## MOVEMENT DISORDERS UPDATE

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#### **Movement Disorders**

Hypokinesia : decreased voluntary and automatic movements

Hyperkinesia: excessive movements

#### **HYPOKINESIAS**

- Parkinson's disease
- Secondary Parkinsonism
- Parkinson's plus syndromes

### PARKINSON'S DISEASE Classical Clinical Features

- Resting Tremor
- Cogwheel Rigidity
- Bradykinesia
- Postural Instability

#### **Parkinsonism**

- Consists of at least 2 of the 4 cardinal signs: (mnemonic=TRAP)
- T Tremor
- R Rigidity
- A Akinesia
- P Postural Impairment

#### PARKINSON'S DISEASE

#### Features supporting diagnosis

- Unilateral symptom onset
- Characteristic resting tremor
- Narrow-based gait with flexed/stooped posture
- Reduced arm swing with tremor when walking
- Sustained and significant levodopa effect

AAN Practice Parameter Recommendations: Clinical features distinguishing other parkinsonian syndromes from PD

- Falls at presentation and early in the disease course
- Poor response to levodopa
- Symmetry at onset
- Rapid progression of postural imbalance and dysfunction
- Lack of tremor
- Early dysautonomia

- Parkinson's disease
- Secondary parkinsonism
   Drug-induced parkinsonism
   Vascular parkinsonism
- Parkinson's plus syndromes
   Multiple system atrophy
   Progressive supranuclear palsy
   Corticobasal Degeneration

#### Secondary parkinsonism

- Post-encephalitic parkinsonism (influenza epidemic 1917)
- Dementia with Lewy bodies
- Idiopathic familial basal ganglionic calcification (Fahr Syndrome)
- Late stages of Alzheimer's disease
- Some patients with Frontal Temporal Lobar Dementia

Drug and toxin-induced parkinsonism

- Drugs: Classic antipsychiotic agents and some antiemetic drugs.
- Toxins: Carbon bisulfide, carbon monoxide cyanide, manganese, and some organic solvents.

Vascular parkinsonism

- Small vessel vascular disease
- Mostly parkinson gait-like disorder

#### PARKINSON PLUS SYDROMES

#### Multiple System Atrophy

- Olivopontocerebellar atrophy, striatonigral degeneration and Shy-Drager
- Parkinsonism with associated dysautonomia, cerebellar ataxia, and/or pyramidal signs
- Poor response to levodopa
- Usually well preserved cognitive function

#### PARKINSON PLUS SYNDROMES

#### Corticobasal Degeneration

- Progressive asymmetric onst in one limb associated with akinesia, rigidity, dystonia, focal myoclonus, apraxia, and alien limb phenomenon
- Cognitive impairment common, may be presenting feature

#### PARKINSON PLUS SYNDROMES

#### Progressive supranuclear palsy

- Early gait difficulty with frequent falls
- Progressive loss of ocular motility
- Later dysarthria, dysphagia, rigidity & dementia
- Manikin-frown, procerus sign
- May initially look very much like Parkinson's disease

# Treatment of Parkinson's Disease and Parkinsonian Syndromes

#### DOPAMINERGIC AGENTS

- Dopamine Precursors: Levadopa
- Decarboxylase Inhibitors: Carbidopa
- Dopamine Agonists: Pramipexole (Mirapax) ropinorole(Requip) apomorphine, rotigotine (Neupro)
- Catechol-o-methyltransferase inhibitors: entacapone(Comptan), tolcapone(Fosmar)
- •MAO-B inhibitors: selegilne (Deprenyl) rosagiline(Azilect)
- Dopamine releaser: amantadine (Symmetrel)

### NON-DOPAMINERGIC AGENTS FOR MOTORSYMPTOMS

Antimuscarinics: Trihexyphenidyl (Artane)

benztropine(Cogentin)

#### TREMOR

Involuntary, somewhat rhythmic, muscle contraction and relaxation involving to-and-fro movements (oscillations or twitching) of one or more parts of the body

It is the most common of all movement disorders and can affect all parts of the body

#### **TREMOR**

- Most common movement disorder in the elderly
- Affects men and women equally
- Rhythmic shaking of hands, arms, head, legs, voice
- Dysfunction of muscle control and coordination of agonist & antagonist muscles
- Triggered by or become exaggerated during stress or strong emotion, physical exhaustion, or with certain postures or movements
- Many causes: idiopathic, brain injury, drug-induced, alcohol, toxin (mercury), metabolic (thyroid & liver diseases)

#### **CLASSIFICATION OF TREMORS**

- Resting tremor
  - Parkinsonian tremor
- Action tremor
  - Postural tremor
  - Kinetic tremor
- Psychogenic tremor

#### RESTING TREMOR

- Idiopathic Parkinson's disease
- Pronation/supination
- Slower rate (5-6Hz)
- Present when walking
- Reemergence tremor

#### **ACTION TREMOR**

- Essential tremor
- Physiologic tremor
- Dystonic tremor
- Cerebellar tremor
- Orthostatic tremor
- Task-specific tremor
- Psychogenic tremor

#### **ESSENTIAL TREMOR**

- Present in about 4% of population over 65
- Frequently hereditary, but exact etiology and pathology are unknown
- Affects both sides of body, but usually asymmetric: hands, arms, head, voice (tongue, legs, trunk)
- Typically high frequency tremor involving flexion/extension muscles
- Severity increases with age
- Reduction with alcohol and a positive family history are supportive information
- Treatment with beta-blocker, primidone, botulinum, toxin injection or deep brain stimulation

#### PHYSIOLOGIC TREMOR

- Very fine, high frequency, low amplitude tremor
- Occurs in normal individual & typically without clinical significance
- Enhanced physiologic tremor may heighten Parkinson's tremor to more visible levels
- Caused by strong emotion, physical exhaustion, hypoglycemia, hyperthyroidism, heavy metal poisoning, stimulants, alcohol withdrawal, or fever
- Usually reversible once the cause is corrected

#### DYSTONIC TREMOR

- Occurs in individuals with dystonia: sustained involuntary muscle contractions causing twisting and repetitive motions and/or painful and abnormal postures or positions
- Dystonic head tremor
- Occurs when the individual is in a certain position or moves a certain way
- Occurs irregularly and often relieved by complete rest
- Touching the affected body part or muscle may reduce tremor severity (geste antagoniste)
- Responds well to botulinum toxin treatment

#### CEREBELLAR TREMOR

- Slow, high amplitude, irregular tremor that occurs at the end of a purposeful movement
- Caused by lesion in or damage to the cerebellum and its outgoing nerves by stroke, tumor, multiple sclerosis, degenerative diseases, alcoholism, and certain medications
- Often most prominent when the individual is active or is maintaining a particular posture
- May be accompanied by dysarthria, nystagmus, and gait ataxia

#### ORTHOSTATIC TREMOR

- Rhythmic contractions that occur in the legs and trunk immediately after standing
- Cramps in thighs and legs
- Shakes uncontrollably when the individual is asked to stand in one spot
- No clinical signs or symptoms when the individual sits or is lifted off the ground

#### TASK-SPECIFIC TREMOR

- Also known as focal tremor or occupational tremor
- Occurs mostly in hands when in a certain position or performing a certain task: writing, throwing a ball, bowing the violin, swinging a golf club ("Yips"in golf"),gripping a glass
- May benefit from beta-blocker, anticholinergic, or botulinum toxin injection

#### PSYCHOGENIC TREMOR

- Occurs at rest or during postural or kinetic movements
- Sudden onset and remission, increased incidence with stress
- Bizarre movements that are distractible, variable and inconsistent
- Associated with conversion disorder and psychiatric disease

### SYMPTOMATIC TREATMENT OF ACTION TREMOR

- Pharmacological therapy
  - Beta-blocker (propanolol, Inderal), (primidone Mysoline)
  - alpha-2-delta ligand (gabipentin, Neurontin), pregabalin, Lyrica
- Physical rehabilitation
  - Weighted bracelets, occupational therapy
- Botulinum toxin treatment
  - Intramuscular injection to weaken dominant muscles
- Neurosurgical procedure
  - Deep brain stimulation

#### RESTLESS LEGS SYNDROME

- Very common movement disorder
- Occurrence and intensity increase with age
- 10-12% of adults and >19% in those 80 years or older
- Disagreeable and troublesome sensation in legs
- Diagnostic criteria:
  - Urge to move legs
  - Worsening of symptoms with rest
  - Relief with activity
  - Intensification during the evening

#### RESTLESS LEGS SYNDROME

- Causes anxiety, sleep deprivation, malaise, fatigue
- Associated with iron deficiency, peripheral neuropathy, kidney or liver disease, and offending medications
- Recognizing symptoms and seeking medical attention are important
- Accompanied by periodic limb movements of sleep

#### RESTLESS LEGS SYNDROME

- Treatments include:
  - Removing offending medications:
     SSRI, neuroleptics, Lithium, antihistamines,
     MAOI
  - Iron supplement
  - Avoid idleness
  - Good sleep hygiene
  - Dopamine agonist
  - Opiates
  - Alpha-2-delta ligands

- How To Prevent Augmentation:
  - -Use lowest recommended dose of dopaminergic medications
  - -Maximum doses:
    - a.) pramapaxole (Mirapex), 0.5 mg/24 hrs
    - b.) ropinirole (Requip), 3 mg/24 hrs
    - c.) rotogotine (Neupro), 3 mg/24 hrs

### DRUG INDUCED MOVEMENT DISORDERS

#### DRUG INDUCED PARKINSONISM

- Antipsychotics
- Antiemetics-Metaclopiamide (Reglan), Domperidone (Motilium)
- Dopamine Inhibitors-Deutetrabenazine(Austedo), Reserpine (Serpasil)
- Methyldopa (Aldomet)
- Valproate (Depakote)

#### DRUG INDUCED TREMOR

- Antipsychotics
- Anticonvulsants
- Lithuim
- Tricyclic and SSRI antidepressants
- Beta-Adrenergic agonists
- Theophyline

- Amphetamines
- Thyroid Medications
- Caffeine
- Amiodarone
- Corticosteroids
- Cyclosporine

#### DRUG INDUCED CHOREA

- Antipsychotics
- Antiemetics
- Dopamine agonist
- Levadopa
- Antidepressants
- Anticonvulsants

#### DRUG INDUCED DYSTONIA

- Antipsychotics
- Antiemetics
- Antiparkinson medications

### DRUG INDUCED TARDIVE DYSKINESIA

- Classic oral-buccal-lingual-dyskinesia
- Tardive dystonia
- Tardive akathisia
- Tardive akathitic movements
- Tardive myoclonus

# Treatment Of Tardive Syndromes

- Mainstay of medical treatment is the use of dopamine depleting drugs such as: Reserpine (Serpasil), Deutetrabenazine(Austedo).
- Reserpine treatment may require high doses (5 to 8mg) to reach maximum effects
- Deutetrabenazine has quicker onset and fewer side effects

#### MOVEMENT DISORDERS

#### Summary

- Movement disorders are common.
- Many of them share features of slowness, stiffness, tremor, unsteadiness
- Distinguishing them from normal aging process can be challenging
- Recognizing them and seeking appropriate treatment are important
- Some of them are highly treatable

#### REFERENCES

Adams and Victor's Principles of Neurology, 10<sup>th</sup> Edition

Neurology.org

http://www.neurology.org/