

Agents for Medication Induced Movement Disorders

A. FDA Approved Indications in Psychiatry:

1. Extrapyramidal Symptoms: (Benztropine, Amantadine, Trihexyphenidyl, Diphenhydramine)
2. Dystonic Reactions, Acute: Benztropine IM/IV
3. **Tardive Dyskinesia (Valbenazine, Deuterabenazine)**
4. Parkinsonism (Neuroleptic Induced Parkinsonism): Amantadine
5. Insomnia short-term treatment: Diphenhydramine
6. Sedation: Diphenhydramine

B. Non-FDA approved indications commonly used in Psychiatry: (Documentation Required)

1. Parkinsonism: Benztropine, Trihexyphenidyl, Diphenhydramine
2. Akathisia: Propranolol (FDA approved for Essential Tremor), Benzodiazepines

C. Minimal documentation:

1. All standard outpatient & inpatient requirements
2. Document risk versus benefit in context of the following:
 - a) To prevent EPS; antipsychotics should be used judiciously, at minimum effective dose and for the shortest possible duration.
 - b) When EPS presents:
 - I. Reduce the dose if clinically indicated.
 - II. Switch to another agent with lower liability to cause EPS.
 - III. If I & II above fails or clinically inappropriate, then management with anticholinergics may be initiated at the lowest effective dose for a short duration i.e., three months. It is recommended to taper to discontinuation after the management of acute EPS. This is intended to minimize, polypharmacy, risk of side effects including cognitive impairment and the risk of abuse.
 - IV. Anticholinergics worsen Tardive Dyskinesia; therefore, they should be avoided and discontinued if TD develops. **However, anticholinergics are second line if focal dystonia is present (e.g., Cervical dystonia, blepharospasm).** See section E below for more details.
 - V. Prophylactic use is ONLY recommended for patients at high risk for developing EPS i.e., first psychotic episode in young severely ill. When used prophylactically, anticholinergics may be used at the lowest effective dose, short duration, and documentation of the response to above **strategies** i.e., consideration of agents with lower liability to cause EPS.

D. Maximum dosage – see Table 1: Maximum Daily Dose

E. Clinical considerations and Management of new-onset tardive dyskinesia (TD):

Ref.: 2020 APA Practice Guidelines for the Treatment of Patients with Schizophrenia

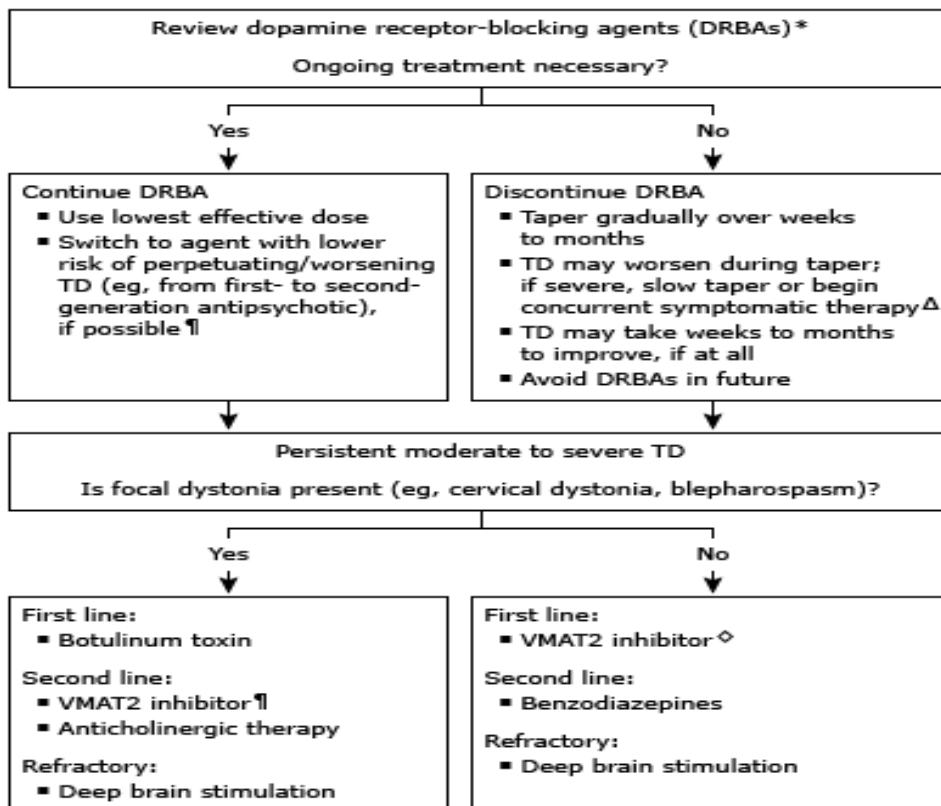
Risk factors for Acute Dystonia:

- Up to 10% of patients on FGA may experience Acute Dystonia vs. <2% with SGA
- Young age
- Males
- Recent Cocaine use
- High medication dose
- IM route of administration

Risk factors for TD:

- TD has been reported after exposure to any of the available antipsychotics.
- 4-8% per year in adults treated with FGAs and the risk is 3x less with SGAs.
- Age>55y.o.
- Females
- White or African race/ethnicity
- Presence of a mood disorder
- Intellectual disability
- CNS injury
- Past or current Akathisia
- Clinically significant parkinsonism
- H/o acute dystonic reactions

Management of New Onset Tardive Dyskinesia (TD)



F. Duration

1. For Outpatient: Document rationale when making any medication change.
2. For Inpatient: Document rationale when making more than 3 changes in any 7-day period.

G. Polypharmacy

Concurrent use of two anticholinergic agents with the same mechanism of action is considered polypharmacy. When considering addition of more than one agent within a class, it is recommended to first titrate the initial agent to maximum tolerated dose; then provide clear supportive rationale for the additional agent(s).

H. Standard laboratory and examination requirements

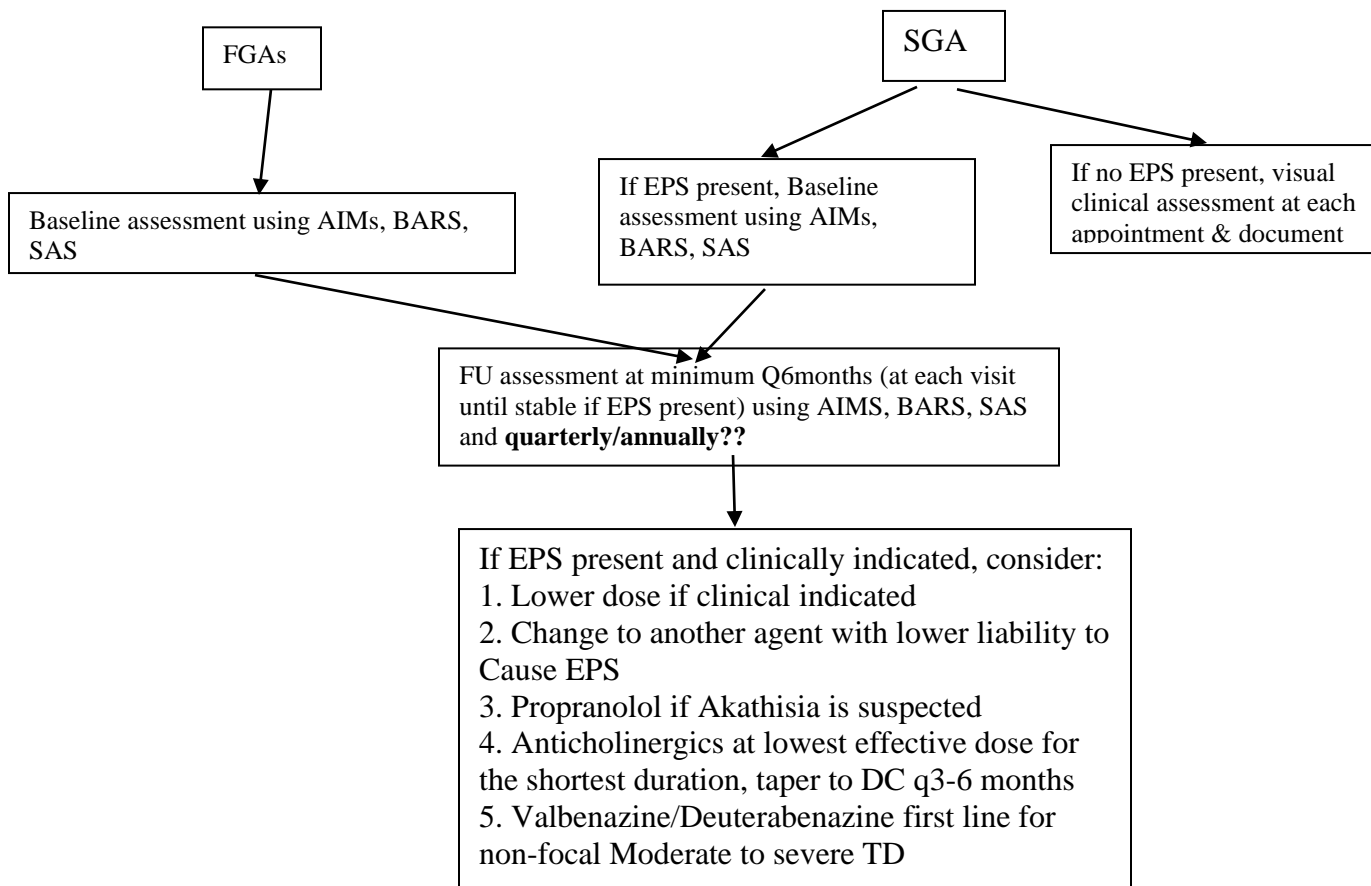
A. For inpatient: Basic laboratory studies on admission

B. For outpatient:

	Initial or baseline assessments	Follow-up assessments
Antipsychotic-induced movement disorders	Clinical assessment of akathisia, dystonia, parkinsonism, and other abnormal involuntary movements, including tardive dyskinesia Assessment with a structured instrument (e.g., AIMS, DISCUS) if such movements are present	Clinical assessment of akathisia, dystonia, parkinsonism, and other abnormal involuntary movements, including tardive dyskinesia, at each visit until stable, then at least q6months Assessment with a structured instrument (e.g., AIMS, DISCUS) at a minimum of every 6 months in patients at high risk of tardive dyskinesia and at least every 12 months in other patients as well as if a new onset or exacerbation of preexisting movements is detected at any visit

Reference: 2020 APA Practice Guideline for the Treatment of Patients with Schizophrenia

Management & Monitoring of EPS



1. Clinical assessments of EPS (using a structured instrument if movements are present) When managing EPS, routine **monitoring** using an appropriate scale is required. **Depending on the symptoms presented; AIMS, BARS or Simson-Angus Scale (SAS) may be used. When anticholinergics are used, monitoring is required at minimum at baseline and annually.**

2. Frequency of Follow-Up: Assessment with a structure instrument at lease every 6 months in patients at high risk of TD and at least every 12 months in other patients (and if new onset or exacerbation of preexisting movements is detected at any visit)

3. When using anticholinergics prophylactically and patient develops EPS: dosage reduction of neuroleptic or switching to a different neuroleptic with lower liability to cause EPS need to be considered while continuing with the anticholinergic agent, increasing its dose or switching to a different anticholinergic agent. In such cases a follow up monitoring by completing one of the above scales is required within three months from the time EPS were detected.

1. Additional monitoring should be considered depending on the clinical situation and whenever there is a change in the patient's status.

I. BBW: Please see PIs for details

J. Warning & Precautions

1. Glaucoma, angle-closure

2. BPH
3. Caution in elderly patients
4. Caution if cardiovascular disease
5. Caution in renal & hepatic disease
6. Caution if high environmental temperature
7. Caution if GI/GU obstruction
8. Caution with alcohol use
9. Caution with Depression
10. **Specific to Velbenazine:**

- A. **Somnolence: May impair patient’s ability to drive or operate hazardous machinery.**
- B. **QT Prolongation: May cause an increase in QT interval. Avoid use in patients with congenital long QY syndrome or with arrhythmias associated with a prolonged QT interval.**

11. Specific to Deuterbenazine;

Deuterbenazine is contraindicated in patients with a h/o depression or prior suicide attempts or ideation. Also contraindicated in patients who are suicidal, and in patients with untreated or inadequately treated depression.

K. Drug-Drug Interactions – Refer to www.epocrates.com for details:

1. Specific to Velbenazine: Dose adjustments due to drug interactions:

Factors	Dosage adjustment for Velbenazine
Use of MAOIs with Velbenazine	Avoid concomitant use with MAOIs
Use of strong CYP3A4 inducers	Concomitant use is not recommended
Use of strong CYP3A4 Inhibitors	Reduce dose to 40mg
Use with strong CYP2D6 inhibitors	Consider dose reduction based on tolerability

2. Specific to Deuterbenazine:

- **Concomitant use of strong CYP2D6 inhibitors: Maximum recommended dose is 36mg per day (18mg twice daily)**
- **Alcohol or other sedation drugs: May have additive sedation and somnolence.**

L. Serious adverse effects:

- Tachycardia
- Psychosis, anticholinergic
- Heat stroke
- Glaucoma
- Neuroleptic Malignant syndrome
- Tardive dyskinesia
- Bradycardia, paradoxical

Amantadine:

- CHF
- Arrhythmia, cardiac arrest
- Respiratory failure, pulmonary edema

- Hematologic (agranulocytosis, neutropenia, leukopenia)
- Seizure
- Oculogyric crisis
- Heat stroke
- Sudden sleep episodes
- NMS-like sx if abrupt DC

Valbenazine:

- QT prolongation
- Parkinsonism
- Hypersensitivity rxn

Deutertrabenazine:

- Suicidality
- Depression
- QT prolongation
- NMS
- Akathisia
- Parkinsonism

M. Common adverse effects:

- Xerostomia
- Constipation
- Urinary retention
- Tachycardia
- Sedation
- GI: N/V/anorexia/flatulence/abdominal pain
- Anxiety

M. Pregnancy and Lactation (Table 2)

Attachments:

Table 1: Maximum Daily Dose

Table 2: Pregnancy and Lactation

Abnormal Involuntary Movement Scale (AIMS)

Barnes Akathisia Rating Scale (BARS)

Simpson-Angus Scale (SAS)

References:

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3. Epocrates & Micromedex
4. WHO 1990 statement: Role of anticholinergic medications in patients requiring long-term antipsychotic treatment for psychotic disorders
5. <http://www.ncbi.nlm.nih.gov/pmc/articles/PMC2306233/pdf/canfamphys00259-0169.pdf>

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7. <http://www.ncbi.nlm.nih.gov/pmc/articles/PMC3325428/>
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10. <http://www.ncbi.nlm.nih.gov/pmc/articles/PMC3018852/> Movement disorders induced by antipsychotic drugs: Implication of the CATIE Schizophrenia Trial
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12. Julia Eve Desmarais, et al, Antinolinergics in the era of atypical antipsychotics: short-term or long-term treatment?, Journal of psychopharmacology, 26(9) 1167-1174, 2012.
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Agents for Medication Induced Movement Disorder

Table 1: Maximum Daily Dose

Agents	Brand	Max Daily Dose	Comments
Amantadine	Symmetrel	300 mg	
Benzotropine	Cogentin	8 mg	
Biperiden	Akineton	16 mg	
Trihexyphenidyl	Artane	15 mg	
Velbenazine	Ingrezza	80mg	<ul style="list-style-type: none"> -Initial dose is 40mg qday. After one week, increase to the recommended 80mg qday. -Can be taken +/-food -Recommended dose for patients with moderate or severe hepatic impairment is 40mg qday -Consider dose reduction based on tolerability in know CYP2D6 poor metabolizers.
Deuterabenazine	Austedo	48mg daily	<ul style="list-style-type: none"> -Titrate weekly by 6mg/day based on reduction of TD and tolerability to max. of 48mg. -Administer total daily dose in two divided doses

References: Epocrates, Micromedex, PIs

**Santa Clara County Department of Mental Health Department Medication Practice Guidelines
Agents for Medication Induced Movement Disorder**

Table 2: Pregnancy Information & Nursing Mother

Agents	Pregnancy Information	Nursing Mother
Amantadine (Symmetrel®)	There are no adequate and well controlled studies in pregnant women. Human data regarding teratogenicity after maternal use of amantadine is scarce. Tetralogy of Fallot and tibial hemimelia (normal karyotype) occurred in an infant exposed to amantadine during the first trimester of pregnancy (100 mg P.O. for 7 days during the 6th and 7th week of gestation). Cardiovascular maldevelopment (single ventricle with pulmonary atresia) was associated with maternal exposure to amantadine (100 mg/d) 9 administered during the first 2 weeks of pregnancy. SYMMETREL should be used during pregnancy only if the potential benefit justifies the potential risk to the embryo or fetus.	SYMMETREL is excreted in human milk. Use is not recommended in nursing mothers.
BENZTROPINE (Cogentin®)	The use of this drug in pregnancy has not been studied.	Lactation: Safety Unknown Inadequate literature available to assess risk, Caution advised.
Biperiden (Akineton®)	Animal reproduction studies have not been conducted with AKINETON. It is also not known whether AKINETON can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. AKINETON should be given to a pregnant woman only if clearly needed.	It is not known whether this drug is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when AKINETON is administered to a nursing woman.
Trihexyphenidyl (Artane®)	Animal reproduction studies to evaluate teratogenic and embryotoxic potential have not been conducted with ARTANE. It is also not known whether ARTANE can cause fetal harm when administered to a pregnant woman or can affect reproduction capacity. ARTANE should be given to a pregnant woman only if clearly needed.	It is not known whether this drug is excreted in human milk. Because many drugs are excreted in human milk, caution should be exercised when ARTANE is administered to a nursing woman. As with other anticholinergics, trihexyphenidyl may cause suppression of lactation. Therefore, trihexyphenidyl should only be used if the expected benefit to the mother outweighs the potential risk to the infant.
Diphenhydramine* (Benadryl®)	Reproduction studies have been performed in rats and rabbits at doses up to 5 times the human dose and have revealed no evidence of impaired fertility or harm to the fetus due to Diphenhydramine hydrochloride. There are, however, no adequate and well-controlled studies in pregnant women. Because animal reproduction studies are not always predictive of human response, this drug should be used during pregnancy only if clearly needed.	Lactation: Probably safe Limited information in animals and/or humans demonstrates no risk/minimal risk of adverse effects to infant/breast milk production, Caution advised.

Propranolol* (Inderal®)	There are no adequate and well-controlled studies in pregnant women. Intrauterine growth retardation, small placentas, and congenital abnormalities have been reported in neonates whose mothers' received propranolol during pregnancy. Neonates whose mothers received propranolol at parturition have exhibited bradycardia, hypoglycemia, and/or respiratory depression. Adequate facilities for monitoring such infants at birth should be available. Inderal should be used during pregnancy only if the potential benefit justifies the potential risk to the fetus.	Propranolol is excreted in human milk. Caution should be exercised when Inderal is administered to a nursing woman.
Velbenazine (Ingrezza®)	May cause fetal harm	Advise not to breastfeed
Deuterabenazine (Austedo®)	Based on animal data, may cause fetal harm	There are no data on the presence of deutetrabenazine or its metabolites in human milk, the effects on the breastfed infant, or the effects of the drug on milk production. The developmental and health benefits of breastfeeding should be considered along with the mother's clinical need for AUSTEDO and any potential adverse effects on the breastfed infant from AUSTEDO or from the underlying maternal condition.

* Off Label

AIMS Examination Procedure

Instructions:

- Should be completed before entering the ratings on the AIMS form.
 - Either before or after completing the Examination Procedure, observe the patient unobtrusively at rest (i.e., in waiting room).
 - The chair to be used in this examination should be a hard, firm one without arms
-
1. Ask patient whether there is anything in his/her mouth (i.e., gum, candy, etc) and if there is, to remove it.
 2. Ask patient about the current condition of his/her teeth. Ask patient if he/she wears dentures. Do teeth or dentures bother patient now?
 3. Ask patient whether he/she notices any movements in mouth, face, hands, or feet. If yes, ask to describe and to what extent they currently bother patient or interfere with his/her activities.
 4. Have patient sit in chair with hands on knees, legs slightly apart, and feet flat on floor. (Look at entire body for movements while in this position).
 5. Ask patient to sit with hands hanging unsupported. If male, between legs, if female, and wearing a dress, hanging over knees. (Observe hands and other body areas.)
 6. Ask patient to open mouth. (Observe tongue at rest within mouth.) Do this twice.
 7. Ask patient to protrude tongue. (Observe abnormalities of tongue movement.)
 8. **Ask patient to tap thumb, with each finger, as rapidly as possible for 10-15 seconds: separately with right hand, then with left hand. (Observe facial and leg movements.)
 9. Flex and extend patient's left and right arms, one at a time. (Note any rigidity and rate it.)
 10. Ask patient to stand up. (Observe in profile. Observe all body areas again, hips included.)
 11. **Ask patient to extend both arms outstretched in front with palms down. (Observe trunk, legs, and mouth.)
 12. **Have patient walk a few paces, turn, and walk back to chair. (Observe hands and gait.) Do this twice.

**Activated movements.

SANTA CLARA COUNTY MENTAL HEALTH

Confidential Patient Information
See Welfare & Institution Code 5328

<p>Patient Name _____ (Last, First, MI) Unicare # _____ Provider _____</p>

ABNORMAL INVOLUNTARY MOVEMENT SCALE (AIMS)

Instructions: Complete Examination Procedure before making ratings.
Code: 0=None, 1=Minimal, may be extreme normal, 2=Mild, 3=Moderate, 4=Severe

MOVEMENT RATINGS: Rate highest severity observed. Rate movements that occur upon activation one less than those observed spontaneously. Circle movement as well as code number that applies.		
Facial and oral movements	1. Muscles of Facial Expression e.g. Movements of forehead, eyebrows, periorbital area, cheeks, including frowning, blinking, smiling, and grimacing.	0 1 2 3 4
	2. Lips and Perioral Area e.g. puckering, pouting, smacking	0 1 2 3 4
	3. Jaw e.g. biting, clenching, chewing, mouth opening, lateral movement	0 1 2 3 4
	4. Tongue Rate only increases in movement both in and out of mouth. NOT inability to sustain movement. Darting in and out of mouth.	0 1 2 3 4
Extremity Movements	5. Upper (arms, wrists, hands, fingers) Include choreic movements (e.g. rapid, objectively purposeless, irregular, complex, serpentine). DO NOT INCLUDE TREMOR (e.g. repetitive, regular, rhythmic)	0 1 2 3 4
	6. Lower (legs, knees, ankles, toes) e.g. lateral knee movement, foot tapping, heel dropping, foot squirming, inversion and eversion of foot.	0 1 2 3 4
Trunk Movements	7. Neck, shoulders, hip e.g. rocking, twisting, squirming, pelvic gyrations	0 1 2 3 4
Global Judgments	8. Severity of abnormal movements overall	0 1 2 3 4
	9. Incapacitation due to abnormal movements	0 1 2 3 4
	10. Patient's awareness of abnormal movements 0=No awareness, 1=Aware, no distress, 2=Aware, mild distress, 3=Aware, moderate distress, 4=Aware, severe distress	0 1 2 3 4
Dental Status	11. Current problems with teeth and/or dentures	No Yes
	12. Are dentures usually worn?	No Yes
	13. Edentia?	No Yes
	14. If known, do movements disappear in sleep?	No Yes NA

Doctor Signature: _____

Date: _____

Name: _____

Date: _____

Barnes Akathisia Rating Scale (BARS)

Instructions: Patient should be observed while they are seated, and then standing while engaged in neutral conversation (for a minimum of two minutes in each position). Symptoms observed in other situations, for example while engaged in activity on the ward, may also be rated. Subsequently, the subjective phenomena should be elicited by direct questioning.

Objective

- 0 Normal, occasional fidgety movements of the limbs
- 1 Presence of characteristic restless movements: shuffling or tramping movements of the legs/feet, or swinging of one leg while sitting, *and/or* rocking from foot to foot or "walking on the spot" when standing, but movements present for less than half the time observed
- 2 Observed phenomena, as described in (1) above, which are present for at least half the observation period
- 3 Patient is constantly engaged in characteristic restless movements, *and/or* has the inability to remain seated or standing without walking or pacing, during the time observed

Subjective

Awareness of restlessness

- 0 Absence of inner restlessness
- 1 Non-specific sense of inner restlessness
- 2 The patient is aware of an inability to keep the legs still, or a desire to move the legs, *and/or* complains of inner restlessness aggravated specifically by being required to stand still
- 3 Awareness of intense compulsion to move most of the time *and/or* reports strong desire to walk or pace most of the time

Distress related to restlessness

- 0 No distress
- 1 Mild
- 2 Moderate
- 3 Severe

Global Clinical Assessment of Akathisia

- 0 *Absent.* No evidence of awareness of restlessness. Observation of characteristic movements of akathisia in the absence of a subjective report of inner restlessness or compulsive desire to move the legs should be classified as pseudoakathisia
- 1 *Questionable.* Non-specific inner tension and fidgety movements
- 2 *Mild akathisia.* Awareness of restlessness in the legs *and/or* inner restlessness worse when required to stand still. Fidgety movements present, but characteristic restless movements of akathisia not necessarily observed. Condition causes little or no distress.
- 3 *Moderate akathisia.* Awareness of restlessness as described for mild akathisia above, combined with characteristic restless movements such as rocking from foot to foot when standing. Patient finds the condition distressing
- 4 *Marked akathisia.* Subjective experience of restlessness includes a compulsive desire to walk or pace. However, the patient is able to remain seated for at least five minutes. The condition is obviously distressing.
- 5 *Severe akathisia.* The patient reports a strong compulsion to pace up and down most of the time. Unable to sit or lie down for more than a few minutes. Constant restlessness which is associated with intense distress and insomnia.

Scoring the Barnes Akathisia Rating Scale (BARS)

The Barnes Akathisia Rating Scale is scored as follows:

Objective Akathisia, Subjective Awareness of Restlessness and Subjective Distress Related to Restlessness are rated on a 4-point scale from 0 – 3 and are summed yielding a total score ranging from 0 to 9.

The Global Clinical Assessment of Akathisia uses a 5-point scale ranging from 0 – 4.

Citation: Barnes TR. A rating scale for drug-induced akathisia. *British Journal of Psychiatry* 1989;154(5):672-676.
This scale can be reproduced freely.

Patient Name: _____ Rater Name and Date: _____

MODIFIED SIMPSON-ANGUS SCALE (MSAS)

Extrapyramidal Side Effects Scale

Each item is rated on a 5-point scale of severity (0 = normal; 4 = most severe; NR = not rated). Circle the rating that best describes the subject's present condition (3 is upper limit for patients without EPS).

1. Gait: The patient is examined as he walks into the examining room: his gait, the swing of his arms, his general posture all form the basis for an overall score for this item. This is rated as follows:

- 0 = Normal
- 1 = Diminution in swing while the subject is walking
- 2 = Marked diminution in swing with obvious rigidity in the arm
- 3 = Stiff gait with arms held rigidly before the abdomen
- 4 = Stooped, shuffling gait with propulsion and retropulsion
- NR = Not ratable

2. Arm Dropping: The patient and the examiner both raise their arms to shoulder height and let them fall to their sides. In a normal subject, a stout slap is heard as the arms hit the sides. In the patient with extreme Parkinson's Syndrome, the arms fall very slowly.

- 0 = Normal, free fall with loud slap and rebound
- 1 = Fall slowed slightly with less audible contact and little rebound
- 2 = Fall slowed, no rebound
- 3 = Marked slowing, no slap at all
- 4 = Arms fall as though against resistance, as though through glue
- NR = Not ratable

3. Shoulder Shaking: The subject's arms are bent at a right angle at the elbow and are taken one at a time by the examiner, who also grasps one hand and also clasps the other around the subject's elbow. The subject's upper arm is pushed to and fro, and the humerus is externally rotated. The degree of resistance from normal to extreme rigidity is scored as follows:

- 0 = Normal
- 1 = Slight stiffness and resistance
- 2 = Moderate stiffness and resistance
- 3 = Marked rigidity with difficulty in passive movement
- 4 = Extreme stiffness and rigidity with almost a frozen joint
- NR = Not ratable

4. Elbow Rigidity: The elbow joints are separately bent at right angles and passively extended and flexed, with the subject's biceps observed and simultaneously palpated. The resistance to this procedure is rated. (The presence of cogwheel rigidity is noted overall but not rated as a separate item.)

- 0 = Normal
- 1 = Slight stiffness and resistance
- 2 = Moderate stiffness and resistance
- 3 = Marked rigidity with difficulty in passive movement
- 4 = Extreme stiffness and rigidity with almost a frozen joint
- NR = Not ratable

5. Wrist Rigidity or Fixation of Position: The wrist is held in one hand and the fingers held by the examiner's other hand, with the wrist moved to extension, flexion, and ulnar and radial deviation, or the extended wrist is allowed to fall under its own weight, or the arm can be grasped above the wrist and shaken to and fro. A "1" score would be a hand that extends easily, falls loosely, or flaps easily upwards and downwards.

- 0 = Normal
- 1 = Slight stiffness and resistance
- 2 = Moderate stiffness and resistance
- 3 = Marked rigidity with difficulty in passive movement
- 4 = Extreme stiffness and rigidity with almost a frozen joint
- NR = Not ratable

6. Head Rotation: The subject sits or stands and is told that the examiner will move his head from side to side, that it will not hurt, and that he should try and relax. (Questions about pain in the cervical area or difficulty in moving his head should be obtained to avoid causing any pain.) Clasp the subject's head between the two hands with the fingers on the back of the neck. Gently rotate the head in a circular motion 3 times and evaluate the muscular resistance to this movement.

- 0 = Loose, no resistance
- 1 = Slight resistance to movement
- 2 = Resistance is apparent and the time of rotation is shortened
- 3 = Resistance is obvious and rotation is slowed
- 4 = Head appears stiff and rotation is difficult to carry out
- NR = Not ratable

Patient Name: _____ Rater Name and Date: _____

7. Glabella Tap: The subject is told to open his eyes and not to blink. The glabella region is tapped at a steady, rapid speed. Note the number of times that the subject blinks in succession. Take care to stand behind the subject so that he does not observe the movement of the tapping finger. A full blink need not be observed; there may be a contraction of the infraorbital muscle producing a twitch each time a stimulus is delivered. Vary the speed of tapping to assure that the muscle contraction is related to the tap.

- 0 = 0 to 5 blinks
- 1 = 6 to 10 blinks
- 2 = 11 to 15 blinks
- 3 = 16 to 20 blinks
- 4 = 21 or more blinks
- NR = Not ratable

8. Tremor: The subject is observed walking into the examining room and then is re-examined for this item with his arms extended at right angles to the body and the fingers spread out as far as possible.

- 0 = Normal
- 1 = Mild finger tremor, obvious to sight and touch
- 2 = Tremor of hand or arm occurring spasmodically
- 3 = Persistent tremor of one or more limbs
- 4 = Whole body tremor
- NR = Not ratable

9. Salivation: The subject is observed while talking and then asked to open his mouth to elevate his tongue.

- 0 = Normal
- 1 = Excess salivation so that drooling takes place if mouth is opened and tongue is raised
- 2 = Excess salivation is present and might occasionally result in difficulty in speaking
- 3 = Speaking with difficulty because of excess drooling
- 4 = Frank drooling
- NR = Not ratable

10. Akathisia: The subject is observed for restlessness. If restlessness is noted, ask, "Do you feel restless or jittery inside; is it difficult to sit still?" Subjective response is not necessary for scoring, but subject report can help make the assessment.

- 0 = No restlessness reported or observed
- 1 = Mild restlessness observed, e.g., occasional jiggling of the foot occurs when the subject is seated
- 2 = Moderate restlessness observed, e.g., on several occasions, the subject jiggles his foot, crosses and uncrosses his legs, or twists a part of the body
- 3 = Restlessness is frequently observed, e.g., the subject's foot or legs are moving most of the time
- 4 = Restlessness persistently observed, e.g., the subject cannot sit still, might get up and walk
- NR = Not ratable

TOTAL SCORE: _____

Total Score Severity:

- Less than 3 = normal
- 3 to 5 = minimal degree of movement disorder
- 6 to 11 = clinically significant degree of movement disorder
- 12 to 17 = severe degree of movement disorder is present

References:

Simpson GM, Angus JWS. A rating scale for extrapyramidal side effects. *Acta Psychiatrica Scandinavica* 1970;212(Suppl 44): 11-19.

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