

Psychiatric Medications: Quick & Dirty review

Terry Broda, Consultant, RN[EC], BScN, NP-PHC, CDDN



RCN document

"Health problems might be accompanied by unusual signs and symptoms, for example someone with severe learning disabilities might demonstrate discomfort by self-injuring."



Challenging Behaviors

- SIB
- Aggression
- Refusals
- Withdrawal or irritability
- Yelling
- "Non-compliance": changes in sleep pattern, appetite, or activity level



Pain Assessment *

Indicators of pain:

- SIB or aggression
- Refusals
- Withdrawal or irritability
- Yelling
- "Non-compliance": changes in sleep pattern, appetite, or activity level
- Denial, inability to communicate or high pain tolerance?



Things that make you say Hmmm...

- Prevalence of mental illness in individuals with developmental disabilities is high.
- Use of psychiatric medication has been reported as approaching 26-40% in community residential placements and 35-50% in institutions in North America.
- Aggression, self-injurious behaviour, over activity, and sleep disturbances are all common.

Mental Illness O'Hara, McCarthy & Bouras, 2010; p.167-169, p.171		
Mental illness	Prevalence	
ADHD	15-60 %	
Anxiety disorders	3-35 %	
Pica	9-25 %	
Mood disorders	4-20 %	
Psychotic disorders	1-4,4 %	
Dementia	8-21 %	
Personality disorders	1-22 %	
OCD	0,2 (community) - 40 % (severe/profound)	
Severe CB	10-22,5 %	
Substance Abuse (ETOH)	Same rate as general population	



Things that make you say Hmmm...

Behavioural changes are often linked to underlying cognitive (thinking) changes, and mood changes occurring in the context of:

- adverse reactions to prescribed medications
- distress arising from a physical illness
- distress arising from mental illness

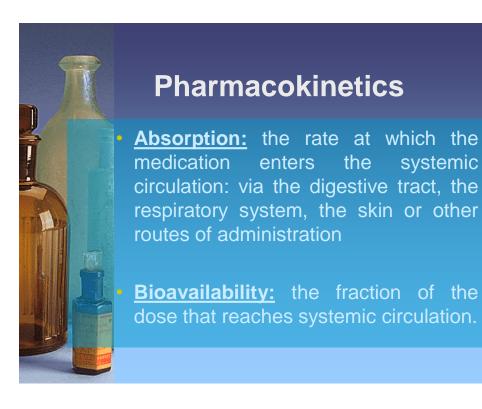


Pharmacokinetics

 Def'n: the study of the metabolism & action of drugs, with particular emphasis on the time required for absorption, duration of action, distribution in the body & method of excretion

(Taber's medical dictionary, 1989)







Absorption:

GI tract:

- Oral
- Sublingual
- Buccal
- Rectal

Respiratory:

- Inhalation
- Intranasal

Integumentary:

- Transdermal
- Subcutaneous
- Intramuscular
- Intravenous

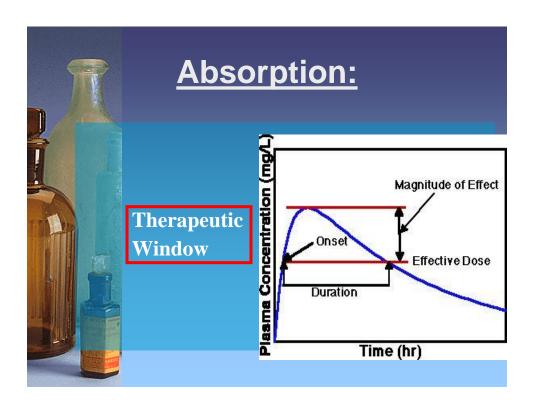
Other:

- Epidural intrathecal
- Intrasynovial, intracardiac
- Eye, ear, vagina, urethra



Psychotropic medications

- **onset of action** time required for medication to have an optimal effect.
- duration of action determines appropriate dosing intervals (minimum time between doses of medication).
- therapeutic range level of medication in the blood & brain achieved over a period of time by prescription of a specific dose of medication. This range is characterized by:
 - a) a therapeutic threshold below which the drug has a suboptimal effect
 - b) a toxic threshold above which adverse effects increase in the absence of any further positive effects







Pharmacokinetics

Distribution: the rate at which a drug moves from the central compartment (blood & highly perfused organs: brain, heart, liver, lungs, kidneys) into the peripheral compartments (muscle, bone, tendons).



Metabolism:

- Hepatic enzymes = Cytochrome P450 system (CYP-450) (about 30 different enzymes).
- Drugs metabolized by an enzyme are substrates of that enzyme



Metabolism:

Possible drug/metabolite interactions:

<u>Competition</u>: substrates compete for same enzyme (2nd substrate can be less 'effective')

<u>Inhibition</u>: blocking enzyme activity (may cause toxicity)

<u>Induction</u>: accelerated metabolism of drugs or their substrates (decreases drug effect as it is metabolized quicker & then eliminated, ex. smoking & clozaril)



Pharmacokinetics

Excretion: the elimination of drugs and their metabolites from the body. Most excretion takes place in the kidney (urine) but excretion also occurs via skin (sweat), lungs (breath) & the digestive (biliary) system (feces).



Pharmacokinetics

Factors that can influence plasma concentration:

- Physiologic factors: age, weight, sex
- Drug-drug interactions (competition for binding at receptor sites, effects of cigarettes & ETOH)
- Patient's health status:
 - especially GI, cardiac, hepatic & renal diseases



Pharmacokinetics

Other factors that can influence plasma concentration:

- Drug absorption variation & presence of food in stomach
- Differences in the pt's ability to metabolize & eliminate the drug (genetics, race)





<u>Definitions of</u> <u>Mood-Stabilizer:</u>

- Substance which is effective for one pole without inducing the other.
- Substance which is effective for both poles of the illness.
- Substance which is effective for both poles of the illness and for prophylaxis of recurrences.



Psychotropic Medication Classes

Mood Stabilizers

- Lithium Carbonate
- Carbamazepine (Tegretol)
- Valproic Acid (Epival, Depakene)
- Lamotrigine (Lamictal)
- Topiramate (Topamax)





Lithium

- Therapeutic Range: 0.7 1.2 mEq/L
- Clearance predominantly through kidneys (95%)
- Dosing adjusted based on renal function
 - Individuals with chronic renal insufficiency must be closely monitored
 - Reabsorption of lithium is increased and toxicity more likely in patients who are hyponatremic or volume depleted (ex. vomiting, diarrhea, diuretics)

Half life

- 12 27 hours
- Increases to 36 hours in elderly persons (**renal function)
- May be considered longer with long-term lithium use (up to 58 hr after one year of therapy)



Lithium Toxicity

- Closely related to concentration of lithium in the blood
 - * Serum concentrations in excess of 2mmol/L
- Preceded by appearance/aggravation of:
 - Sluggishness, drowsiness, lethargy, coarse hand tremor or muscle twitching, loss of appetite, vomiting and diarrhea
 - **repeated episodes of lithium toxicity can cause kidney damage



Lithium Toxicity

Treatment:

- D/C lithium therapy
- Support resp & cardiac functions
- Depending on mental status, use ipecac syrup or gastric lavage
- Follow with charcoal and saline cathartic if multiple ingestion
- Restore fluid and electrolyte balance
- Hemodialysis is treatment of choice when above measures fail



Important considerations:

- 30-50% of persons with DD have epilepsy, so they may be receiving AEDs (Devinsky, 2002)
- Persons with DD may be 3-4 X more likely to have a psychiatric illness (Hellings, 1999)
- Persons with DD are more prone to drug side effects & are also often unable to articulate the effects of the drugs
- 40-60% of persons in general population show inadequate response to mood stabilizer Tx alone & require additional Rx (antipsychotics) (Hellings, 1999)

Bipolar Mood Chart

Client's Name: Month:

Each day, assess the client's mood state for that day by circling the appropriate mood scale item. Your rating should be based on observations for the entire day and evening. If the client is both manic and depressed during the day, carry out separate ratings based upon 12 hour time periods. When completing the log, please use the following anchor points for your mood rating.

- +3 = markedly manic +2 = moderately manic +1 = mildly manic 0 = normal mood for the day
- -1 = mildly depressed -2 = moderately depressed -3 = markedly depressed

Developed by Hurley & Sovner



Antidepressants in Bipolar Depression

- **DO NOT** use in monotherapy
- Combine antidepressant with 1 or more mood stabilizers
- Try first SSRI or buproprion (wellbutrin)
- Beware of noradrenergic drugs and tricyclics
- Discontinue add-on SSRI after response (3-6 months) or may flip into (hypo)mania
- Consider combination with atypica antipsychotics



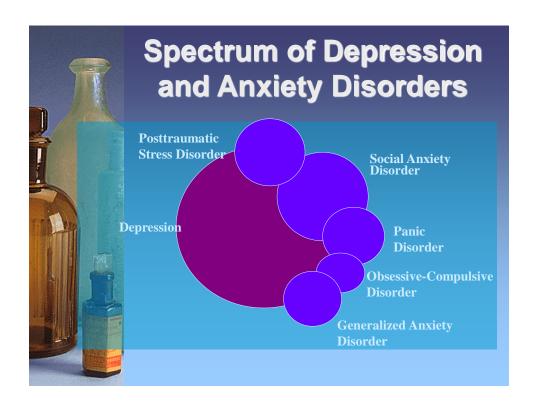


Medication	Systemic/Physical Effects	CNS Effects
Phenobarbital	Rash Sleep problems ↓ Vit D & K Rare: blood dyscrasias, liver toxicity	Sedation, ataxia, dizziness Nystagmus ↓ concentration & cognition Behavior △, irritability (kids)
Phenytoin (Dilantin)	Hirsutism Acne Gingival hyperplasia (50%) ↓ folate/T4/Vitamin D & K levels Rash Osteomalacia ↑ LFTs Blood dyscrasias	Ataxia, dizziness Nystagmus ↓ concentration Sedation Dyskinesia, tremor Arrhythmia N & V, diarrhea
Ethosuximide (Zarontin)	Anorexia <u>Rare:</u> Rash (SJS), blood dyscrasias, behavioral ∆ (kids)	Drowsiness, dizziness Hiccups Headache N & V, diarrhea

Medication	Systemic/Physical Effects	CNS Effects
Clonazepam (Rivotril)	Drooling Rare: Rash Paradoxical anger Thrombocytopenia Depression	Sedation, dizziness Risk of aspiration Paradoxical reaction: disinhibition ↓ concentration Anterograde amnesia Ataxia Nystagmus
Carbamazepine (Tegretol) *CR tab < GI & CNS effects	Pruritic rash ↓ WBC, ↓ Vit D Rare: Aplastic anemia, ↑ LFTs (GGT/ALK), Hyponatremia (SIADH) Cardiac abnormalities ↓ T3/T4/Vit K Alopecia, ocular effects, Osteomalacia	N & V Diplopia Ataxia Sedation, dizziness Dyskinesia Nystagmus

Medication	Systemic/Physical Effects	CNS Effects
Valproic Acid (Depakene) (VPA > GI SE) Divalproex (Epival)	Alopecia Abdominal cramps Hyperammonemia Menstrual disturbances Rare: ↓ platelet & WBC Hepatotoxicity Pancreatitis Low carnitine CAUTION: PCOS Obesity (more common in ♀) *SJS w/ Lamotrigine	Sedation, fatigue Dizziness, ataxia N & V Confusion Headache Tremor
Gabapentin (Neurontin)	Edema Weight gain Rash Behavior ∆, irritability (kids) ↓ WBC Low platelets (rare) ECG changes (rare)	Lethargy, fatigue Dizziness, ataxia Headache N & V Diplopia Tremor Slurred speech

Medication	Systemic/Physical Effects	CNS Effects
Lamotrigine (Lamictal)	**Rash (1st month: gen. red morbilliform) Abdominal pain Alopecia Rare: SJS & toxic epidermal necrolysis Hepatotoxicity Tics in kids	Dizziness, Ataxia N & V Asthenia Headache Lethargy, fatigue Blurred vision, diplopia
Topiramate (Topamax)	Diarrhea Weight loss Kidney stones Glaucoma Rare: ↑ LFTs	Drowsiness, fatigue Headache Dizziness, ataxia Agitation Behavioral Δ Paresthesias (fingers, toes) Cognitive deficits (memory, concentration, wordfinding)







Pharmacotherapy for Obsessive-Compulsive Disorder

- First-Line
 - SSRI's
- Second-line
 - Clomipramine

Considerations

- Higher mean doses
- Delayed onset of response
- Residual symptoms common
- Often long-term (maintenance)

Pharmacologic Management of OCD			
Drug	Dose Range (Frequency	Target Symptoms	Common Adverse Effect
Clomipramine	10-300 mg/d (qhs)	Obsessions, compulsions, ADHD, Nocturnal enuresis	Dry mouth, blurred vision, constipation, sexual dysfunction, orthostatic hypotension
Fluoxetine	10-80 mg/d (qam)	Obsessions, compulsions	Insomnia, nausea, headache, agitation, sexual dysfunction
Fluvoxamine	50-300 mg/d (qhs or bid)	Obsessions, compulsions	As above
Sertraline	50-200 mg/d (qam or bid	Obsessions, compulsions	As above
Paroxetine	10-40 mg/d (qam or bid)	Obsessions, compulsions	As above
Citalopram	10-40 mg/d (qam or bid)	Obsessions, compulsions	As above
		From S	andor P. (1995). Pg. 580



Rationale for Antidepressant Use in Generalized Anxiety Disorder

- GAD is comorbid w/ major depression in 62% of cases
- Clinical goal: treat both anxiety and depression

When you see the anxiety,
don't miss the depression
When you see the depression,
don't miss the anxiety



Psychotropic Medication Classes

Antidepressants (Tx : Panic disorder, OCD, social phobia, bulimia)

Selective serotonin reuptake inhibitors

Fluoxetine (Prozac), paroxetine (Paxil), sertraline (Zoloft), fluvoxamine (Luvox), citalopram (Celexa)

Novel antidepressants

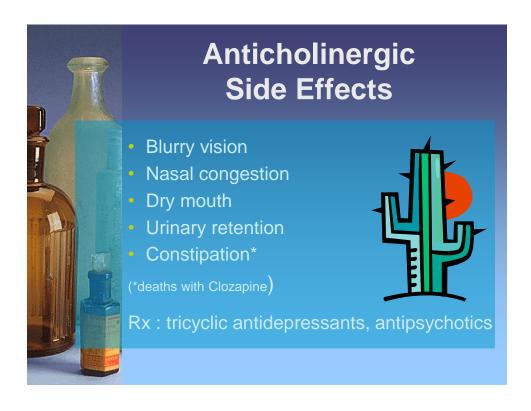
Venlafaxine (Effexor), Nefazodone (Serzone), Moclobemide (Manerix), Bupropion (Wellbutrin)

Tricyclic antidepressants

Amitriptyline (Elavil), Imipramine (Tofranil), Sinequan (Doxepin), Clomipramine (Anafranil)











Psychotropic Medication Classes

Benzodiazepines

Target psychomotor agitation, anxious and fearful affects, and have a calming or sleep-inducing effect

Examples include:

- ·Lorazepam (Ativan),
- Diazepam (Valium),
- Oxazepam (Serax),
- Alprazolam (Xanax),
- •Clonazepam (Rivotril),
- Midazolam (Versed)

Use of Benzodiazepines Useful in many patients but not recommended first-line Use only for short periods of time (less than 4 months) Side effect profile Sedation Reduced coordination Increased risk of falls Impaired cognition Risk of dependency/tolerance Withdrawal symptoms/rebound anxiety **(decrease gradually: 10 - 25% every 1 - 4 weeks)

Benzodiazepines			
<u>Class</u>	<u>Drug</u>		
1. Long half-life (>13hrs) & high potency	Clonazepam (Rivotril) Clobazam (Frisium) (*AED)		
2. Long half-life (>13hrs) & low potency	Chlordiazepoxide (Librium) Diazepam (Valium) Flurazepam (Dalmane) Nitrazepam (Mogadon)		
3. Short half-life (<13hrs) & high potency	Lorazepam (Ativan) Alprazolam (Xanax)		
4. Short half-life (<13hrs) & low potency	Oxazepam (Serax) Temazepam (Restoril)		

Indications for the Use of Benzodiazepines			
ESTABLISHED INDICATIONS:	PROBABLE INDICATIONS:	POSSIBLE INDICATIONS:	
Panic disorderGADSocial phobiaMania/excited schizophrenia	 Adjustment disorder w/ anxiety Acute stress- related insomnia Circadian rhythm disturbances 	•Akathisia •Tourette syndrome •Severely excited states (ER)	



Psychotropic Medication Classes

Antipsychotics

Target psychomotor agitation & aggressive behaviour, particularly in the presence of psychotic symptoms (hallucinations, delusions, and disorganized behaviour)

Traditional

Haloperidol (Haldol), Chlorpromazine (Thorazine/Largactil), Methotrimeprazine (Nozinan), Trifluoperazine (Stelazine), Loxapine (Loxapac)

Atypical

Clozapine (Clozaril), Risperidone (Risperdal), Paliperidone (Invega), Olanzapine (Zyprexa), Quetiapine (Seroquel), Ziprasidone (Zeldox/Geodon), Aripiprazole (Abilify)



	Acute D	ystonia	1
	CI	inical Signs	/Symptoms
Motor Symptoms	Psychological Symptoms	Differential Diagnosis	Risk
Briefly sustained or fixed abnormal movement e.g., torticollis (30%) tongue (25%) trismus/jaw (14.6%) oculogyric crisis (6%)	fearanxiety	malingeringseizurecatatonia	 high potency first- generation antipsychotic s (FGAP) young males first exposure to FGAP



	Akathisia Clinical Signs/Symptoms			
Motor Symptoms	Psychological Symptoms	Differential Diagnosis	Risk	
Foot shiftingPacingRocking	Agitation Restlessness Decreased concentration	Psychotic exacerbation	 High potency first-generation antipsychotics (FGAP) Elderly Female 	

Parkinsonism				
	Clinical Sign	s/Symptoms	5	
Motor Symptoms	Psychological Symptoms	Differential Diagnosis	Risk	
 Tremor Bradykinesia Rigidity Akinesia (masked facies, decreased arm swing) 	Poor concentration attention Bradyphrenia	 Depression Negative symptoms of psychosis 	 High potency first-generation antipsychotics (FGAP) Elderly Female Neurological disorders 	



Tardive Dyskinesia (TD)

Diagnostic Criteria:

History of 3 months total cumulative neuroleptic use

Dyskinesia of lingual-facial-buccal muscle (most common), upper face, limb, trunk

Movements which are repetitive, stereotyped in appearance and distribution

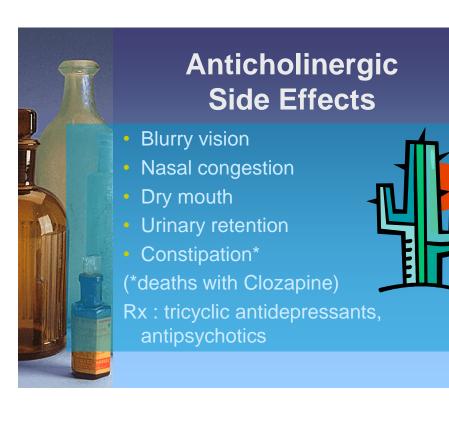
Most common is choreoathetoid movements (classical TD)

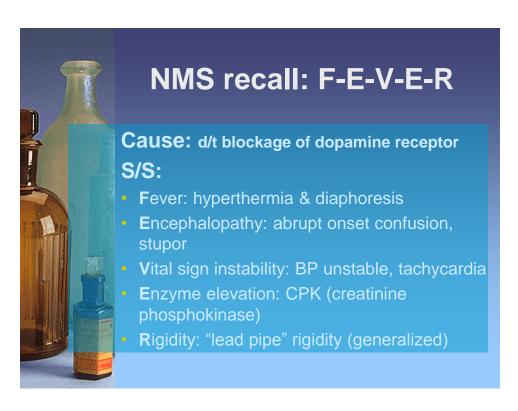
Gait is usually not affected

TD Risk Factors			
Variable	Factor	Determinant of Increased Risk	
Patient Characteristics	AgeGenderDiagnosisPrevious EPSDiabetes	 Increased risk w/ age (>55) Female (slightly higher) Affective disorder Risk 2 to 3 times higher Risk 50-100% higher 	
Drug Characteristics	Type of neurolepticDose/DurationContinuous vs. intermittent	 Typical neuroleptics have similar liability Positive correlation with total drug exposure Higher with intermittent treatment 	

• Abnormal birth • Abnormal
development Neurological disorders Diagnosis of a developmental disability Male, younger age Earlier onset













Side effects – Stimulants

- Nervousness, irritability
- Insomnia
- Anorexia & weight loss (*growth may be effected)
- Headache
- Hypertension, tachycardia
- Tics
- Dry mouth, blurry vision



Strattera: atomoxetine

- Blocks recapture of NE (↑attention, ↓impulsivity, activity)
- With/without meals
- Takes effect in 4 weeks
- No withdrawal symptoms noted
- SE: headache, N & V, abdominal discomfort, anorexia (weight loss), labile mood, fatigue
- Precautions: hypertension, cardiovascular disease, hypotension, liver disorders, glaucoma



Side effects - Strattera

- N & V, abdominal discomfort
- Loss of appetite
- Headache, dizziness
- Insomnia
- Fatigue, lethargy
- Anticholinergic side effects
- Irritability, aggressiveness
- Palpitations
- Sexual dysfunction



Clonidine

- Vs hyperactivity & impulsivity
- Inhibition of noradrenergic transmission
 Dosage :
 - ADHD: 0,05-0,3mg/day
 - Aggression: 0,15-0,4mg/day
 - Anxiety: 0,15-0,5mg/day
- Takes effect in: 30-60 minutes (patch: 2-3jrs)
- Duration: 8 hours (patch: 7 days)
- SE: fatigue, hypotension, vertigo, dermatitis (patch), agitation, depression
- *withdrawal symptoms



Naltrexone

- Opiate Antagonist (blocks the sites)
- Used in severe cases of SIB(& in alcoholism)

SE:

N & V, abdominal discomfort, weight loss, insomnia, anxiety, depression, confusion, fatigue, headache, rare cases of panic attacks.



INDIVIDUALIZED Treatments!

Non pharmacological

- Multimodal approach
- Decrease stress / anxiety:
 - Sensory
 - Environmental modifications
 - Staff support & training
- Communication aids
- CBT , Psychotherapy

Pharmacological

- Antidepressants
- Mood stabilizers
- Benzodiazepines
- Anxiolytics
- Antipsychotics
- Stimulants
- Monitoring side effects!
 O'Hara, et al., 2010, Chapter 16



Monitoring of side effects

- Medication side effect monitoring
- MOSES
- SSRI side effect monitoring tool



Caregiver role

- Observe for particular signs
 - Grimacing
 - Body posturing/positions
 - New posture
 - Change in regular habits/behaviours
- Note observations & tabulate data
 - charts
 - Sleep, food diary, weight
 - Pain scale/checklist
 - Side effects of meds
- Precision!

