The Johns Hopkins University School of Medicine Psychosis and other unusual behaviors in Parkinson's disease

Presenting: GREG PONTONE, MD, MHS Associate Professor, Departments of Psychiatry and Neurology Johns Hopkins University



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#### Greg Pontone, MD, MHS. DISCLOSURES

> No relevant financial relationships with commercial interests.

The following talk includes unlabeled/unapproved use of medications.



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#### Presentation title

#### "Neuropsychiatric symptoms in Parkinson's Disease"

#### **Today's Goals and Objectives:**

- 1. Become familiar with the prevalence and range of psychotic symptoms in Parkinson's disease
- 2. Discuss the current treatments for psychotic symptoms in Parkinson's disease
- 3. Introduce a range of unusual behaviors in PD such as, impulse control disorders, dopamine dysregulation syndrome, punding, and apathy, and discuss how best to recognize and address them





#### **Psychosis in Parkinson's disease**



#### **Psychosis – a general definition**

 "A severe mental disorder in which contact is lost with external reality"



### **Psychosis – a Parkinson's definition**

- Hallucinations false sensory perceptions in the absence of external stimuli
- Illusions misperception of actual stimuli
- **Passage hallucinations** indefinite object passing through the peripheral visual field
- Sense of presence hallucination a 'feeling' of someone present
- Delusions false, fixed, idiosyncratic belief



### PD psychosis typically occurs in more advanced disease



## Risk factors and correlates of Parkinson's disease psychosis

- Severity and duration of PD
- Cognitive impairment
- Delirium due to medical causes
- Dopaminergic medication for PD
- Alterations in vision (or hearing)
- Anticholinergic, opiate, or benzodiazepine medications



### PD psychosis Management Strategies

- Treat underlying medical illness, if present
- Discontinue medications that may exacerbate psychosis (e.g., pain, bladder, and CNS-acting medications)
- Non-pharmacologic techniques
- Antipsychotic therapy
- Reduction of PD medications
- Treatment with cholinesterase inhibitors



Non-pharmacologic strategies for PD psychosis

- Home modifications (e.g. night lights)
- Discussion, education
- Coping strategies

   -visual techniques
   -cognitive techniques
   -interactive techniques



## Antipsychotic Treatments for PD psychosis

Treatment		Efficacy	Safety	Practice Implications
MDS review designation (2011) <sup>1</sup>	Clozapine	Efficacious	Acceptable risk with specialized monitoring	Clinically useful†
	Quetiapine	Insufficient evidence	Acceptable risk without specialized monitoring	Investigational†
FDA approved for PDP (2016) <sup>2</sup>	Pimavanserin	Efficacious <sup>3</sup>	No impairment of motor function <sup>3</sup> ; increase in QT interval	Clinically useful

<sup>†</sup> Black box warning for typical and atypical APs in elderly patients who have dementia-related psychosis<sup>4</sup>

1. Seppi K, et al. *Mov Disord*. 2011;26(suppl 3):S42-S80. 2. US Food and Drug Administration Web site.

http://www.fda.gov/NewsEvents/Newsroom/PressAnnouncements/ucm498442.htm. Published April 29, 2016. Accessed October 27, 2016. **3.** Cummings J, et al. *Lancet.* 2014;383(9916):533-540. **4.** US Food and Drug Administration Web site. http://www.fda.gov/Drugs/DrugSafety/ucm124830.htm. Published NS HOPKIN June 16, 2008. Accessed November 5, 2016.

### Cholinesterase Inhibitors for Parkinson's disease psychosis

- Emergence of hallucinations may coincide with onset of dementia
- Cholinesterase inhibitors often used for PDP because of its association with dementia<sup>1,2</sup>
- Agents investigated (all off-label) include:
  - Galantamine
  - Rivastigmine
  - Donepezil





### IMPULSE CONTROL DISORDERS IN PARKINSON' S DISEASE

#### **Impulse control disorders in PD**

"Behaviors performed repetitively, excessively, and compulsively to an extent that interferes with life functioning" Impulsive – lack of forethought or consideration of consequences Compulsive – repetitious behaviors with lack of self-control



### Impulse control disorders in PD

- Pathological gambling
- Compulsive buying/shopping
- Hypersexual behaviors
- Binge eating



# Impulse control disorders in PD are often a hidden problem

- Routine screening is not common
- Patients are often:
  - -embarrassed by the behaviors
  - -have limited awareness of behaviors
  - -don't recognize association with PD or PD medications
- Caregivers/spouses/family bring to clinical attention often only *after* consequences are recognized



## Risk factors for impulse control disorders in PD

- Dopamine agonists
- to a lesser extent L-dopa, amantadine and MAOIs e.g. rasagiline
- Men sexual ICDs, Women eating, shopping/buying
- Younger and young onset PD, dyskinesias
- Depression and anxiety



### Dopamine dysregulation syndrome

- Drug addiction-like state marked by selfmedication with inappropriately high doses of dopaminergic medications
- May be more common in early onset PD and males
- Co-occurs with ICDs, psychosis, panic attacks



### Punding

- Repetitive, purposeless behaviors, characterized by an intense preoccupation with specific items or activities – collecting, arranging, or taking apart objects
- Hobbyism higher level repetitive behaviors, e.g. excessive internet use, reading, art work, work on projects





# DISEASE

# **APATHY IN PARKINSON'S**

#### **Apathy in Parkinson's disease**

- May affect more than 1/3 of persons with PD
- Associated with more severe motor symptoms and cognitive impairment; more likely to be men and older age



### Apathy in Parkinson's disease

3+ symptoms lasting more than one month and impacting function

- Reduced initiative and decreased self-driven ideas
- Decreased curiosity and spontaneity
- Difficulty continuing activities to completion
- Indifference or blunted emotional reactions
- Lack of concern about personal problems
- Lack of affectionate behavior



#### Apathy vs depression in PD

Apathetic symptoms Reduced initiative Decreased participation in external activities unless engaged by another person Loss of interest in social events or everyday activities Decreased interest in starting new activities Decreased interest in the world around him or her Emotional indifference Diminished emotional reactivity Less affection than usual Lack of concern for others' feelings or interests

Overlapping symptoms Psychomotor retardation Anhedonia Anergia Less physical activity than usual Decreased enthusiasm about usual interests Emotional symptoms of depression Sadness Feelings of guilt Negative thoughts and feelings Helplessness Hopelessness Pessimism Self-criticism Anxiety Suicidal ideation



### Management of apathy in PD

Non-pharmacological interventions

 -scheduled activities (social and physical)
 -establish clear and achievable goals
 -rewards conditional on completion of goals

-recruit social support for activities

• Dopamine agonists, stimulants, and antidepressants can be tried







