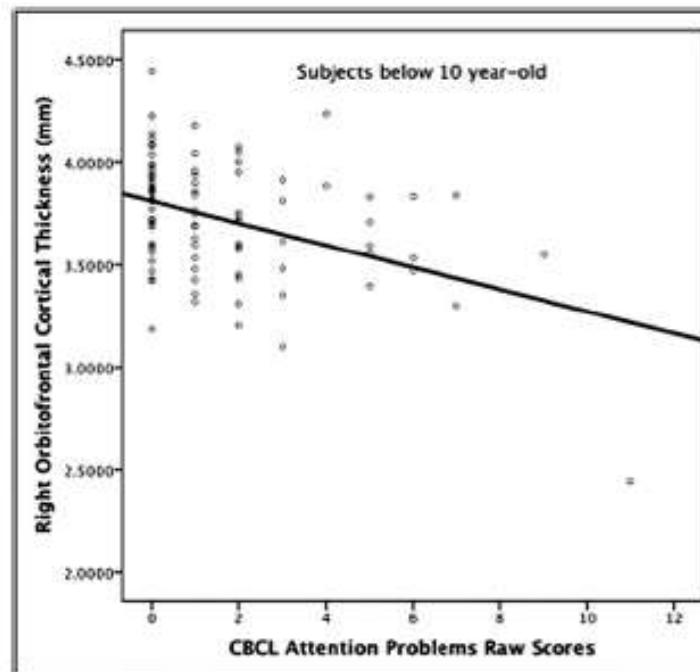


Bipolar I GM Development





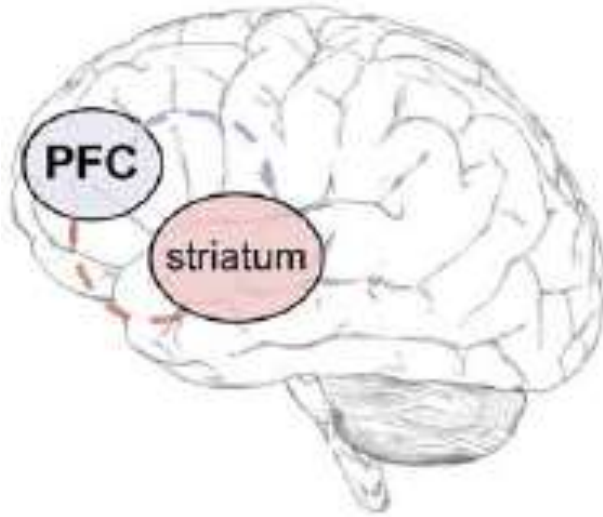
Larr et al 10.09
Widespread Cortical Thinning Is a Robust Anatomical Marker for ADHD



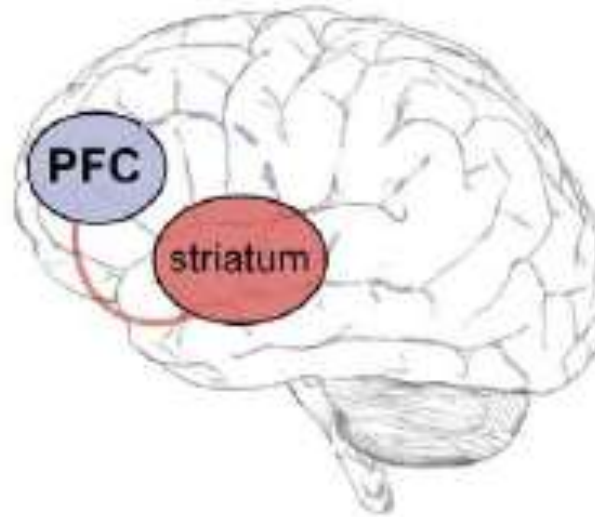
Ducharme et al 1.12

Decreased Regional Cortical Thickness and Thinning Rate
Are Associated with Inattention Symptoms in Healthy Children

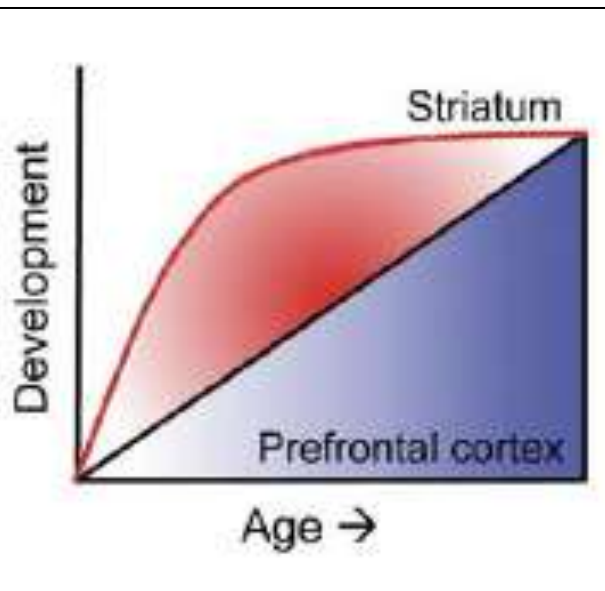
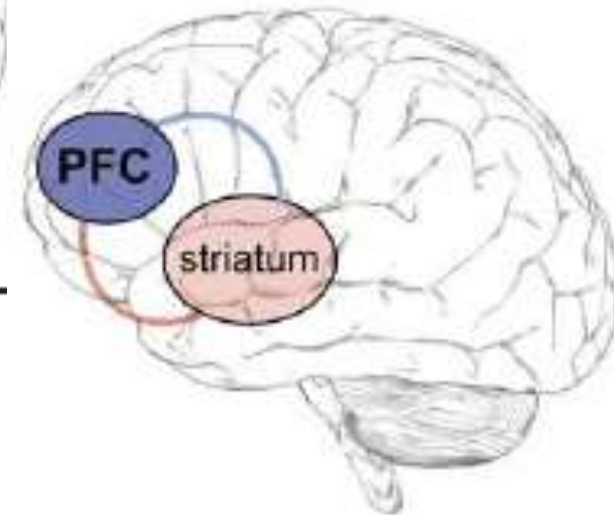
Children



Adolescents



Adults

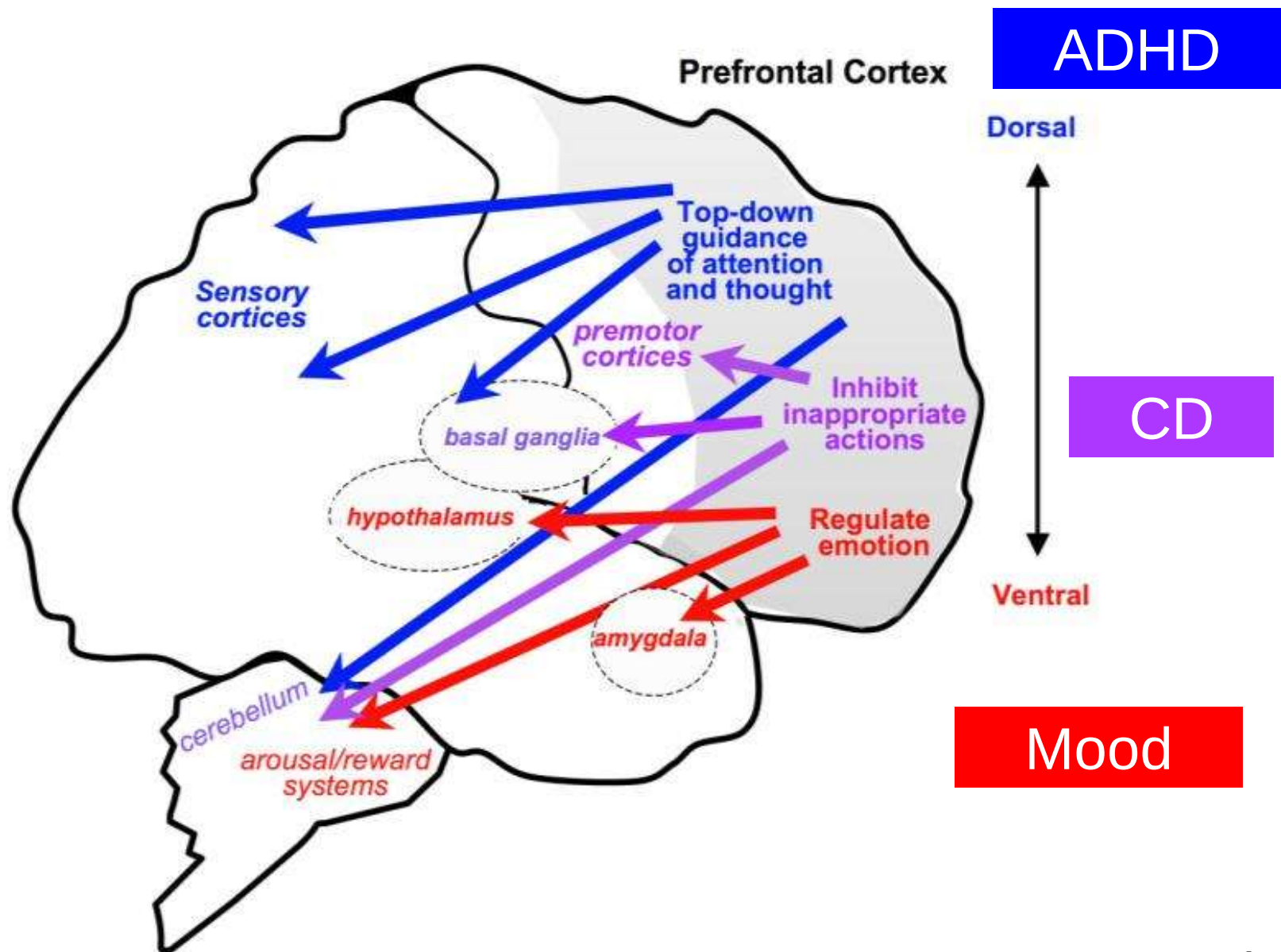


Casey et al 10.10
Neurobiology of the Adolescent Brain and Behavior:
Implications for Substance Abuse Disorders



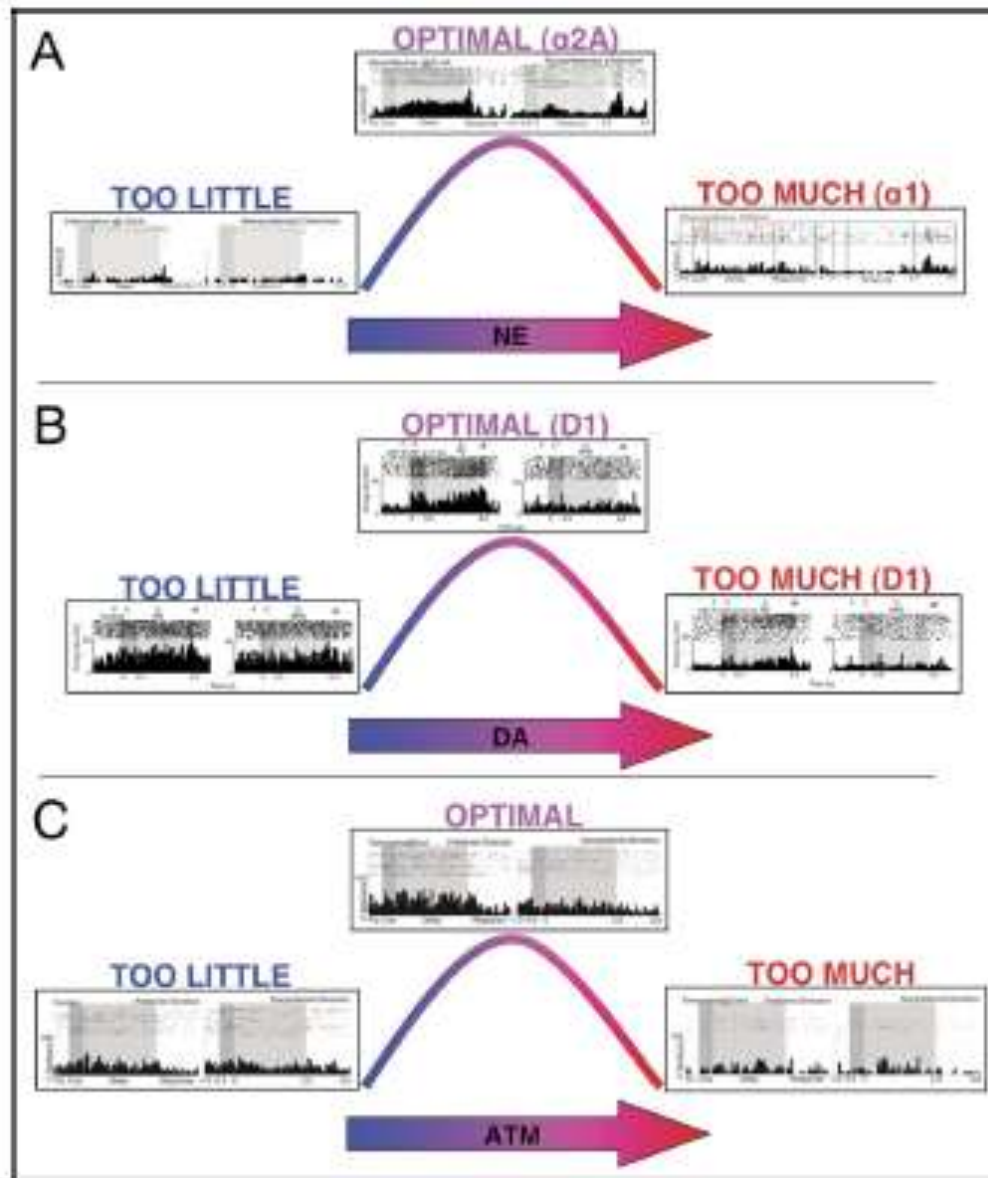
Hechtman 6.11

Prospective Follow-up Studies of ADHD: Helping Establish a Valid Diagnosis in Adults

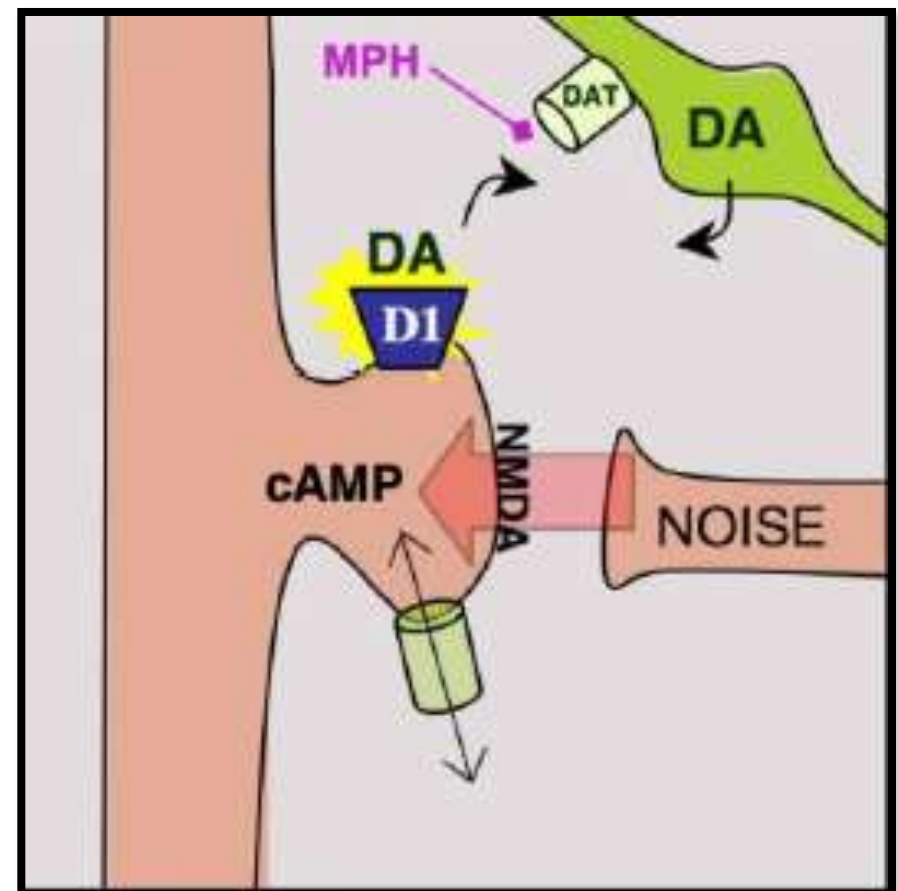
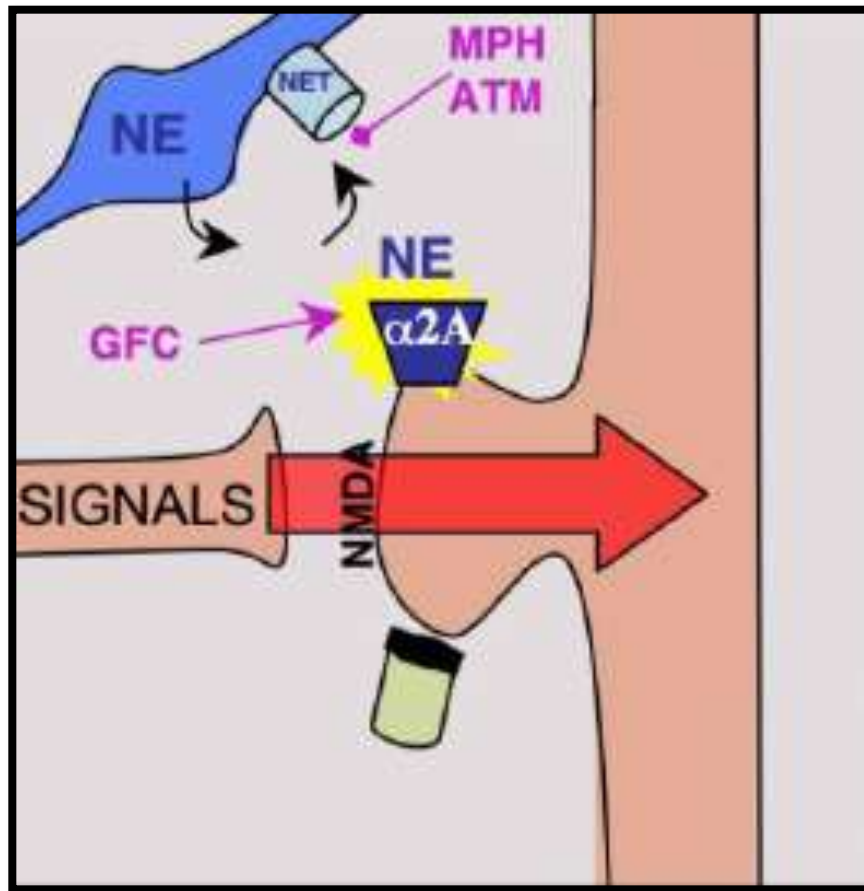


Arnsten et al 4.12

Neurobiological Circuits Regulating Attention, Cognitive control, Motivation and Emotion:
Disruptions in Neurodevelopmental Psychiatric Disorders



Arnsten et al 10.10
Methylphenidate and Atomoxetine Enhance Prefrontal Function
Through α_2 -Adrenergic and Dopamine D₁Receptors



Arnsten et al 4.12

Neurobiological Circuits Regulating Attention, Cognitive control, Motivation and Emotion:
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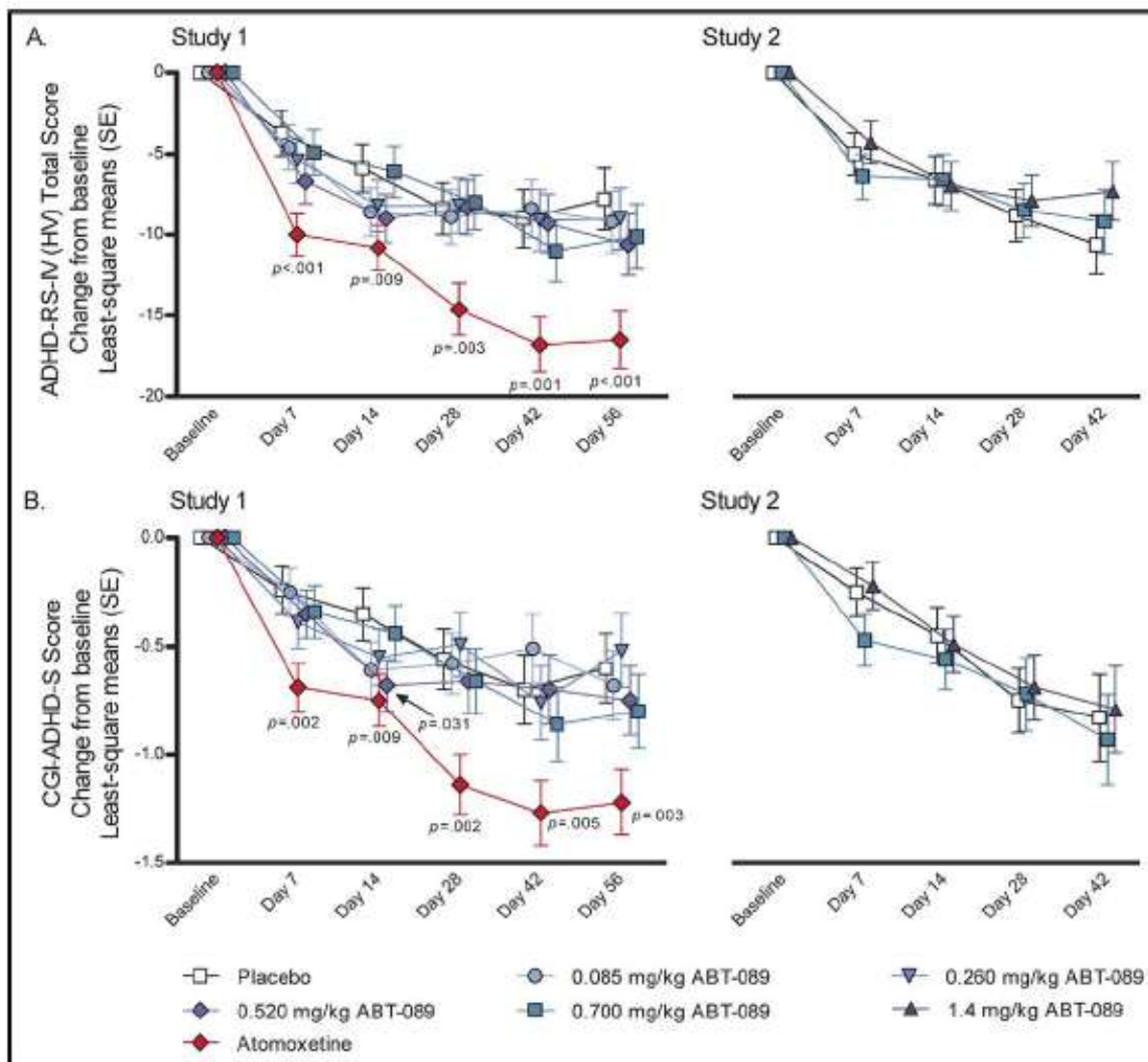
PHARMACEUTICALS

Novartis to shut brain research facility

Drug giant redirects psychiatric efforts to genetics.

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psy- the l
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has vard
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with nold
not- both
sequencing capacity and a la

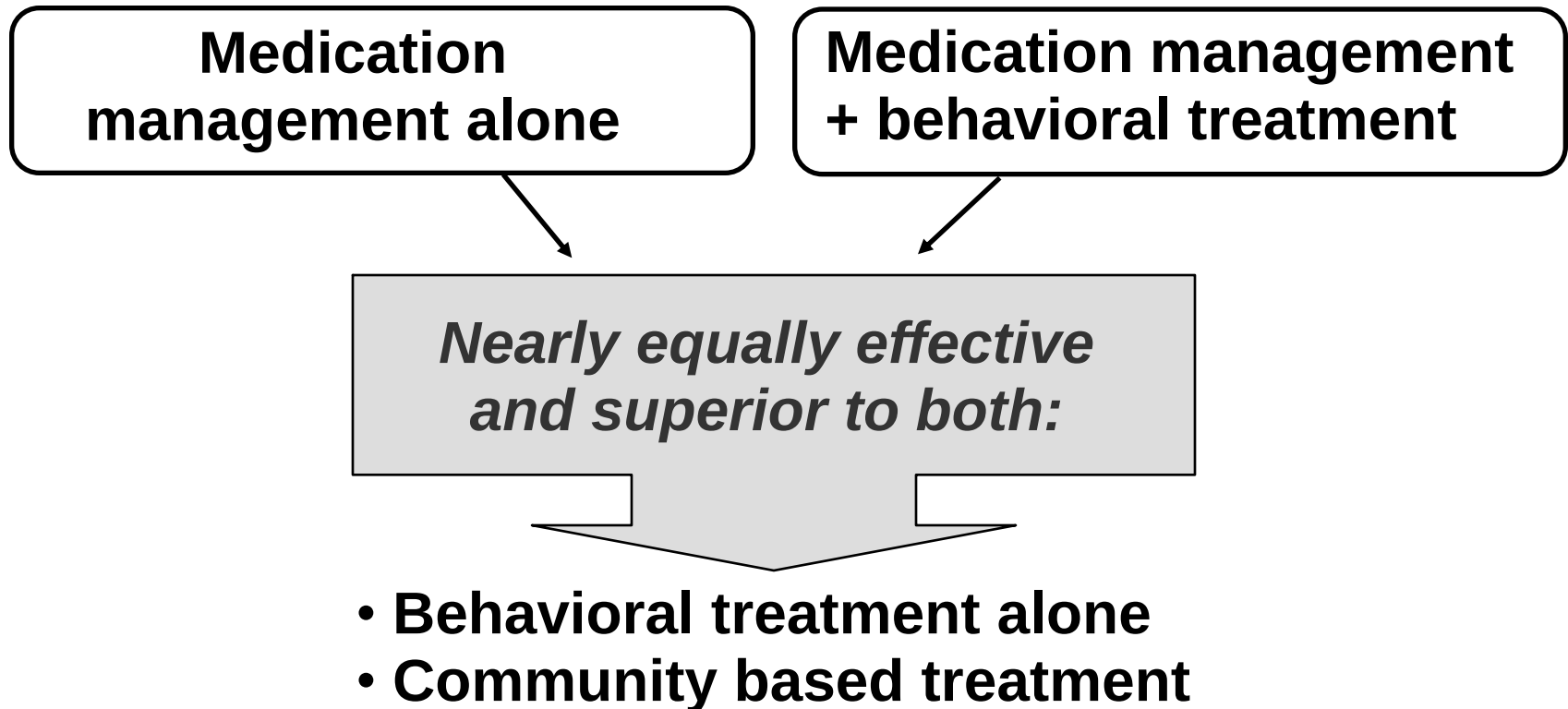
“Developing drugs for the brain has become high-risk, with most candidates failing.”



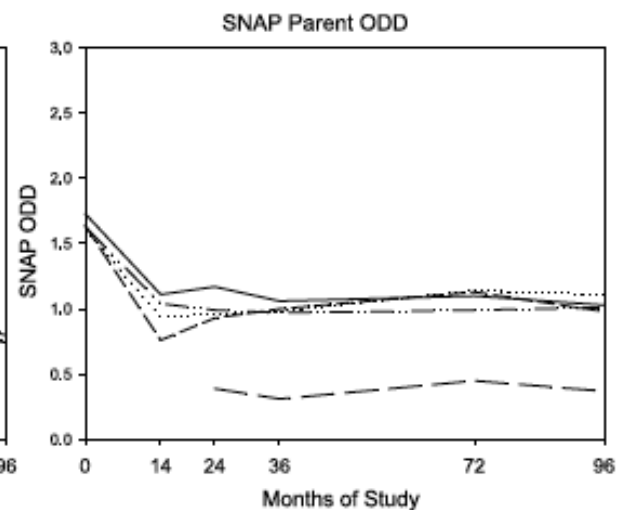
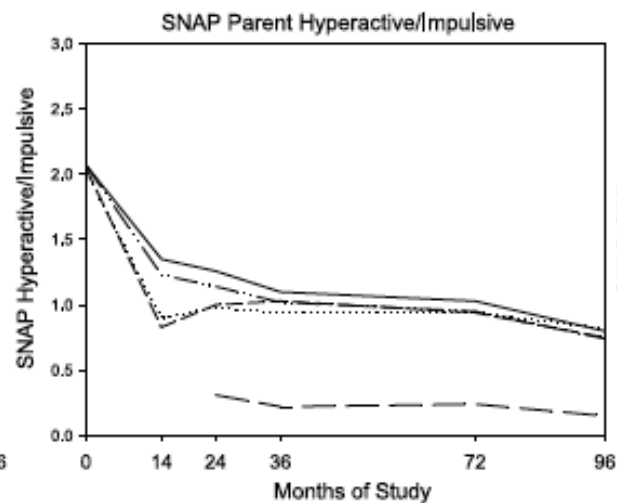
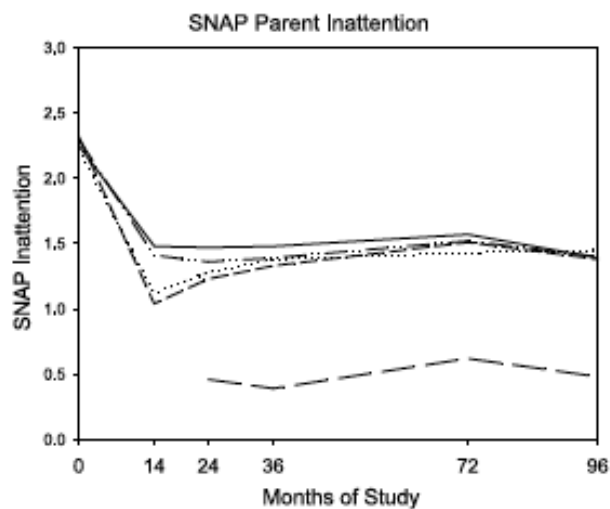
Wilens et al 1.11
Safety and Efficacy of ABT-089 in Pediatric ADHD:
Results from Two Randomized Placebo-Controlled Clinical Trials

ADHD: MTA Results

All treatment arms found to be effective on an absolute basis



(MTA Study Group, Arch Gen Psych, 1999)



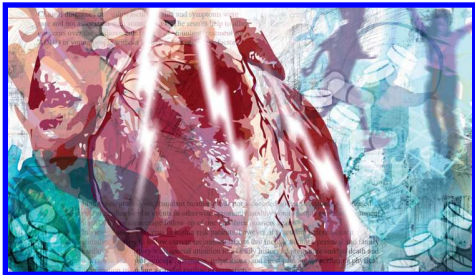
Molina et al 5.09
The MTA at 8 Years: Prospective Follow-up of Children
Treated for Combined-Type ADHD in a Multisite Study

Clinical diagnoses of myocarditis, events and symptoms were rare and not associated with stimulant use. The results help to allay concerns over the downsides of stimulant treatment for ADHD in young people without identifying additional risk factors.

In the present analysis, stimulant treatment was not associated with a substantial increased risk of sudden cardiac events in otherwise apparently healthy young people. The mean duration of follow-up was 1.5 years, with an average follow-up of one and three quarters in view of the high attrition rates. The results are reassuring in higher risk patients, however, it is prudent to follow stimulant treatment in line with current recommendations that include taking a personal and family cardiovascular history with special attention to a family history of premature sudden death and a personal history of syncope, seizures, palpitations, and chest pain, and performing a physical examination including a careful cardiac examination.

Stimulants and cardiovascular risk - 1

- Pharmacoepidemiology study
- Used pharmacy claims
- N=171,126; Ages 6-21
- Outcomes:
 - Cardiac events (eg ED, angina)
 - Cardiac symptoms (eg tachycardia)



Olfson et al 2.12
Stimulants and Cardiovascular Events in Youth with ADHD

Stimulants and cardiovascular risk - 2

- Incidence on stimulants (per million days of use)
 - Events: 0.92
 - Symptoms 3.08
- Relative Risk (vs no use)
 - Events: 0.69 (0.42-1.12)
 - Symptoms: 1.18 (0.83-1.66)



Olfson et al 2.12
Stimulants and Cardiovascular Events in Youth with ADHD

Cardiovascular monitoring



- Before treatment
 - Review personal history
 - Family history, eg sudden / unexpected early deaths
 - HR and BP (per gender / age / height norms)
 - ECG not routinely mandatory
- During treatment
 - Monitor for symptoms
 - Refer for symptoms, pre HTN (90-95 PP), or HTN (>95)
 - Seek consultation as indicated

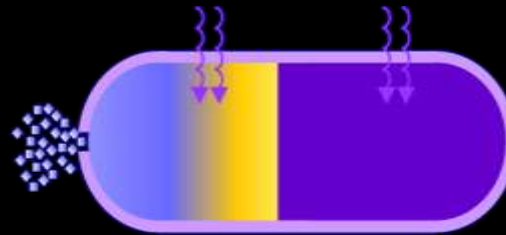
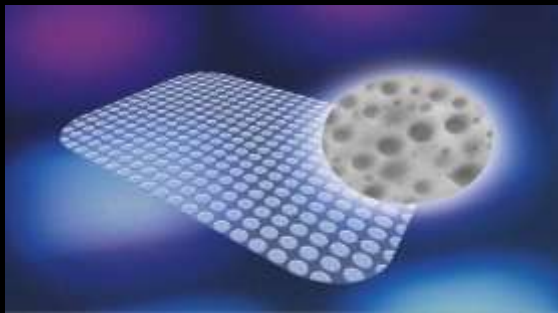


Pumps

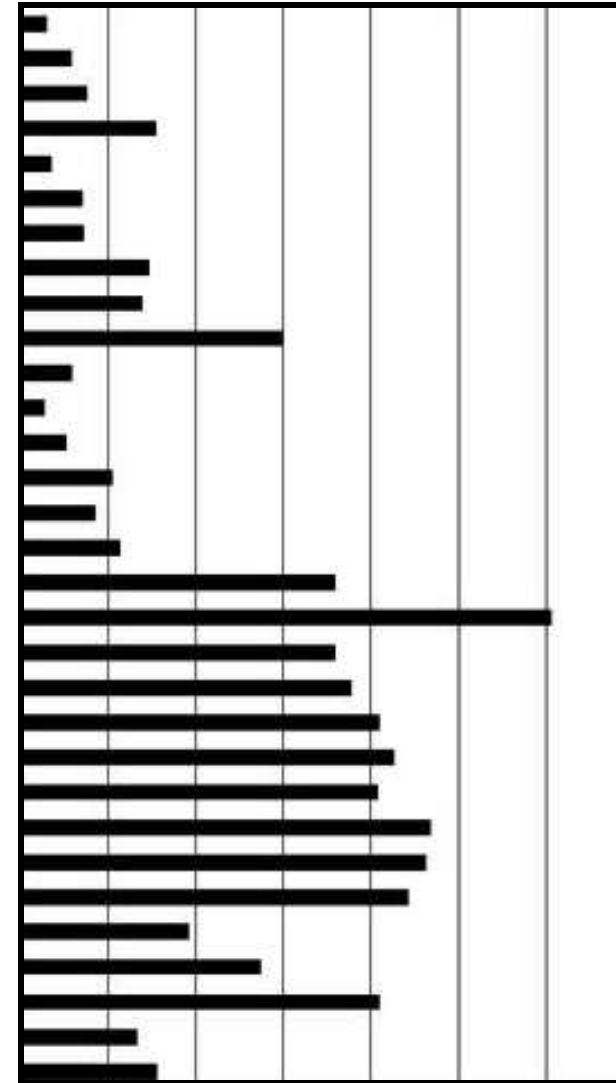
Prodrugs

Pearls

Patches

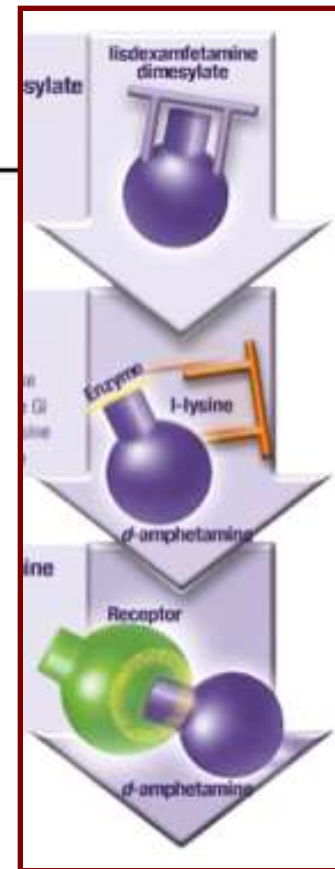
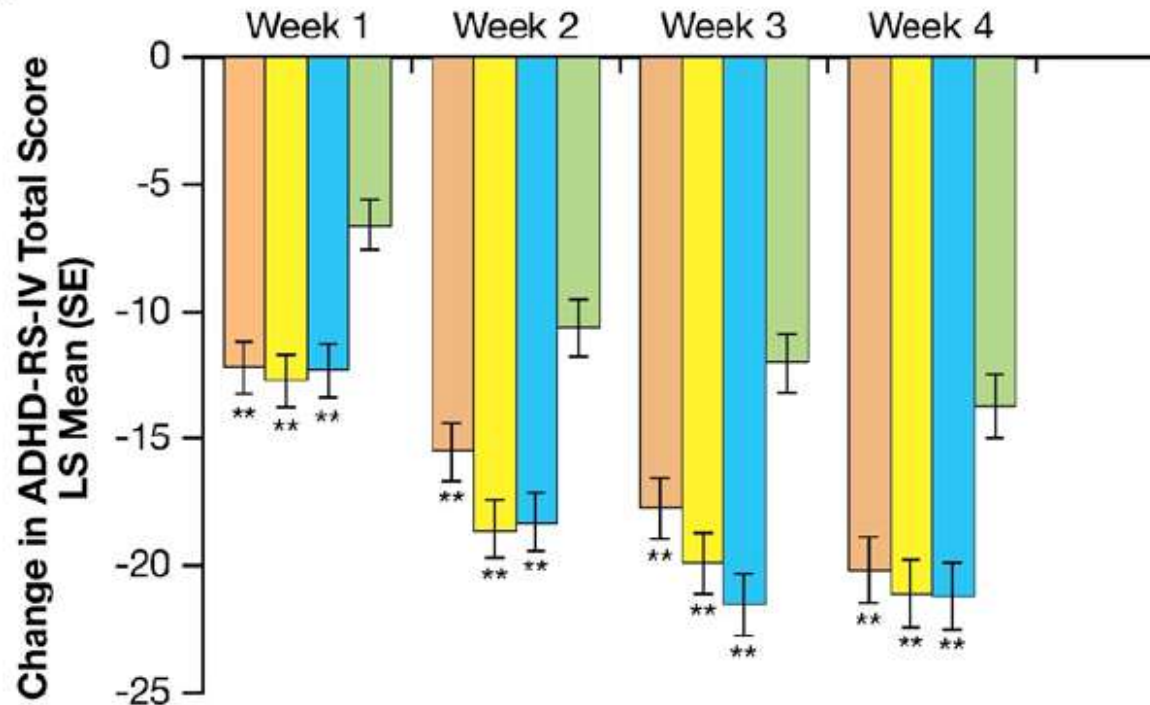


\$400



LDX				
30 mg/d n = 78	50 mg/d n = 77	70 mg/d n = 78	All Doses n = 233	Placebo n = 77

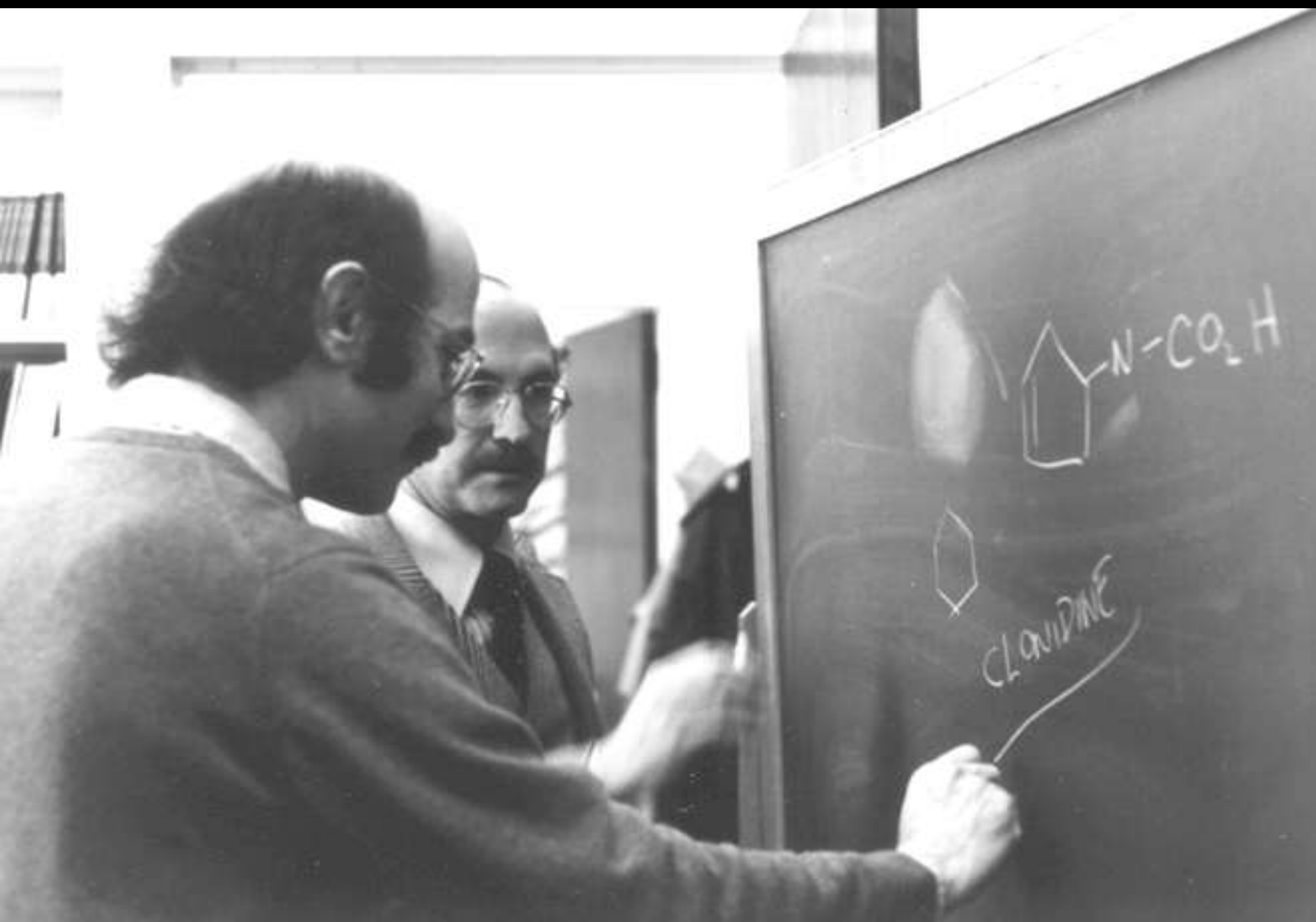
b



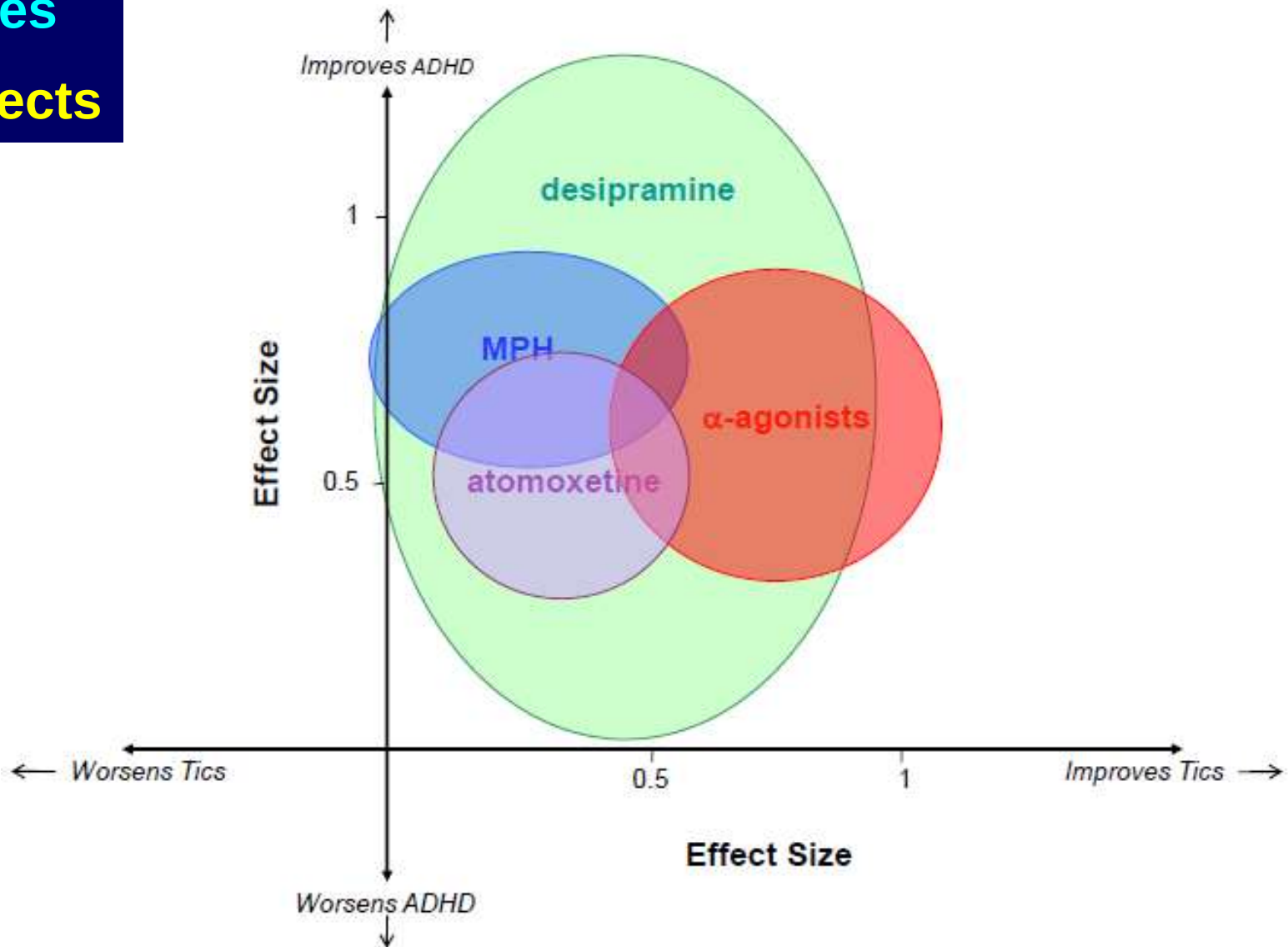
- LDX (30 mg/d)
- LDX (50 mg/d)
- LDX (70 mg/d)
- Placebo

Findling et al 4.11

Efficacy and Safety of Lisdexamfetamine Dimesylate in Adolescents with ADHD



9 studies
477 subjects

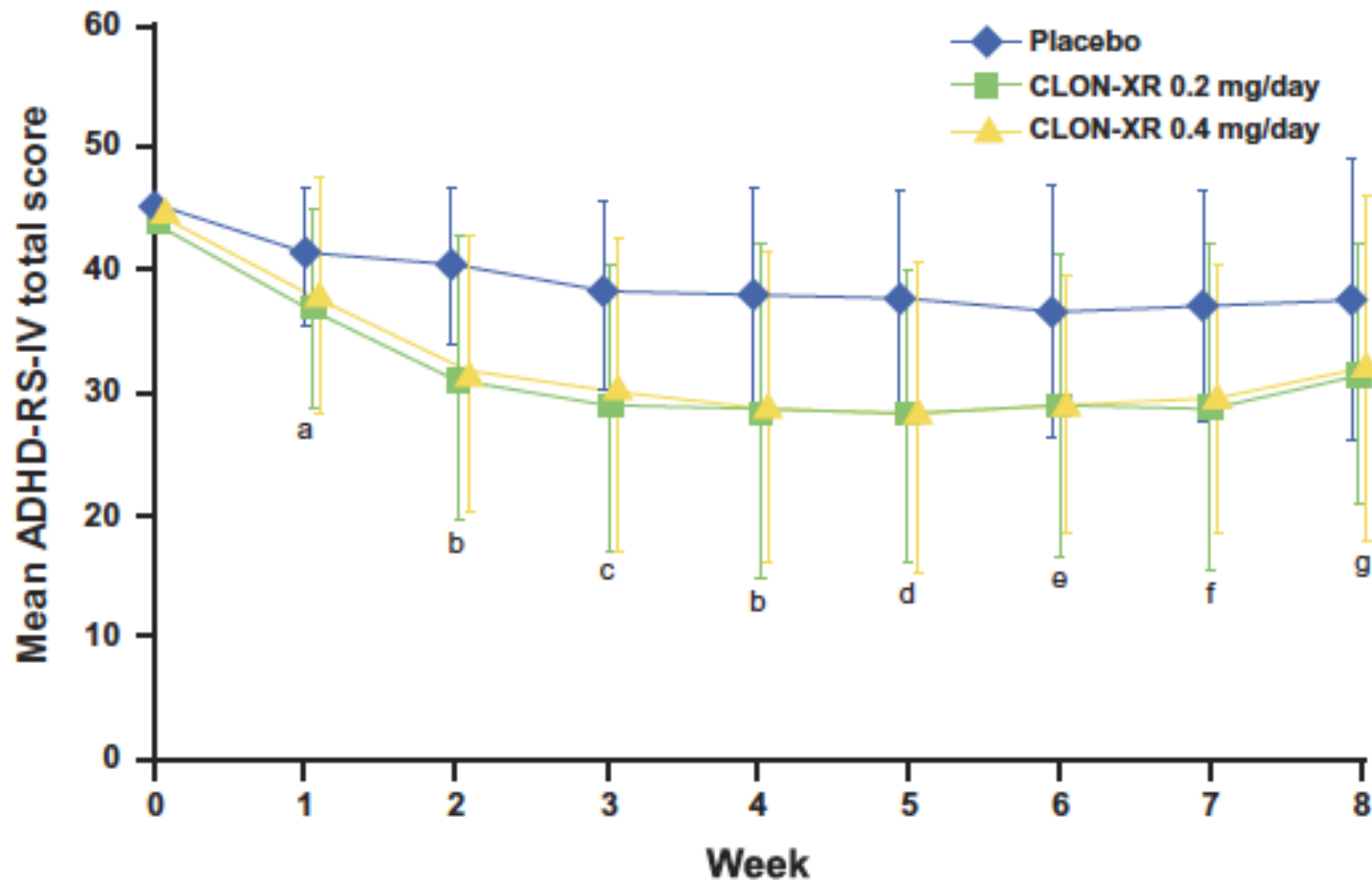


Meta-Analysis: Treatment of Attention-Deficit Hyperactivity Disorder in Children with Comorbid Tic Disorders. Bloch *et al* JAACAP 9.2009

Placebo
(n = 76)

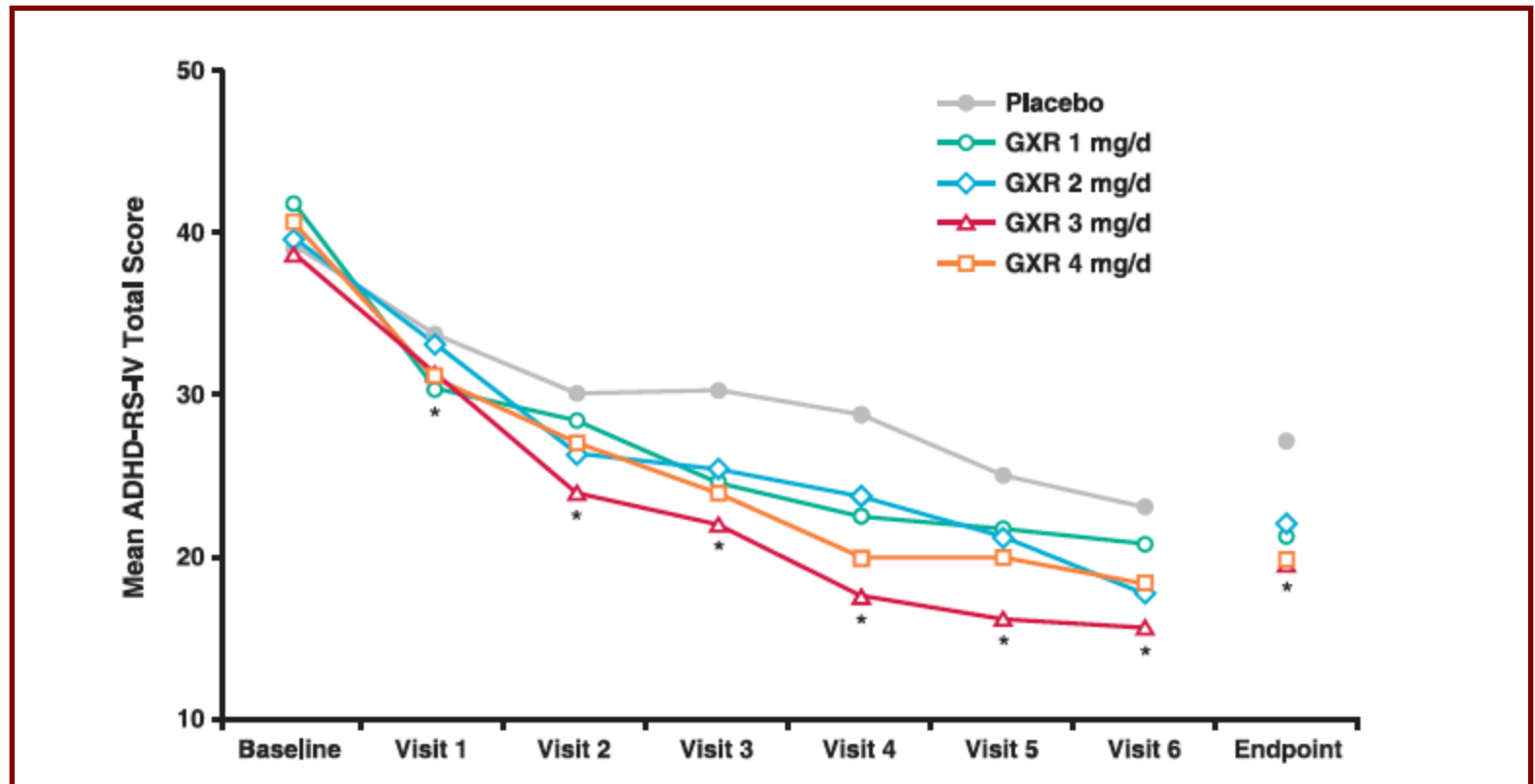
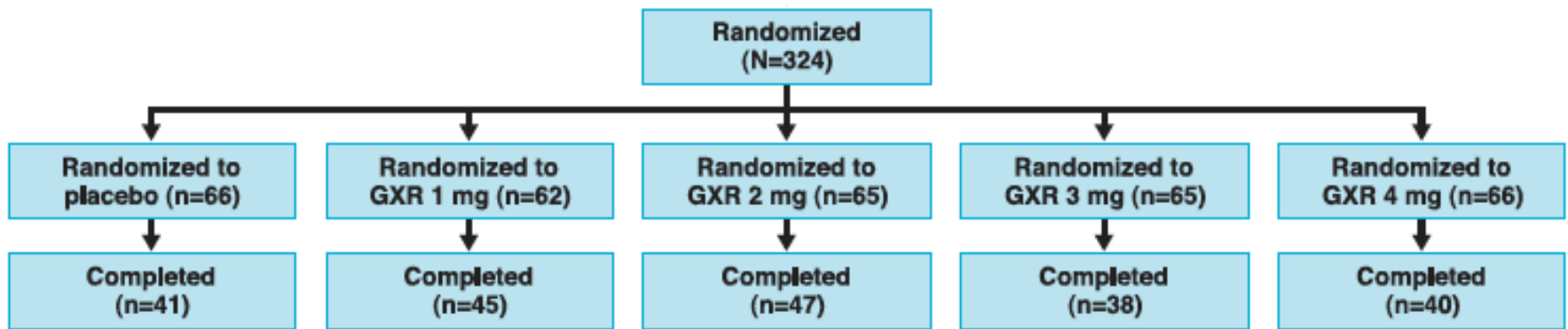
CLON-XR 0.2 mg/day
(n = 74)

CLON-XR 0.4 mg/day
(n = 78)



Jain et al 2.11

Clonidine Extended-Release Tablets for Pediatric Patients with ADHD



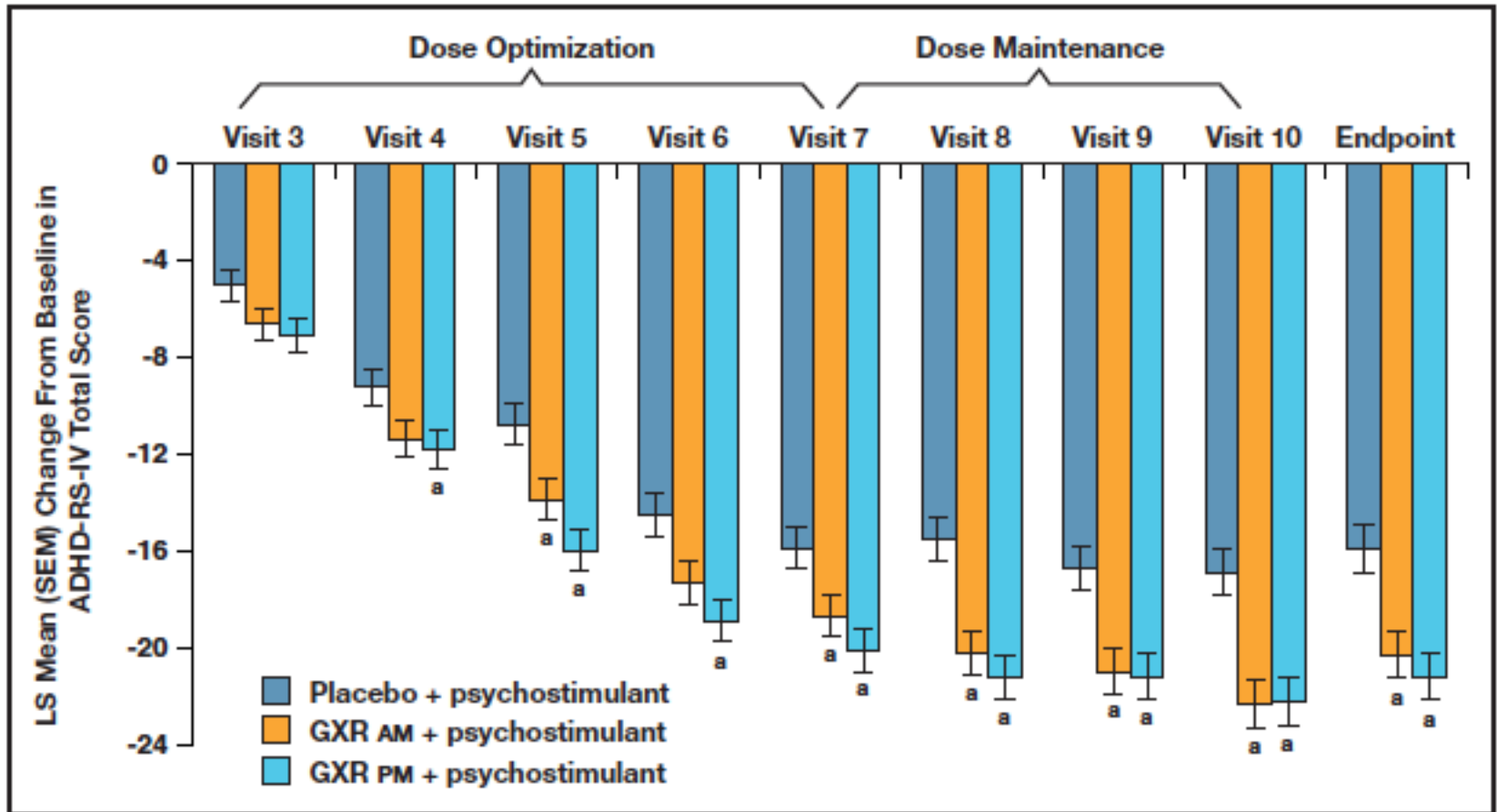
Sallee et al 2.09
 Guanfacine Extended Release in Children and Adolescents With ADHD:
 A Placebo-Controlled Trial

Placebo +
Psychostimulant
(n = 153)

GXR AM +
Psychostimulant
(n = 150)

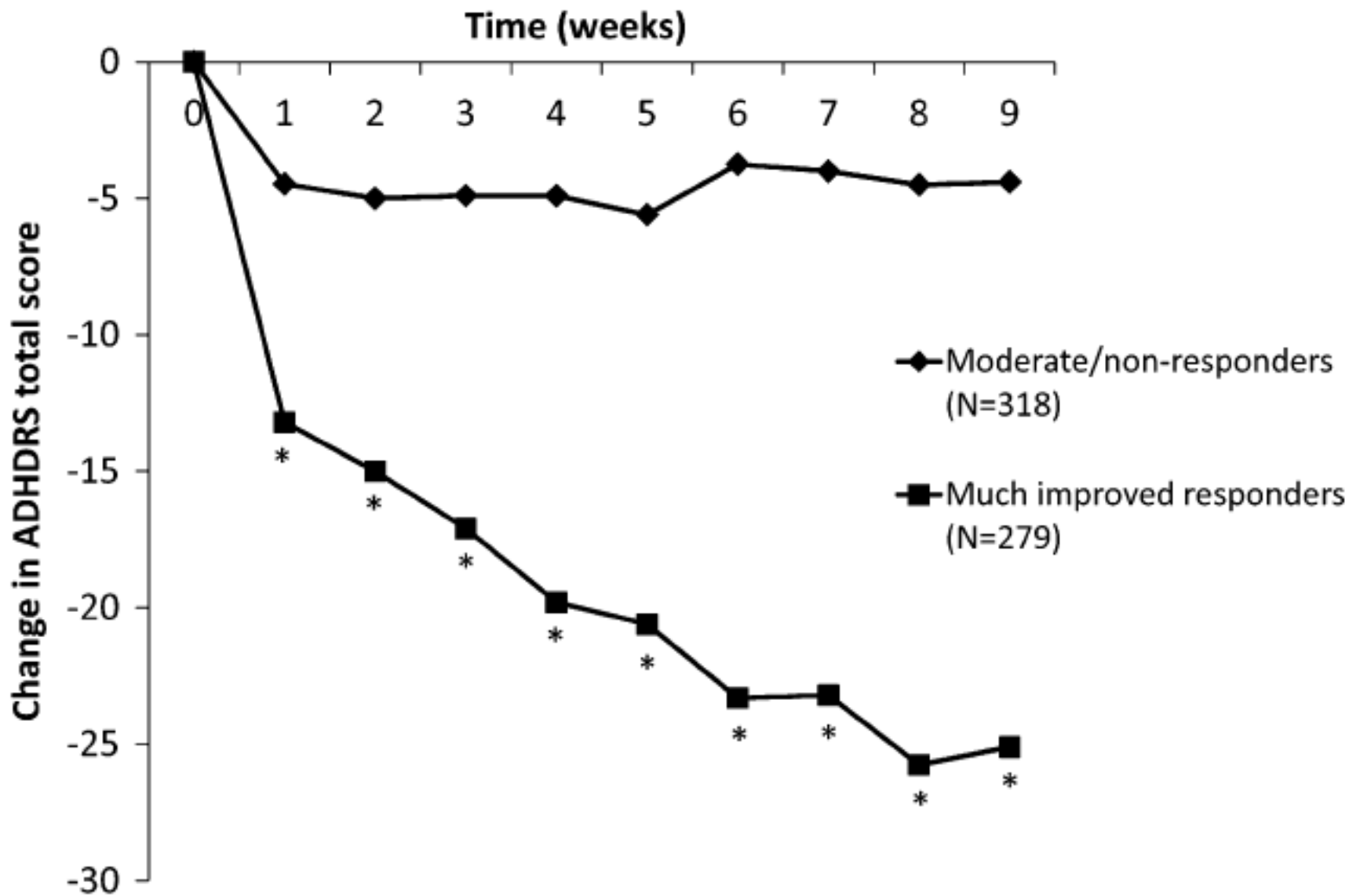
GXR PM +
Psychostimulant
(n = 152)

FAS/Safety
Population
(N = 455)



Wilens et al 1.12

A Controlled Trial of Extended-Release Guanfacine and Psychostimulants for ADHD



Newcorn et al 5.09
Clinical Responses to Atomoxetine in ADHD:
The Integrated Data Exploratory Analysis (IDEA) Study

Atomoxetine + MI/CBT

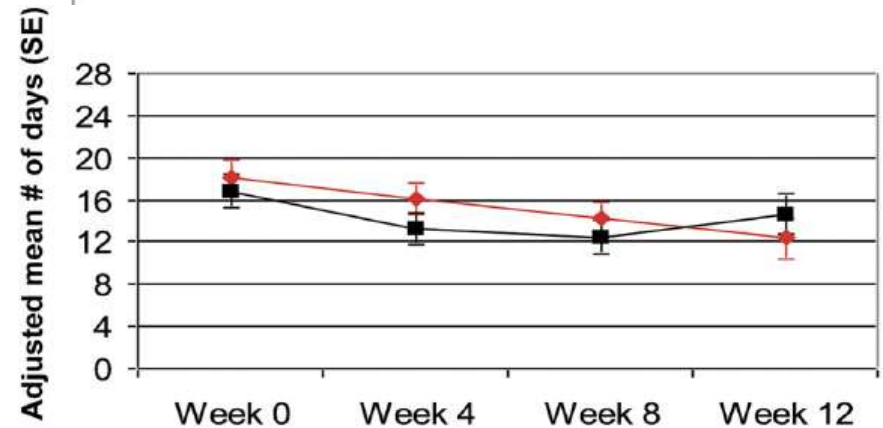
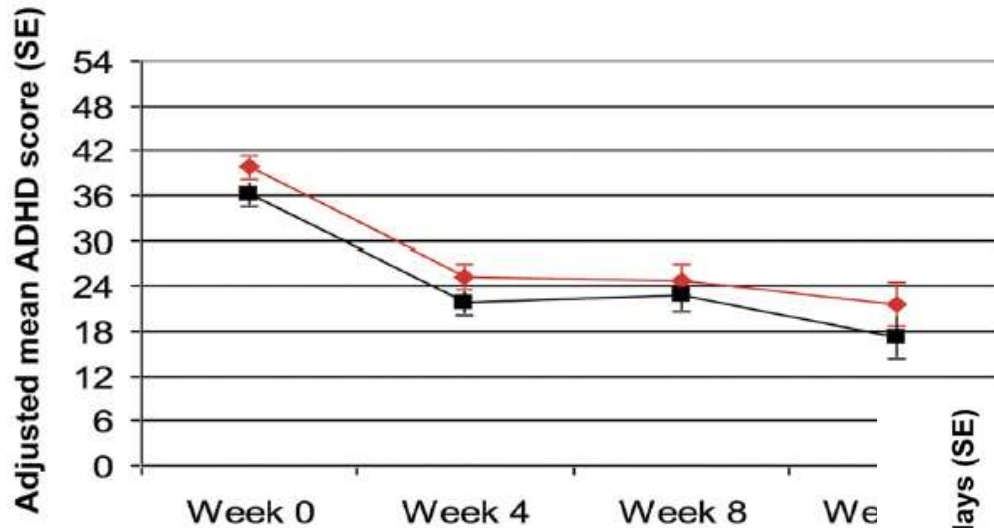
$n = 35$

n (%) or mean (SD)

Placebo + MI/CBT

$n = 35$

n (%) or mean (SD)

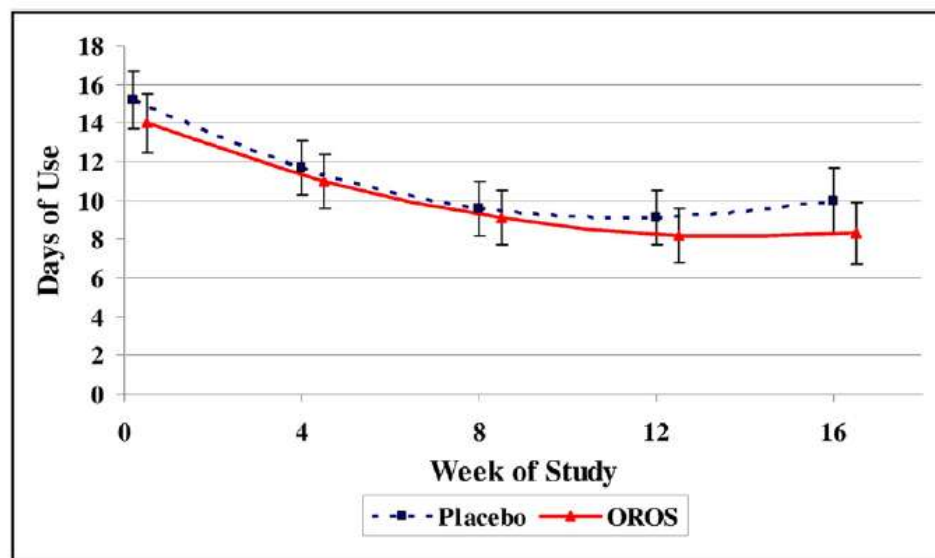
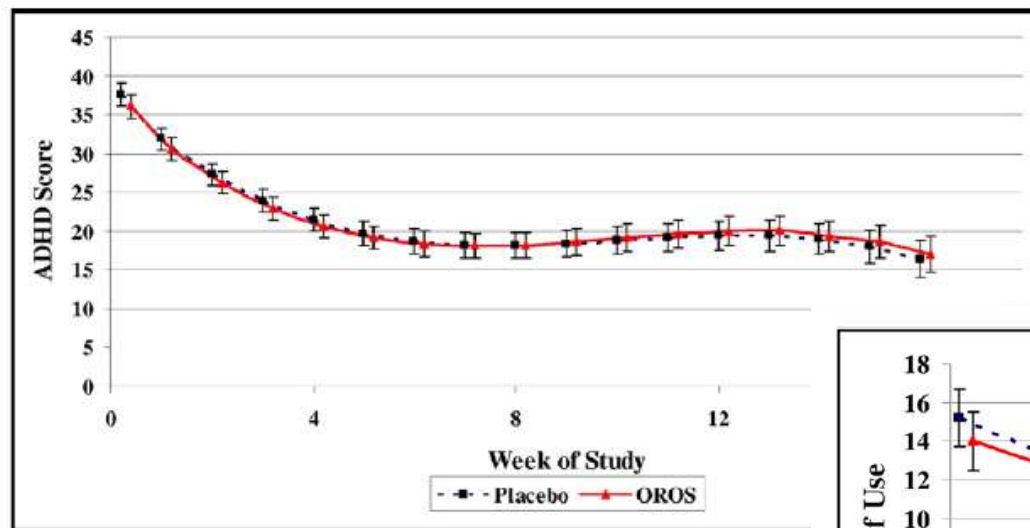


Thurstone et al 6.10
RCT of Atomoxetine for ADHD in Adolescents with Substance Use Disorder

OROS-MPH + CBT
(n = 151)

Placebo + CBT
(n = 152)

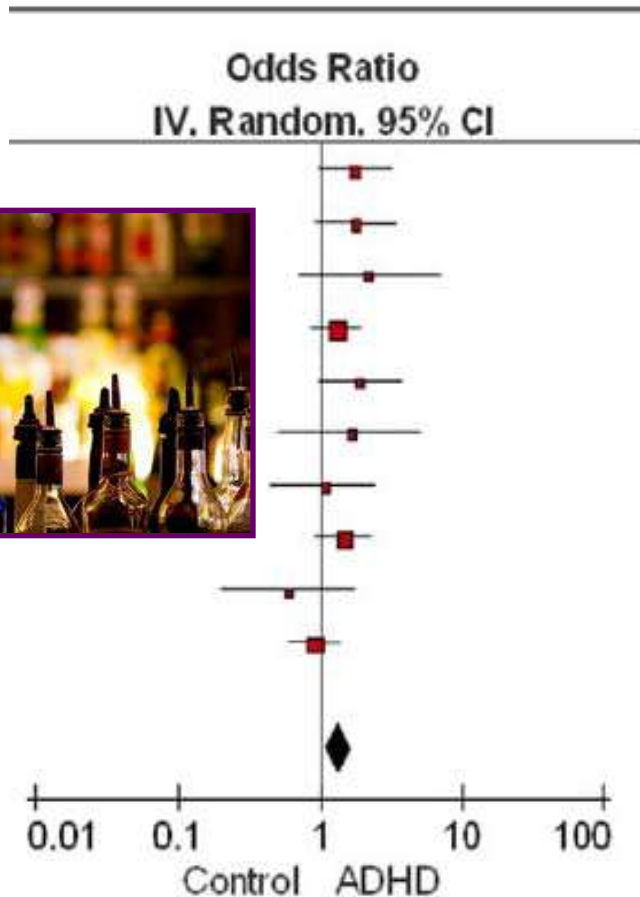
All
(N = 303)



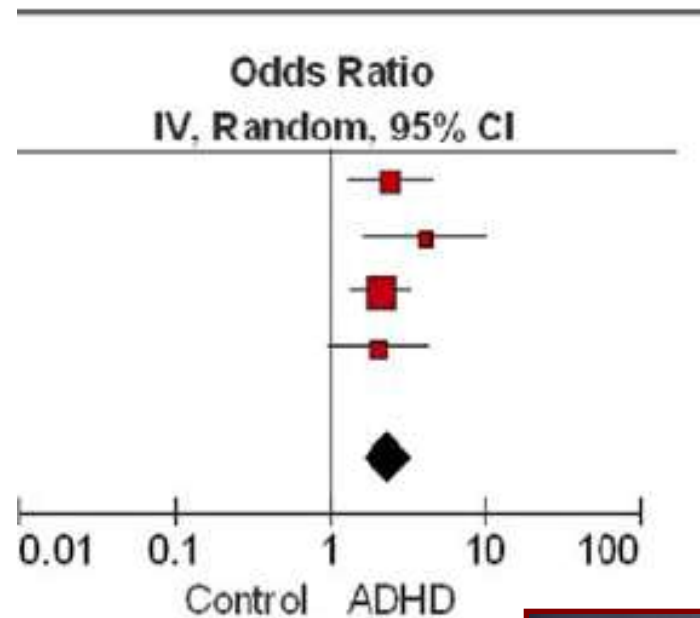
Riggs et al 5.11
RCT of Osmotic-Release Methylphenidate with CBT
in Adolescents with ADHD and Substance Use Disorders

- 21 studies
- 113,104 subjects
- Past-year misuse rates:
 - 5-9% grade and high school
 - 5-35% college
- Lifetime diversion rates:
 - 16-29%
- Risk Factors:
 - Whites>minorities
 - Fraternities / sororities
 - IR>ER forms





ETOH N=3,184
OR 1.35 (1.11, 1.64)



Nicotine N=3,184
OR 1.35 (1.11, 1.64)



Charach et al 1.11
Childhood ADHD and Future Substance Use Disorders: Comparative Meta-Analysis

The Real World: Unmet Need

- Longitudinal cohort, Los Angeles
- N=530, ages 5-11
- Three six-month intervals
- Primary care or MH clinics
- No treatment at 18 months: 34-44%
- Refill persistence: 31-49%

Zima et al 12.10

Quality of Care for Childhood ADHD in a Managed Care Medicaid Program

The Real World: Adherence

- Philadelphia public schools
- Medicaid eligible children
- Pharmacy claims, N=3,543
- Outcome = GPA
- Adherence = >0.7 med. possession
- $r = 0.108$ ($p < 0.001$)
- Only 18.6% adherent



Our Past Informs Our Future



- Pediatric psychopharmacology the front end of a lifespan developmental model
- Electronic medical records + biospecimens = switch from phenotypic to molecular medicine
- Genomics and proteomics revolution = preemptive, preventive and curative care

March 5.11

The Preschool ADHD Treatment Study (PATs) as the Culmination of Twenty Years of Clinical Trials in Pediatric Psychopharmacology









Gracias - y olé !



Publicaciones Relevantes

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DSM ICD