

Pediatric ADHD

Elkhorn Spring Primary Care Conference May 12, 2018





Simon Trepel, MD FRCPC



• Staff Psychiatrist, MATC

- Manitoba's largest provider of Mental Health Services for Children, Youth & Families
- ICRS, GDAAY, PASC Programs



- Assistant Professor
- Department of Psychiatry, Faculty of Medicine, University of Manitoba

Acknowledgement

I acknowledge that today we are on Treaty One territory and on the homeland of the Metis Nation.

I also acknowledge that Manitoba includes the original lands of Anishinaabeg, Cree, Oji-Cree, Dakota and Dene people and Inuit people

We respect the Treaties that were made on these territories and we acknowledge the harms and mistakes of the past. We dedicate ourselves to moving forward in partnership with First Nations, Metis and Inuit communities in a spirit of reconciliation and collaboration.



Outline and Objectives

- Prevalence
- Causes and neurobiology
- DSM5 criteria
- Clinical presentations
- Diagnosis
- Treatment



ADHD

- Single most common psychiatric disorder of childhood
 - 20+ studies (child) US population
 - 30+ studies (child) Non-US population
 - Both studies found the same prevalence ranges



Kessler et al, Am J Psych, 2006; Spencer et al, Ambul Ped. 2007; World Psychiatry 2003 (June); Faraone et al.



Epidemiology

ADHD symptoms	Children/Teens	Adults
Prevalence	6-8%	4.4%
Gender ratio	2.5:1	1:1
Combined subtype	50-75%	30-40%
Predominantly inattentive	20-30%	50-60%
Predominantly hyperactive/impulsive	15%	5%
Percent diagnosed	33%	7%



Estimated ADHD Sufferers in Canada

	ADHD in children/teens (age 5-19)	ADHD in adults (age 20-59)
Total Population	6, 182, 933	18, 567, 976
Prevalence [%]	6%	4%
Patients w/ ADHD	370,976	742,719
% Diagnosed	33%	7%
Patients Untreated	248,544	690,729

Stats Can, 2004 projected to 2005; % diagnosed calculated based on estimate of treated patients in Canada



What are symptoms of ADHD?

- Hyperactivity
- Inattention
- Impulsivity
- Emotional dysregulation



Impact of ADHD beyond symptoms



Mental Health Services for Children, Youth and Families

Causes- Multifactorial

Catecholamines

Less norepinephrine and dopamine in specific brain areas

• Genetics

- 75-80% higher risk if 1st degree relative
- Novelty seeking, poor frustration tolerance
- **CNS insults-** typically during pregnancy and delivery
 - IUGR, LBW, SUD in utero, toxins (eg. lead)
 - TBI, hypoxia, encephalitis
- Environmental factors- may perpetuate symptoms
 - Family dysfunction, abuse, parental difficulty



Heritability of ADHD



Mean heritability of ADHD = .76

Role of Norepinephrine in Attention



- Executive Functions
- Engage and disengage from stimuli
- Change focus to new stimuli
- Increases inhibition
- Inhibit distraction
- Dampens 'background' noise

Solanto. Stimulant Drugs and ADHD. Oxford; 2001.

Role of Dopamine



Nigrostriatal: movement Mesolimbic: pleasure, psychosis Mesocortical: attention, arousal

- Enhances signal
- May improve mood
- Improves attention
 - Focus
 - On-task behavior
 - On-task cognition

What's going on in the ADHD brain?

- Frontal lobe under-activated
- Brainstem reward and movement systems under-activated
- Poor cerebellar and intra-hemispheric communication
 - Coordination problems, delayed milestones
- Smaller brains that are less active
 - Hyperactivity, distractible/novelty seeking, reactive



Neurobiology of adolescence



Variations in the structure and function of the frontoparietal and frontostriatal circuits are responsible for enhanced selfcontrol

Frontal Lobe Underactivated

Poor executive functioning (NE and DA)

- Activation- able to learn at higher level
- Focus- sustaining vs. task switching
- Effort- processing speed
- Emotion- frustration, silliness
- Memory- working memory and recall
- Action- self regulation, impulsivity

Poor modulation of movement and activity (NE and DA)

- Hyperactivity
- <u>Alternate reward seeking behavior (DA)</u>
 - Underwhelmed and offline in low stimulation
 - May often seek 'negative' attention



ADHD: Neurobiologic Basis



Pre Frontal Cortex

- Judgment
- Analysis
- Problem Solving
- Critical Thinking
- Forward Thinking

Dorsolateral PFC

- Organization
- Working Memory
- Attention
- Orbital Frontal Area
 - Impulse Control
 - Socialization



Posner and Raichle. Images of the Mind. Scientific American Books; 1996.



Neuroanatomical differences: the brain and ADHD

- Thinner areas of the dorsolateral prefrontal cortex (DLPC)
- Smaller anterior cingulate cortex (ACC)
- More white matter
- Generally less grey matter
- Larger nucleus accumbens



Reward and movement underactivated

- Less dopamine in nucleus accumbens
 - Looking for anything more interesting all the timeinstant gratification
 - Novelty/stimulating activity seeking behavior
 - Difficult to structure someone always looking for something more interesting than requested task

- Less dopamine in nigrostriatal and basal ganglia
 - Poor regulation of movement



DSM5 Criteria



DSM5 Criteria

Neurodevelopmental condition characterized by a persistent pattern of:

Inattention and/or

Hyperactivity-Impulsivity

Interferes with:

-Functioning and development

-School/work

-Occurs in multiple settings (home/school/social)



Symptoms

- Pervasive many settings
- Persistent duration is 6 months
- Onset before age 12
- Severe
- Maladaptive, affects functioning
- Inconsistent with developmental level
- Clinical significant impairment
- Social, academic, occupational problems



DSM-5 Inattention in ADHD

6 symptoms for 6 months (only 5 needed if over 16)

- Careless Mistakes
- Difficulty sustaining attention in activities
- Doesn't seem to listen
- No follow through
- Disorganized, messy

- Avoids/dislikes tasks requiring sustained mental effort
- Often loses things necessary for tasks
- Often distracted by extraneous stimuli
- Forgetful chores
- * Inattention maladaptive/inconsistent with development level



Hyperfocus

WHY WE AREN'T





DSM-5 Hyperactivity-Impulsivity in ADHD

6 symptoms for 6 months (only 5 needed if over 16)

- Often fidgets squirms in seat
- Can't stay seated
- Runs/climbs excessively
- On the go driven by motor

- Talks excessively
- Blurts out answers
- Difficulty waiting turn
- Interrupts/intrudes on others

Maladaptive and inconsistent with developmental level



Clinical Presentation of ADHD



Clinical Presentation in School Aged Children

- Full of energy and out of seat
- Easily distracted
- Disruptive in class and frequent trips to Principal's office
- Homework poorly organized, careless errors, often incomplete or lost
- Low academic scores
- Often interrupts and intrudes on others
- Low self-esteem



- Displays aggression
- Difficult peer relationships
- Does not wait turns in games
- Perception of "immaturity"
- Unwilling or unable to do chores at home
- Accident prone

Clinical Presentation In Teenagers



- More subtle than for younger children
- Increased boredom
- Significant school underachievement
- Inner restlessness instead of hyperactivity
- Poor independence, self-esteem, peers, authority
- "Risky" behavior- sex, speeding, substance abuse
- More injuries and accidents
- Increased risk of suicide attempt, especially when associated with mood, CD, SUD problems
- 15-20% of children have poor academic outcome
- Almost 30% of ADHD subjects fail grades
- 10% of ADHD pupils suspended
- 25% persistently antisocial

Developmental trajectory of ADHD



Harke A. Bosma. "Introduction à la psychopathologie développementale ", L'orientation scolaire et professionnelle [Online], 35/2 | 2006, posted online September 28, 2009, consulted August 6, 2012. Adapted by A. Dansereau URL: http://osp.revues.org/index1097.html

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Diagnosing ADHD



When is ADHD first diagnosed?

- ADHD most often diagnosed at age 7
- Symptoms appear as early as 3-7 years
 - 4-year-olds settle down, take naps, sit for meals
 - Child with ADHD is 'on the go' all the time
- Commonly diagnosed due to academics
 - Easily quantifiable





Initial screening questions

- Do you feel restless or the need to always move around?
- Do you get easily distracted?
- Do you find it hard to concentrate at school, home, or during certain activities?
- Do you get easily annoyed, called annoying, or are difficult in groups?
- Do you get in trouble for doing things without thinking?
- Does anyone in your family have ADHD?



Red flags

- Having to work or study harder than peers to get the same or worse grades
- Certain subjects which were easy while others extremely difficult
- Needing a lot of support and structure from parents and teachers
- Bored all the time and can't structure self
- Frequent accidents, injuries, or fights
 - Punching walls, breaking game controllers
- Explosive anger followed by apology later



Diagnosing ADHD

- Clinical observations and collateral
 - Ruling out sensory problems, seizure, thyroid
- Academic record
 - Learning assessments and disabilities
- SNAP4
 - Children and adolescents
 - ASRS for adults
- Psychological testing
 - Conner's testing for working memory deficits
 - Cognitive testing for FSIQ and adaptive function
- Occupational therapy assessment
 - Coordination, attention tests, adaptive function
- Functional imaging (lol)



Physical exam

- Vital signs
 - Heart rate
 - Blood pressure
- Weight
 - Underweight kids and stimulants



- Height
 - Stimulants don't have significant effect
- Cardiovascular
 - Arrhythmias
 - Structural defects (HOCM)
 - Consider EKG if no family history or concerning personal/family hx



Differential Diagnosis in ADHD

• Psychiatric disorders

- Age appropriate behavior
- Stress or adjustment disorder
- ODD or conduct
- Intermittent explosive
- Other neurodevelopmental
 - ID, learning, language, ASD
- Mood/anxiety disorders
- Disruptive mood dysregulation
- Tic disorder
- Substance use

Medical disorders

- Fetal alcohol syndrome
- Sleep apnea
- Substance use disorder
- Developmental disorders
- Use of other medications
- Seizure disorder
- Vision or hearing problems
- Thyroid disorder


Comorbidity of ADHD 70% have 1, 30% have none, 20% have >1

- Oppositional defiant (50%)
- Coordination problems (33%)
- Anxiety (30%)
- Learning disability (30%)
- Depression or dysthymia (25%)
- Conduct disorder (20%)
- SUD (15%)



- Tic disorder or Tourette's (10%)
- Communication disorder



ADHD: Simple versus Complex

Simple

- Patients with one subtype of ADHD, but no other learning disabilities or co-morbid disorders
- One in five cases

Complex

- Combined learning disabilities
- Development disorders and / or
- Associated comorbid psychiatric disorders

Canadian ADHD Resource Alliance (CADDRA): Canadian ADHD Practice Guidelines, 3rd Edition, 2011.



Treatment



Functional Impairment in Patients with ADHD



Barkley RA. Attention-Deficit Hyperactivity Disorder. A Handbook for Diagnosis and Treatment, 1998 Barkley RA, et al. J Am Acad Child Adolesc Psychiatry. 1990;29:546-557. Biederman J, et al. Arch Gen Psychiatry. 1996;53:437-446. Weiss G, et al. J Am Acad Child Psychiatry. 1985;24:211-220. Satterfield JH, Schell A. J Am Acad Child Adolesc Psychiatry. 1997;36:1726-1735. Biederman J, et al. Am J Psychiatry. 1995;152:1652-1658.

Untreated & Under-Treated ADHD

Society

Substance Use Disorders:

2 X Risk⁸

Earlier Onset⁹

Less Likely to Quit

in Adulthood¹⁰

Health Care System

50% ↑ in bike accidents¹ 33% ↑ in ER visits²

2-4 x more motor vehicle crashes³⁻⁵

School & Occupation

46% Expelled⁶ 35% Drop Out⁶ Lower Occupational Status⁷

Family

3-5x ↑ Parental Divorce or Separation^{11,12}
2-4 x ↑ Sibling Fights¹³

1. DiScala et al., 1998

2. Liebson et al., 2001

4-5. Barkley et al., 1993; 1996

3. NHTSA. 1997

Sibiling Fights

Barkley, et al., 1990
 Manuzza et al., 1997
 Biederman et al., 1997

9. Pomerleau et al., 1995
 10. Wilens et al., 1995
 11. Barkley, Fischer et al., 1991

Employer
↑ Parental
Absenteeism¹⁴

and

↓ Productivity¹⁴



Start ADHD Treatment here...

- Lifestyle
 - Proper nutrition
 - Good sleep hygiene
 - Positive family time
 - Regular exercise
 - Extracurricular activities
 - Stress management
 - Go outside
 - Screen time



Psychosocial Treatments for ADHD

- Psychoeducation
- Individual Education
 Plan (Academic)
- Organization & time
 management skills
- Behavioral Parent Management Training
- Remediation
- Social Skills

- Anger management
- Relaxation Technique (anger, anxiety)
- Individual psychotherapy (low self-esteem)
- Family therapy to prevent "scapegoating"
- Yoga and meditation currently under study



Off Meds

On Meds



Medication Treatment

80% effective in studies, for kids 6 and older

<u>Stimulants</u>

*Dextro-Amphetamine *Methyl-Phenidate

Non Stimulants

*Atomoxetine (Strattera) *Clonidine and guanfacine (Intuniv XR) Bupropion (Wellbutrin, Zyban) Effexor XR



Methylphenidate- (0.5-1mg/kg/day)

- Ritalin IR (3-4h)
 - 5, 10, 20mg
- Ritalin LA or SR (6-8h)
 - 10, 20mg
- Concerta (10-12h)
 - 18, 27, 36, 54mg

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NET WT. 180Z HLB 20Z

- Non-crushable therefore hard to abuse
- Biphentin (10-12h)
 - 10, 15, 20, 30, 40, 50, 60, 80, 100mg



CONCERTA*: OROS Technology to Clinical Efficacy



Biphentin Multi-layer release (MLR)





Dextroamphetamines- 0.25-0.5mg/kg/day

- Dexedrine (4-6h)
 - 5, 10mg
- Dexedrine spansules (6-8h)
 - 10, 15mg
- Adderal XR (10-12h)
 - 5, 10, 15, 20, 25, 30mg
- Lisdexamphentamine-Vyvanse (13-14h)
 - 20, 30, 40, 50, 60mg





Amphetamine mixed salts (Adderall XR)



d- and *l*amphetamine salt at 3:1 ratio

 Equal quantities of IR and ER beads

Lisdexamfetamine (Vyvanse)





Lisdexamfetamine dimesylate





d-amphetamine

Comparison of Newer Long-acting Stimulants

Drug	Release Mechanism	% of dose immediately released	Durations	Available Doses
MLR methylphenidate	Multi-layer Beads Bi-modal	40%	10-12 hours	8
OROS methyphenidate	Water Absorption push mechanism	22%	10-12 hours	4
Mixed amphetamine salts XR	Two Beads BID delivery	50%	10-12 hours	6
Lisdexamfetamine	Prodrug, lysine is cleaved in blood stream	Continuous	13-14 hours	5

Canadian ADHD Resource Alliance (CADDRA): Canadian ADHD Practice Guidelines, 3rd Edition, 2011.

Benefits of Long Acting Stimulant Formulations

- Core impairments continue all day
- Use of long-acting stimulant formulation may improve medication compliance
- May decrease abuse potential
- Smoother, more consistent coverage

Pliszka SR et al., J Am Acad Child Adolesc Psychiatry. 2006 Jun;45(6):642-57. Swanson J., CNS Drugs, 2003; 17:117-131.

Common stimulant side effects

- Extra energy
- Mild euphoria
 - (first few days)
- Tachycardia
- Sweating
- Headaches

- Insomnia
- Poor appetite
- Anxiety
- Caffeine 'tremor'
- Weight loss



Concerning Stimulant Side Effects

• <u>Seizures</u>

 safe with well controlled epilepsy

Insomnia

 cause of mania if underlying bipolar disorder

<u>Tolerance or craving</u>

- addiction/abuse/diversion
- Panic attacks or dysphoria
 - with comorbid depression

- <u>Unmasking of tics</u>
 - sometimes improves tics
- <u>Psychosis</u>
 - 1/1000 generally, higher in SCZ
- Possible growth suppression
 - small loss and likely reversible with withdrawal of treatment
- <u>Heart attack</u>
 - undiagnosed arrhythmia/HOCM



Atomoxetine (Strattera) NRI 0.5mg/kg/day

- 10, 18, 25, 40, 60, 80, 100mg
- Patients 6-17 not responding to stimulants, significant side effects, or Tourette's Syndrome/Chronic Motor Tic Disorder
- Especially helpful with ADHD and comorbid Anxiety- 24 hour duration
- 65% effective in studies
- 3 total studies (n=558), 3/3 positive
 - 3 double-blind, placebo-controlled tria
 - No crossover studies, no comparator triais
- Side effects include
 - Headache, somnolence, dizziness
 - abdominal pain, nausea, vomit and decreased appetite
 - Rare hepatic failure
 - FDA black box warning for suicidal ideation





Alpha 2 Adrenergic Agonists

- Act on presynaptic autoreceptor inhibiting NE release
- 50% effective, up to 24 hour duration of action
- For kids 6-17 years
- <u>Clonidine</u>- start 0.025-0.05 BID, up to 0.1-0.4mg/d
- <u>Guanfacine</u>
- Guanfacine XR (Intuniv XR) 1, 2, 3, 4mg
- Side effects include:
 - Somnolence, headache, fatigue, dizzy, sedation, insomnia, dry mouth, decreased pulse and BP, rebound tach



Antidepressants

• Wellbutrin (NDRI)

- Increases NE and DA
- 55% effective in studies
- 4 total studies (n=257), 4/4 positive
 - o 2 double-blind, placebo-controlled trials
 - \circ No crossover studies, no comparator trials
 - $\odot\,\textbf{1}$ open-label study with comorbid bipolar disorder
- Effexor XR (SNRI)



Alternative Treatments NOT **Confirmed Efficacious for ADHD**

- Biofeedback
- Fish Oils
- Herbal Remedies

Canadian Pediatric Society. The use of alternative therapies in treating children with attention deficit hyperactivity disorder. 2002.

Drug-drug Interactions of Stimulants

- Caution with monoamine oxidase inhibitors (MAOIs) e.g. phenelzine
- Inhibits metabolism of warfarin, anticonvulsants, tricyclic antidepressants
 need to adjust dose



Therapeutic approaches: ADHD



Treatment algorithms for comorbid (Always treat most severe disorder first)

- **Depression** treat moderate to severe mood first and assess safety, stimulant may help improve low energy and motivation of depression
- **<u>Anxiety</u>** consider Strattera early in treatment
- **<u>Autism spectrum disorder</u>** stimulants may agitate thus helpful to have atypical antipsychotic started first
- **Bipolar and psychosis** treat before adding stimulant
- <u>**Tics</u>** consider clonidine</u>
- **<u>Aggression</u>** treat ADHD first, then add atypical or alpha2 agonist
- **Epilepsy** treat epilepsy first, new onset seizure managed with antiepileptic medication, antiepileptic level may increase with MPH (enzyme inhibition)
- **<u>SUD</u>** important to stabilize the addiction before prescribing stimulant



Disruptive Behaviour Disorder and ADHD

Oppositional Disorder

- Treat both at same time
- Ensure psychosocial interventions
- Severe cases may require combos of psychostimulants and alpha 2 agonist

 Treatment of ADHD symptoms often causes significant improvements in ODD, CD and emotional impulsivity (Barkley)

Conduct Disorder

- Treat both at same time
- Ensure psychosocial interventions
- Severe cases may require combos of psychostimulants and alpha 2 agonist
- Expect legal issues
- Adding an antipsychotic might improve the symptoms of conduct disorder, according to some cases cited in the literature.



What Types of Stimulants are Most Abused?



Bright GM, Medscape J Med, 2008.

Top Drugs among 8th and 12th Graders, Past Year Use



* Only 12th graders surveyed about sedatives use

Source: University of Michigan, 2014 Monitoring the Future Study

ADHD Resources

- www.adhd.net
- www.caddra.ca
- www.aacap.org
- <u>www.caddac.ca</u>

• Parent ADHD support groups



CADDRA Guide to ADHD Pharmacological Treatments in Canada - 2018

Medications available and illustrations	Characteristics	Duration of action 1	Starting dose ²	Dose titration as per product monograph	Dose titration as per CADDRA www.caddra.ca		
AMPHETAMINE-BASED PSYCHOSTIMULANTS							
Dexedrine®	Pill can be	~ 4 h	Tablets = 2.5 to 5 mg BID	↑ 2.5 - 5 mg at weekly intervals;	↑ 2.5 - 5 mg/day at weekly intervals		
Dexedrine® spansules 10, 15 mg	Spansule (not crushable)	~ 6 - 8 h	Spansules = 10 mg q.d. a.m.	Max. dose/day: (q.d. or b.i.d.) All ages = 40 mg	Max. dose/day: (q.d. or b.i.d.) Children and Adolescents = 20 - 30 mg Adults = 50 mg		
Adderall XR® Capsules 5, 10, 15, 20, 25, 30 mg	Sprinkable Granules	~ 12 h	5 - 10 mg q.d. a.m.	↑ 5 - 10 mg at weekly intervals Max. dose/day: Children = 30 mg Adolescents and Adults = 20 - 30 mg	Children: ↑ 5 mg at weekly intervals Max. dose/day = 30 mg Adolescents and Adults: ↑ 5 mg at intervals max. dose/day = 50 mg		
Vyvanse® 540 54	Capsule content can be diluted in water, orange juice and yogurt	~ 13 - 14 h	20 - 30 mg q.d. a.m.	↑ by clinical discretion at weekly intervals Max. dose/day: All ages = 60 mg	↑ 10 mg at weekly intervals Max. dose/day: Children = 60mg Adolescents and Adults = 70 mg		
METHYLPHENIDATE-BASED PSYCHOSTIMULANTS							
Methylphenidate short acting, tablets 5 mg (generic) 10, 20 mg (Ritalin®)	Pill can be crushed ³	~ 3 - 4 h	5 mg b.i.d. to t.i.d. Adult = consider q.i.d.	↑ 5 - 10 mg at weekly intervals Max. dose/day: All ages = 60 mg	↑ 5 mg at weekly intervals Max. dose/day: Children and Adolescents = 60 mg Adults = 100 mg		
Biphentin® (1100) (120) Capsules 10, 15, 20, 30, 40, 50, 60, 80 mg (100)	Sprinkable Granules	~ 10 - 12 h	10 - 20 mg q.d. a.m.	↑ 10 mg at weekly intervals Max. dose/day: Children and Adolescents = 60 mg Adults = 80 mg	↑ 5 - 10 mg at weekly intervals Max. dose/day: Children = 60 mg Adolescents and Adults = 80 mg		
Concerta® Extended Release Tabs 18, 27, 36, 54 mg	Pill needs to swallowed whole to keep delivery mechanism intact	~ 12 h	18 mg q.d. a.m.	↑ 18 mg at weekly intervals Max. dose/day: Children = 54 mg Adolescents = 54 mg / Adults = 72 mg	↑ 9 - 18 mg at weekly intervals Max. dose/day: Children = 72 mg Adolescents = 90 mg / Adults = 108 mg		
Foquest* Capsules 25, 35, 45, 55, P L L L L L L L L L L L L L L L L L L	Sprinkable Granules	~ 16 h	25 mg q.d. a.m.	↑ 10-15 mg in intervals of no less than 5 days Max. dose/day: Adults = 100 mg	↑ 10-15 mg in intervals of no less than 5 days Max. dose/day: Adults = 100 mg		
NON PSYCHOSTIMULANT - SELECTIVE NOREPINEP	HRINE REUPTAKE INHIB	ITOR					
Strattera ^{MD} (Atomoxetine) Capsules 10, 18, 25, 40, 60, 80, 100 mg	Capsule needs to swallowed whole to reduce GI side effects	Up to 24 h	Children and Adolescents : 0.5 mg/kg/day Adults = 40 mg q.d. for 7-14 days	Maintain dose for a minimum of 7 - 14 days before adjusting: Children = 0.8 then 1.2 mg/kg/day 70 kg or Adults = 60 then 80 mg/day Max. dose/day : 1.4 mg/kg/day or 100 mg	Maintain dose for a minimum of 7 - 14 days before adjusting: Children = 0.8 then 1.2 mg/kg/day 70 kg or Adults = 60 then 80 mg/day Max. dose/day: 1.4 mg/kg/day or 100 mg		
NON PSYCHOSTIMULANT - SELECTIVE ALPHA-2A ADRENERGIC RECEPTOR AGONIST							
Intuniv XR® (Guanfacine XR) Extended release tabs 1, 2, 3, 4 mg	Pills need to be swallowed whole to keep delivery mechanism intact	Up to 24 h	1 mg q.d. (morning or evening)	Maintain dose for a minimum of 7 days before adjusting by no more than 1 mg increment weekly Max. dose/day: Monotherapy: 6-12 years = 4 mg, 13-17 years = 7 mg As adjunctive therapy to psychostimulants 6-17 years = 4 mg	Maintain dose for a minimum of 7 days before adjusting by no more than 1 mg increment weekly Max. dose/day: Monotherapy: 6-12 years = 4 mg, 13-17 years = 7 mg As adjunctive therapy to psychostimulants 6-17 years = 4 mg		

Note: Illustrations do not reflect real size of pills/capsules. For specific details on how to start, adjust and switch ADHD medications, clinicians are invited to refer to the Canadian ADHD Practice Guidelines (www.caddra.ca) ¹ Pharmacokinetics and pharmacodynamic response vary from individual. The clinician must use clinical judgement as to the duration of efficacy and not solely rely on reported values for PK and duration of effect. ² Starting doses are from product monographs. CADDRA recommends generally starting with the lowest dose available. ³ Higher abuse potential. * Vyvanse 70 mg is an off-label dosage for ADHD treatment in Canada. *Document developed by Annick Vincent MD (www.attentiondeficit-info.com) and Direction des communications et de la philanthropie, Laval University, with the special collaboration of CADDRA.*





Version : February 2018

Pharmacological treatment for ADHD must be integrated in a multimodal approach and needs to include medical evaluation and follow-up. Comorbid disorders and co-administration of other medications must be taken into account. Here is a brief summary of contraindications and possible drug interactions.

CONTRAINDICATIONS TO PSYCHOSTIMULANTS*

- Treatment with MAO inhibitors and for up to 14 days after discontinuation
- Symptomatic cardiovascular disease
- Glaucoma
- Advanced arteriosclerosis
- Untreated hyperthyroidism
- Known hypersensitivity or allergy to the products
- Acute psychiatric conditions such as mania or psychosis
- Moderate to severe hypertension

Contraindications to Atomoxetine (Strattera)

- Treatment with MAOI and for up to 14 days after discontinuation
- Narrow angle glaucoma
- Uncontrolled hyperthyroidism
- Pheochromocytoma
- Moderate to severe hypertension
- Symptomatic cardiovascular disease
- Severe cardiovascular disorders
- Advanced arteriosclerosis
- Known hypersensitivity or allergy to the products

Contraindications to Guanfacine XR (Intuniv XR)

- Known hypersensitivity or allergy to the products
- · Precautions are advised for those with a history of bradycardia, cardiovascular disease, heart block, hypotension, and syncope.
- * For contraindications to quanfacine XR and atomoxetine hydrochloride, see chapter 5, Canadian ADHD Practice Guidelines 4th Edition, 2018, www.caddra.ca

MAIN POTENTIAL DRUG INTERACTIONS

Psychostimulants

- Monoamine oxidase inhibitors are contraindicated
- SSRIs and SNRIs possible increased risk of serotonin syndrome
- TCAs amphetamines and methylphenidate may interact with TCAs by different mechanisms
- Antipsychotics (e.g. chlorpromazine, fluphenazine) may reduce the effect of amphetamines
- Anticonvulsants methylphenidate may increase the level of phenytoin, primidone and phenobarbital
- Warfarin methylphenidate may increase serum concentrations of warfarin

Atomoxetine (Strattera)

- Monoamine oxidase inhibitors are contraindicated.
- Inhibitors of CYP2D6 (e.g., paroxetine, fluoxetine, bupropion, guinidine) may increase atomoxetine serum concentrations.
- Decongestants (e.g. pseudoephedrine) possible increase in blood pressure and heart rate.

Guanfacine XR (Intuniv XR)

- QT prolonging drugs (e.g. quetiapine, quinidine) since quanfacine XR may cause a decrease in heart rate, concomitant use with QT prolonging drugs is not recommended.
- Beta-blockers may increase risk of rebound hypertensive effect if guanfacine XR is stopped abruptly.
- Anticonvulsants guanfacine XR may
 rerum concentrations of valproic acid. Carbamazepine, phenobarbital and phenytoin may + serum concentrations of guanfacine XR through CYP3A4 induction.
- CYP3A4 inducers or inhibitors (e.g. rifampin, fluconazole, ritonavir) Inducers may + serum concentrations of guanfacine XR Inhibitors may *f* serum concentrations of guanfacine XR.

Additional information: Chapter 5, Canadian ADHD Practice Guidelines 4th Edition, 2018, www.caddra.ca

How can CADDRA help you in your practice?

- . The Canadian ADHD Practice Guidelines: Written and reviewed by a multidisciplinary team of medical experts, the Guidelines provide practical information on how to screen, assess and treat ADHD in children, adolescents and adults.
- ADHD Assessment Toolkit: This is a step-by-step quide to ADHD assessment, provides information on differential diagnosis and comorbid disorders, and includes all required forms and handouts.
- CADDRA eLearning Portal: www.adhdlearning.caddra.ca is a virtual library of resources, including video presentations, podcasts, ePosters and documents on ADHD.

- Education and Training programs: Training on ADHD and comorbid disorders across the lifespan.
- Benefits of becoming a Member: Join a network of health professionals working in the field of ADHD, receive newsletters, updates and notifications, obtain a discount of 20% on the cost of our annual conference; get premium access to our ADHD Learning and receive a printed copy of the Canadian ADHD Practice Guidelines in French or English.
- During our annual conferences, you have an opportunity to hear the top international experts in the field of ADHD speaking on topical subjects. to participate in practical and interactive workshops on ADHD and take part in networking sessions.

www.caddra.ca





Clinicians are invited to refer to the Canadian ADHD Practice Guidelines, 4th edition, for more information on ADHD diagnosis and treatments, www.caddra.ca Document developed by Annick Vincent MD (www.attentiondeficit-info.com) and Direction des communications et de la philanthropie, Laval University.



Patient Name: Date of Birth: Physician Name:

MRN/File No: Date:

SNAP-IV 26 – Teacher and Parent Rating Scale

Name:	Gender:		Age:	
Grade: Ethnicity: 🗆 African-American 🗆 Asian 🗆	Caucasian	🗆 Hispanic	Other:	
Completed by:	_ Type of Class	:	Class size:	
For each item, check the column which best describes this child:	Not At All	Just A Little	Quite A Bit	Very Much
 Often fails to give close attention to details or makes careless mistakes in schoolwork or tasks 				
2. Often has difficulty sustaining attention in tasks or play activities				
3. Often does not seem to listen when spoken to directly				
Often does not follow through on instructions and fails to finish schoolwork, chores, or duties				
5. Often has difficulty organizing tasks and activities				
Often avoids, dislikes, or reluctantly engages in tasks requiring sustained mental effort				
 Often loses things necessary for activities (e.g., toys, school assignments, pencils, or books) 				
8. Often is distracted by extraneous stimuli				
9. Often is forgetful in daily activities				
10. Often fidgets with hands or feet or squirms in seat				
11. Often leaves seat in classroom or in other situations in which remaining seated is expected				
12. Often runs about or climbs excessively in situations in which it is inappropriate				
13. Often has difficulty playing or engaging in leisure activities quietly				
14. Often is "on the go" or often acts as if "driven by a motor"				
15. Often talks excessively				
16. Often blurts out answers before questions have been completed				
17. Often has difficulty awaiting turn				
18. Often interrupts or intrudes on others (e.g. butts into conversations/ games)				
19. Often loses temper				
20. Often argues with adults				
21. Often actively defies or refuses adult requests or rules				
22. Often deliberately does things that annoy other people				
23. Often blames others for his or her mistakes or misbehavior				
24. Often touchy or easily annoyed by others				
25. Often is angry and resentful				
26. Often is spiteful or vindictive				

THE END

Hope you enjoyed the show



Guidelines for ADHD Children and Adolescents

Canadian ADHD Resource Alliance (CADDRA)

www.caddra.ca/pdfs/caddraGuidelines2011.pdf

- American Academy of Paediatrics
 http://pediatrics.aappublications.org/content/early/2011/10/14/peds.2011-2654.full.pdf
- American Academy of Child & Adolescent Psychiatry http://download.journals.elsevierhealth.com/pdfs/journals/0890-8567/PIIS0890856709621821.pdf
- British Association for Psychopharmacology http://www.bap.org.uk/pdfs/ADHD_Guidelines.pdf
- National Institute for Clinical Excellence (NICE) http://guidance.nice.org.uk/CG72

Management of Common Stimulant-induced Adverse Effects

Anorexia, nausea, weight loss

- Administer medication with meals
- Use caloric supplements
- Encourage frequent smaller meals
- Eat early in the morning
- Allow to eat late in the evening (when appetite comes back)
- Encourage healthy snacks
Management of Common Stimulant-induced Adverse Effects

<u>Insomnia</u>

- Administer earlier in day
- Change to short acting
- Discontinue late dosing
- Consider adjunctive treatment (i.e. melatonin)

Management of Common Stimulant-induced Adverse Effects

Rebound phenomena

- Up to 30% of inpatient ADHD youth
- Change to long acting preparation or combine
- Overlap stimulant dosing

Management of Common Stimulant-induced Adverse Effects

<u>Irritability</u>

- Assess timing of phenomena (peak or withdrawal)
- Evaluate comorbid symptoms
- Change preparation
- Consider adjunctive or alternate treatment
- Determine it is not from hypoglycemia (secondary to not eating)

Cardiac Effects of ADHD Medications

• ADHD Drugs and Serious Cardiovascular Events

- Large, retrospective cohort study in children and young adults (2-24yrs) treated with ADHD medication
- No association found between use of certain ADHD medications and adverse cardiovascular events, e.g., stroke, heart attack, and sudden cardiac death
- Medications studied include stimulants (amphetamine products and methylphenidate) and atomoxetine

Recommendations

- Continue to prescribe drugs used for the treatment of ADHD according to the professional prescribing directions
- Stimulant products and atomoxetine should generally not be used in patients with serious heart problems, or for whom an increase in blood pressure or heart rate would be problematic
- Patients treated with ADHD medications should be periodically monitored for changes in heart rate or blood pressure



Results of Multi-modal Treatment Analysis (MTA)

- Medication management alone
 OR
- Medication with behavioral treatment WAS FOUND TO BE
- Nearly equally effective and superior to
 - Behavioral treatment alone
 - Community based treatment

