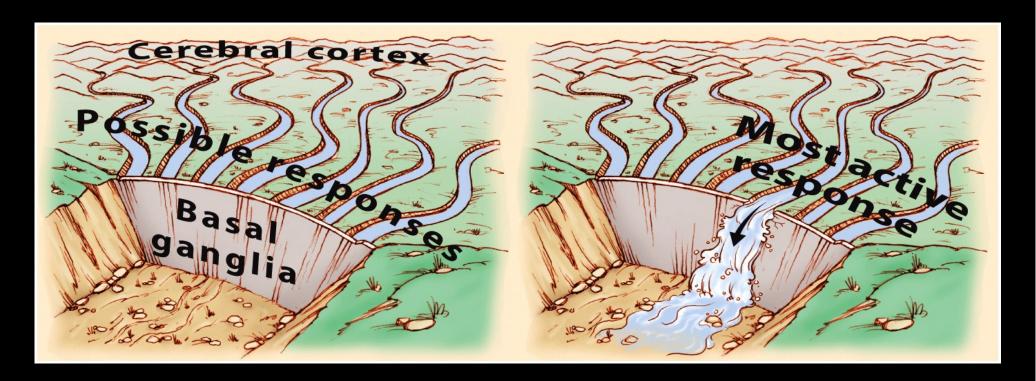
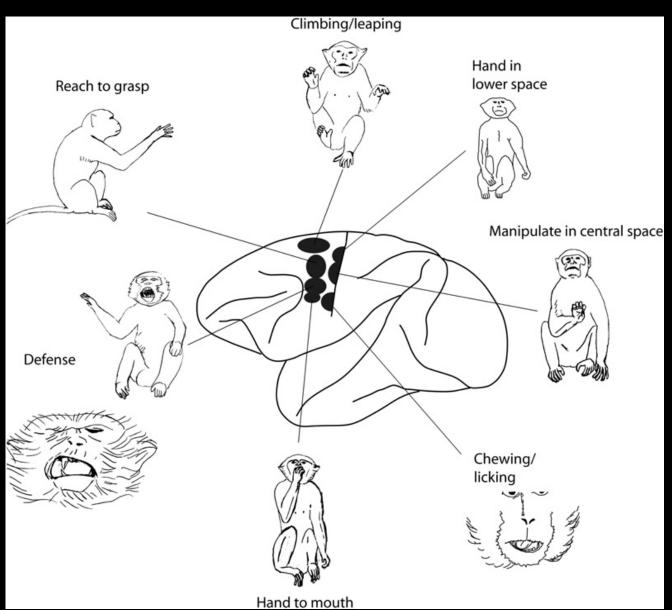
### Integrated Motor Systems

Michael S. Beauchamp, Ph.D.
Department of Neurobiology and Anatomy
University of Texas Health Science Center at
Houston
Houston, TX

Michael.S.Beauchamp@uth.tmc.edu



### **Action Zones**

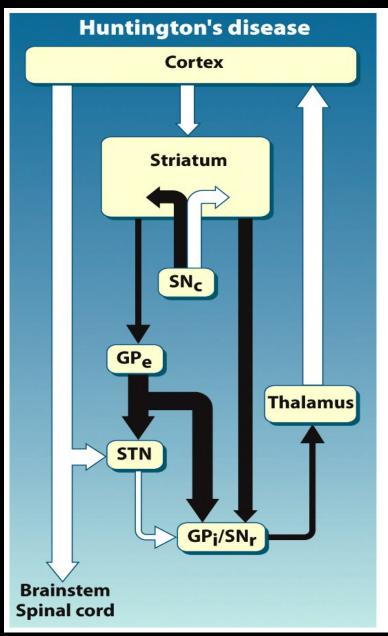


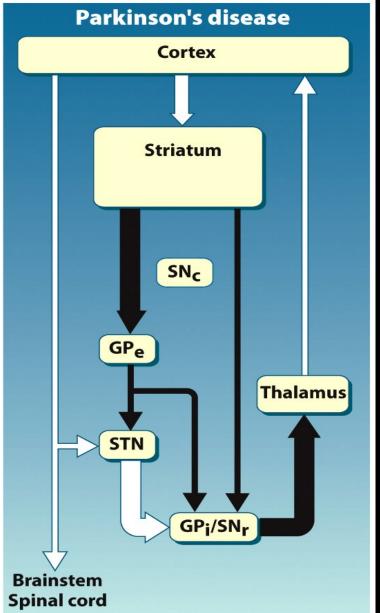
Aflalo & Graziano, *Neuron,* 2007



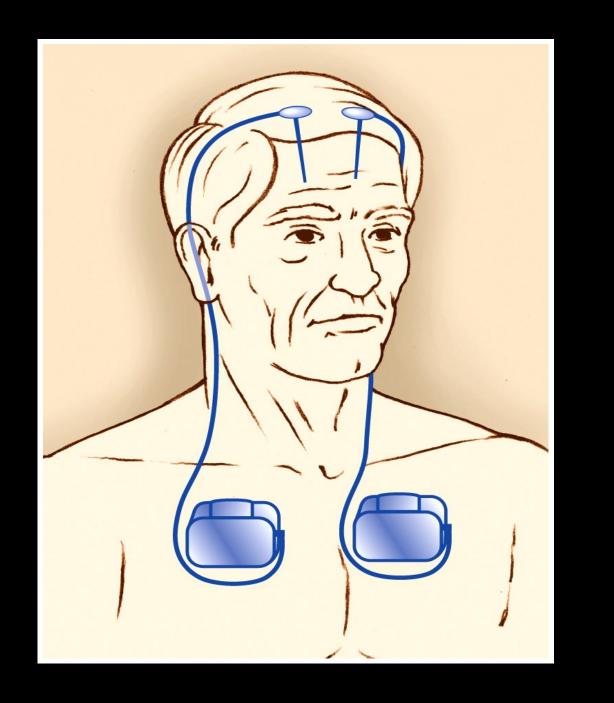
# **Disorders of Basal Ganglia**

- 1. Huntington's Disease
- 2. Parkinson's Disease





(b) Parkinsonian subject (a) Healthy subject



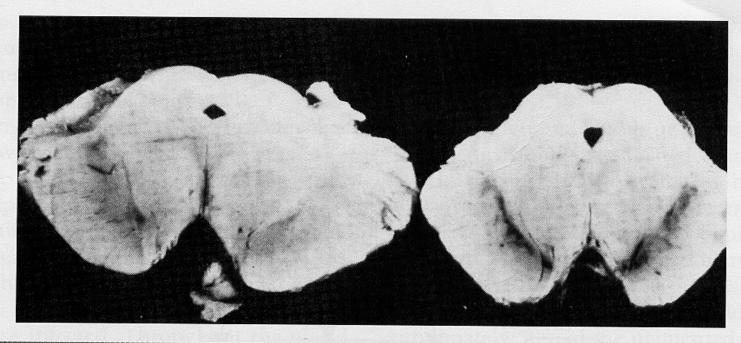
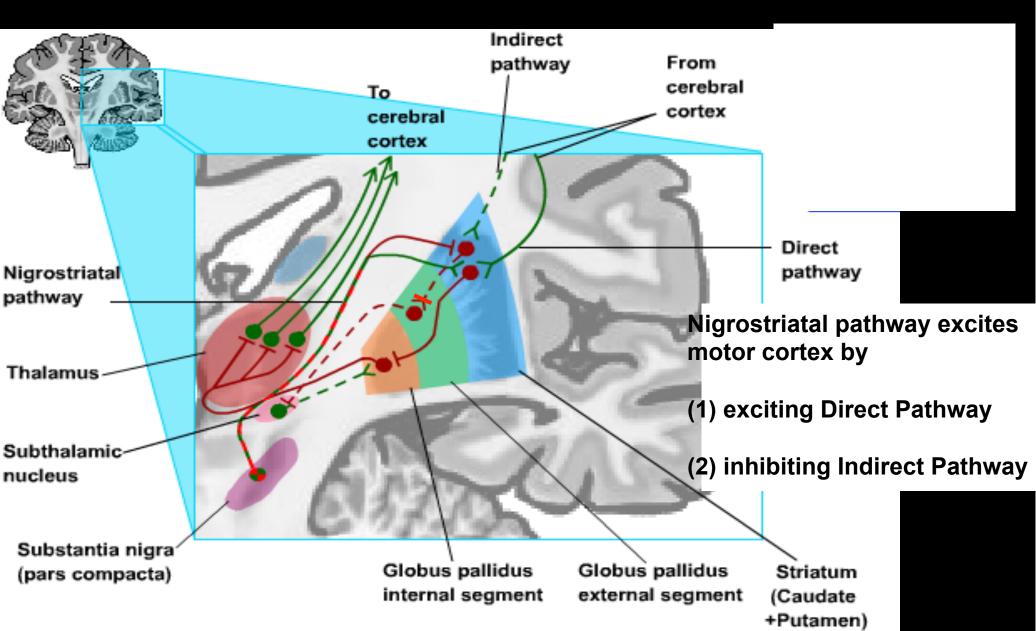
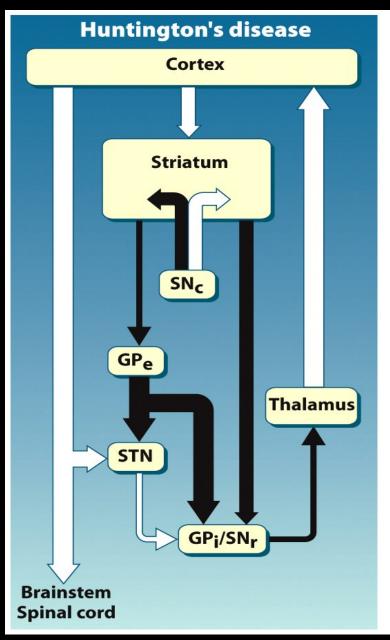
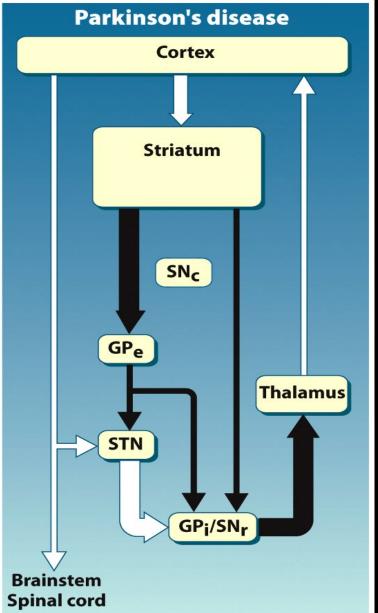


Figure 8.13 Parkinson's disease pathology

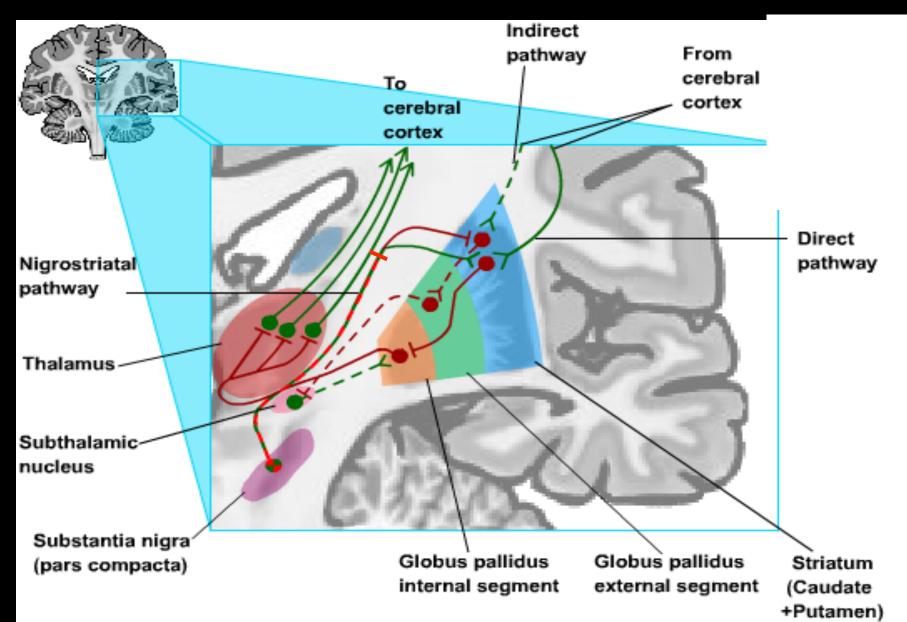
### Dopaminergic Input from Substantia Nigra



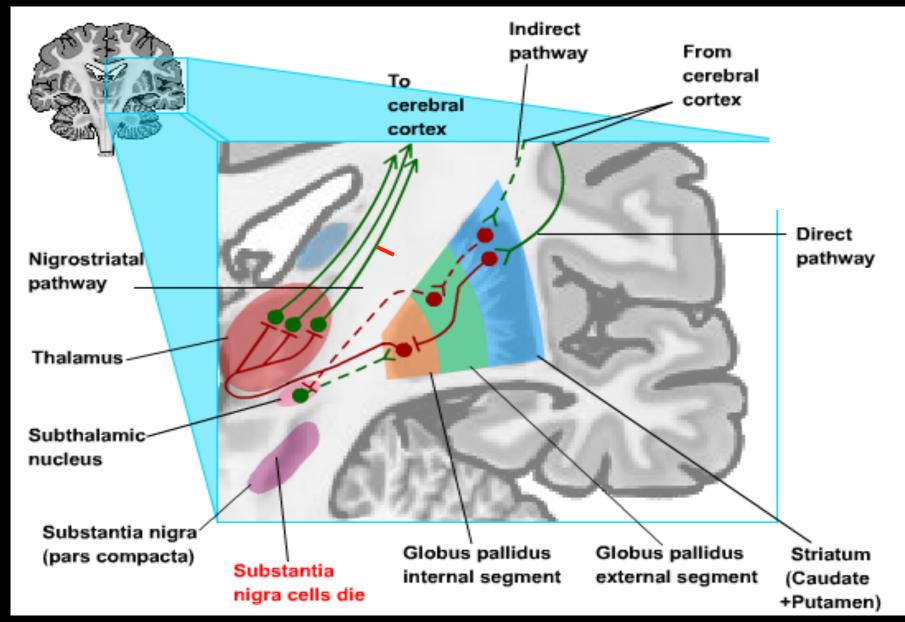




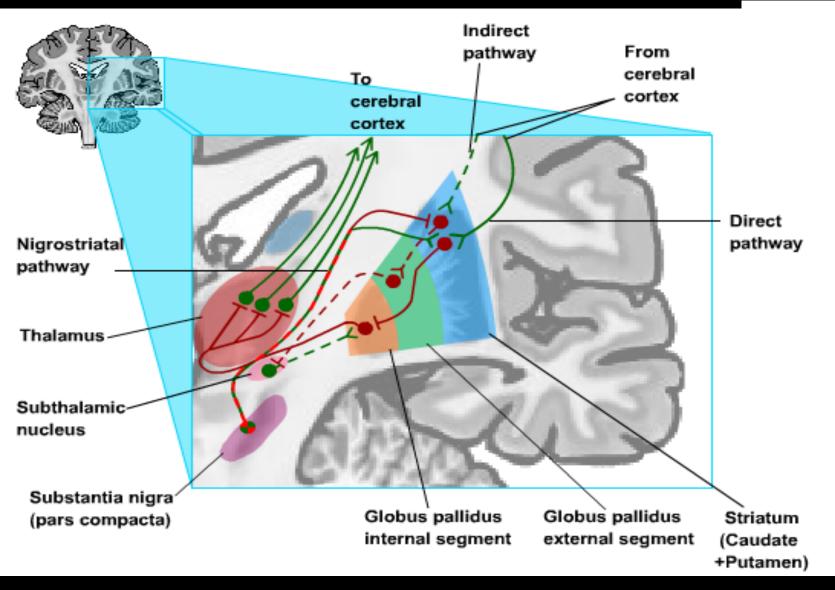
### Parkinson's Disease



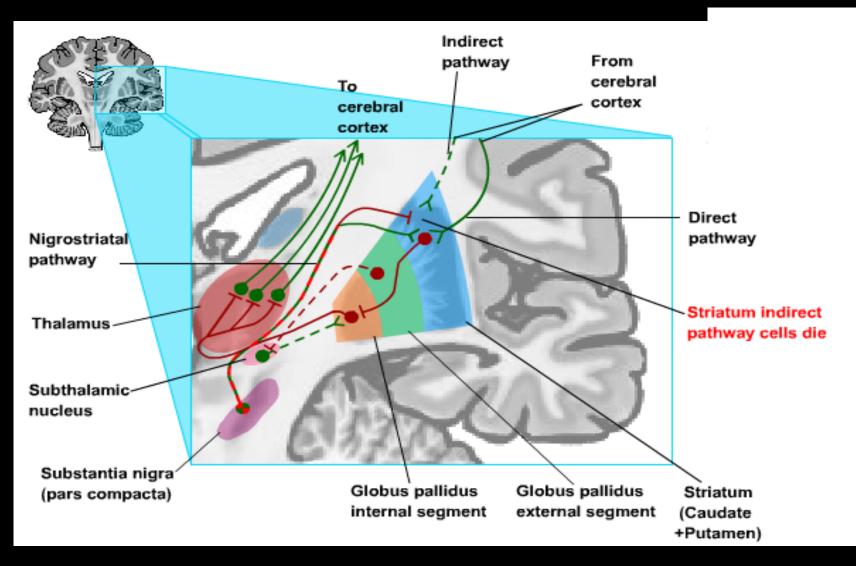
## Parkinson's Disease

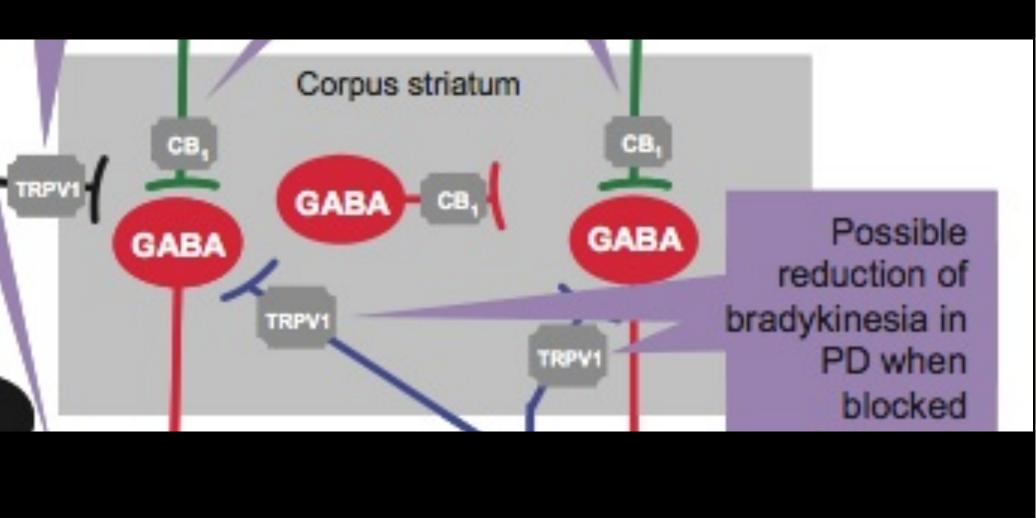


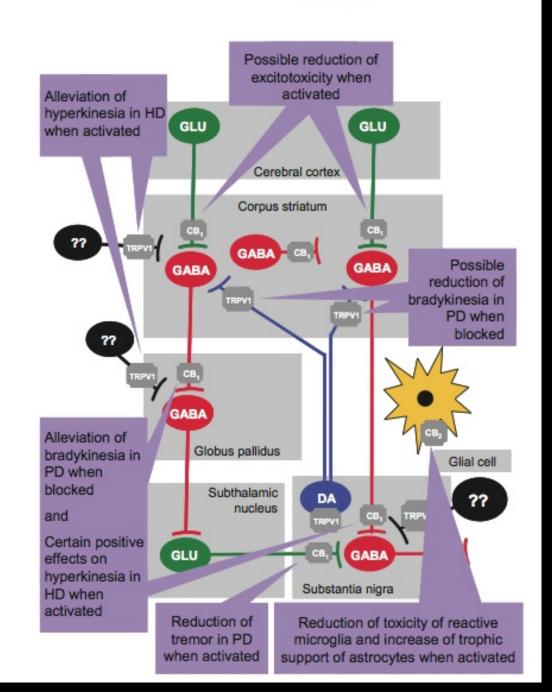
# **Huntington's Disease**

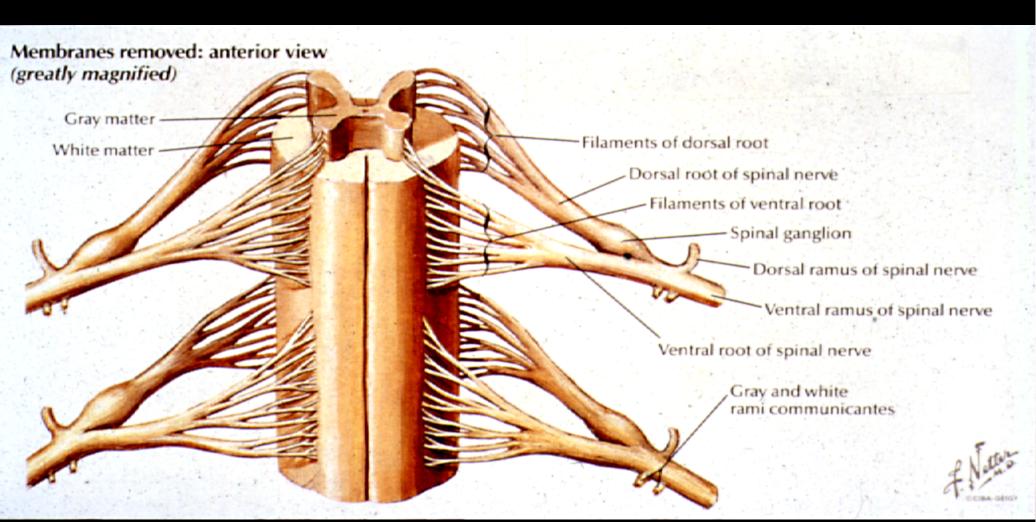


# **Huntington's Disease**

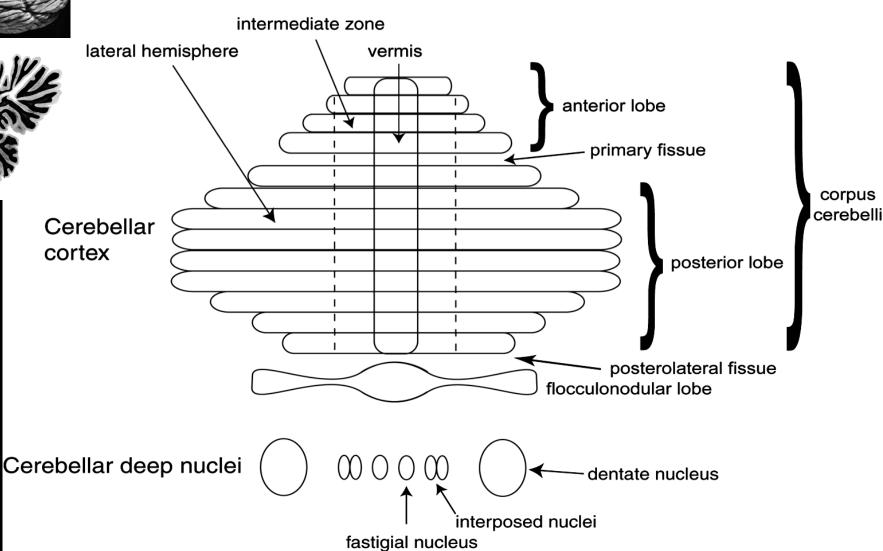








#### **Gross Anatomical Organization of Cerebellum**



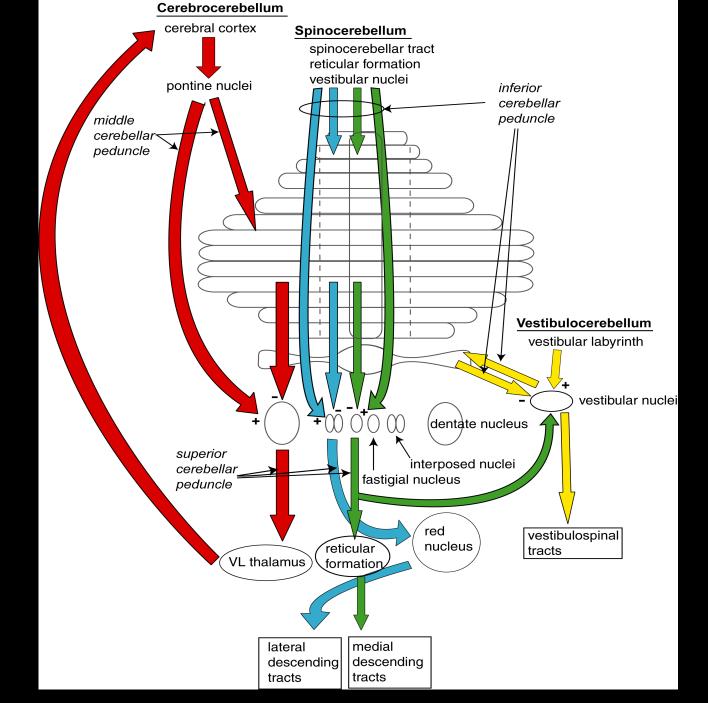


Also Lateral Vestibular

**Fastigial** 

Interposed

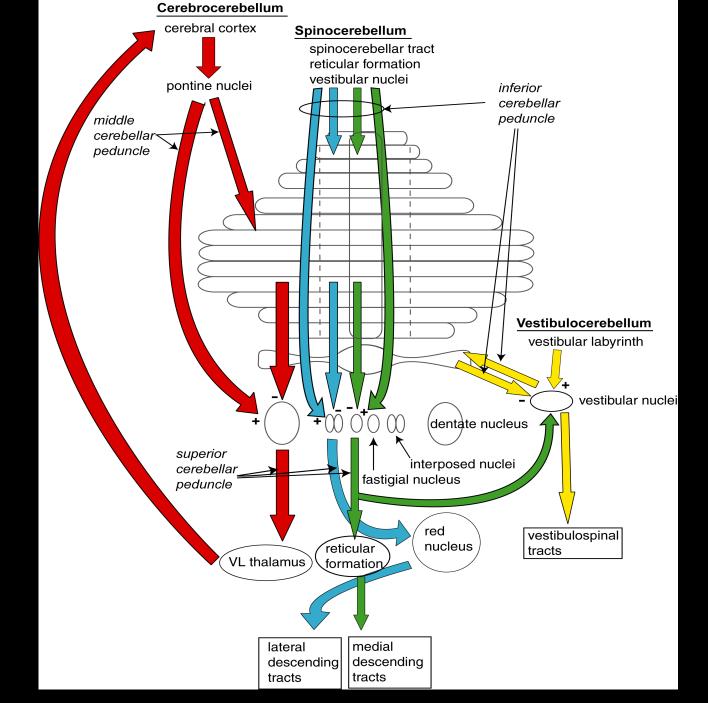
**Dentate** 





#### **Fastigial Nucleus**

- inhibitory input from cerebellar cortex: vermis
- excitatory input: vestibular, proximal somatosensory, auditory, visual
- projects to vestibular nuclei, reticular formation

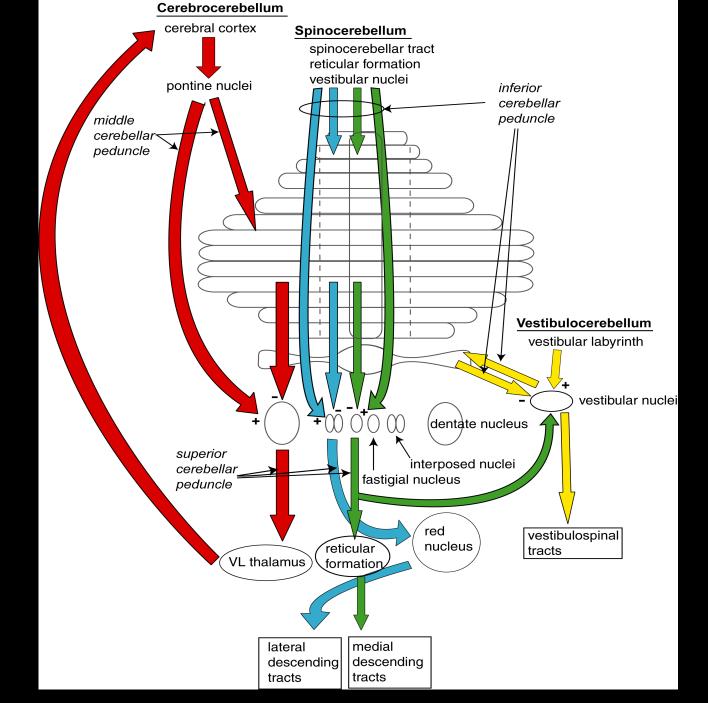


#### **Fastigial Nucleus**

- inhibitory input from cerebellar cortex : vermis
- excitatory input: vestibular, proximal somatosensory, auditory, visual
- projects to <u>vestibular nuclei, reticular formation</u>

#### Interposed

- inhibitory input from cerebellar cortex : intermediate zone
- excitatory input: spinal, proximal somatosensory, auditory, visual
- projects to <u>contralateral red nucleus</u>



#### **Fastigial Nucleus**

- inhibitory input from cerebellar cortex : vermis
- excitatory input: vestibular, proximal somatosensory, auditory, visual
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#### Interposed

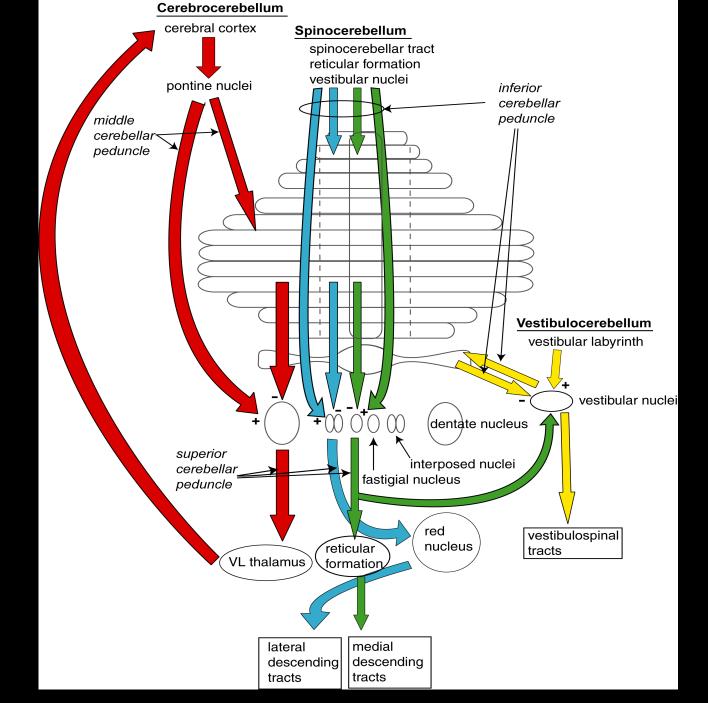
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- excitatory input: spinal, proximal somatosensory, auditory, visual
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#### **Dentate**

- inhibitory input from cerebellar cortex: <u>lateral hemisphere</u>
- excitatory input: cortex via pontine nuclei
- projects to contralateral red nucleus, thalamus (VL)



DeArmond Fig. 81



#### **Fastigial Nucleus**

- inhibitory input from cerebellar cortex : vermis
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- projects to <u>vestibular nuclei, reticular formation</u>

#### **Interposed**

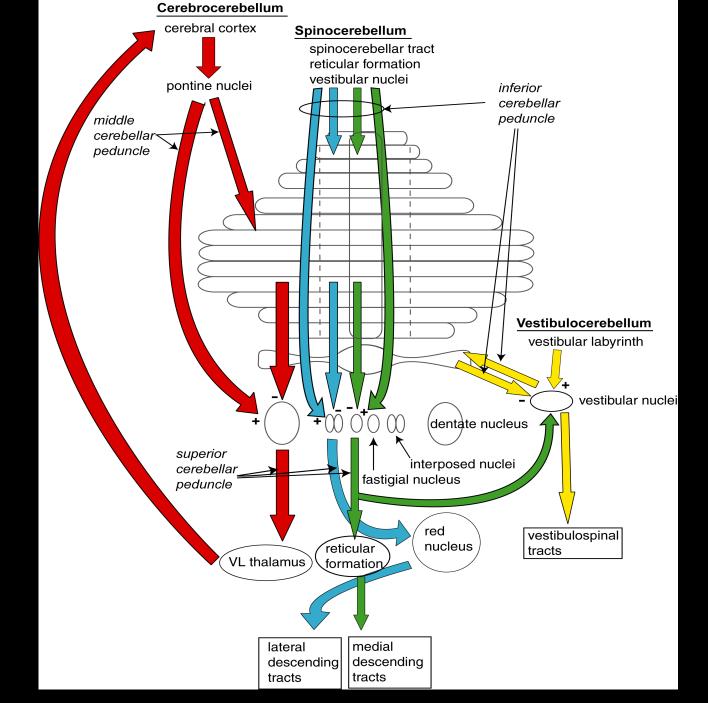
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- excitatory input: spinal, proximal somatosensory, auditory, visual
- projects to <u>contralateral red nucleus</u>

#### **Dentate**

- inhibitory input from cerebellar cortex: <u>lateral hemisphere</u>
- excitatory input: cortex via pontine nuclei
- projects to contralateral red nucleus, thalamus (VL)

#### Vestibular

- inhibitory input from cerebellar cortex : flocculonodular lobe
- excitatory input: vestibular labyrinth
- projects to motor nuclei



#### **Cerebellar Peduncles**

Input and output tracts of cerebellum

Inferior Cerebellar Peduncle (restiform body)
Primarily afferent fibers from medulla

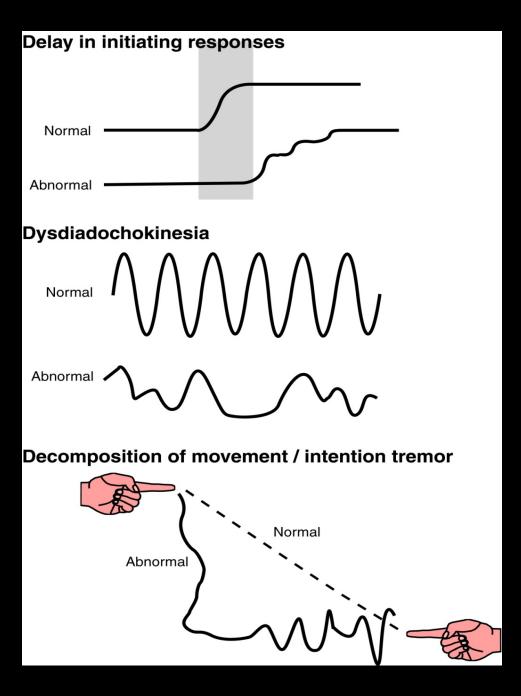
Middle Cerebellar Peduncle (brachium pontis)
Primarily afferent fibers from pons

Superior Cerebellar Peduncle (brachium conjunctivum)

Primarily efferent fibers from the cerebellar nuclei

Cerebellum controls the IPSILATERAL side of the body

# **Cerebellum Produces Movement Disorders**



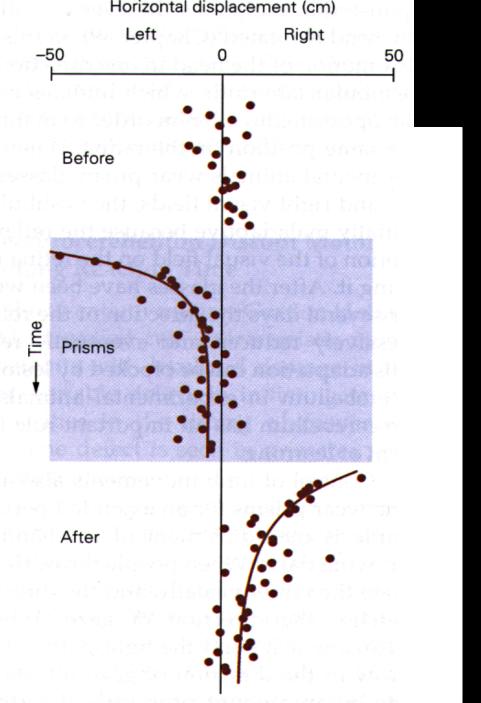
### Cerebellum and Motor Learning

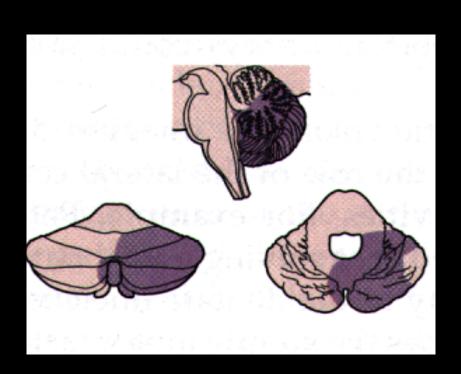
Vestibulo-ocular reflex

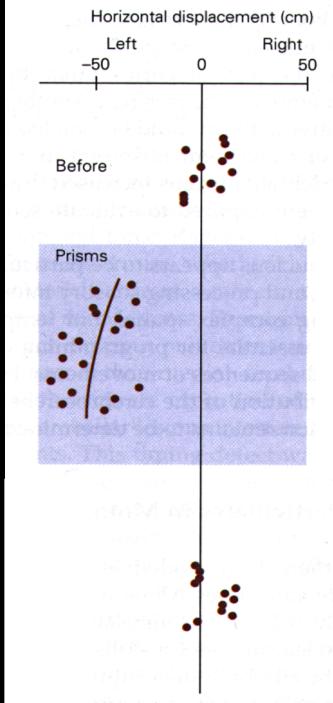
**Coordination of movements** 

**Pavlovian Classical Conditioning** 







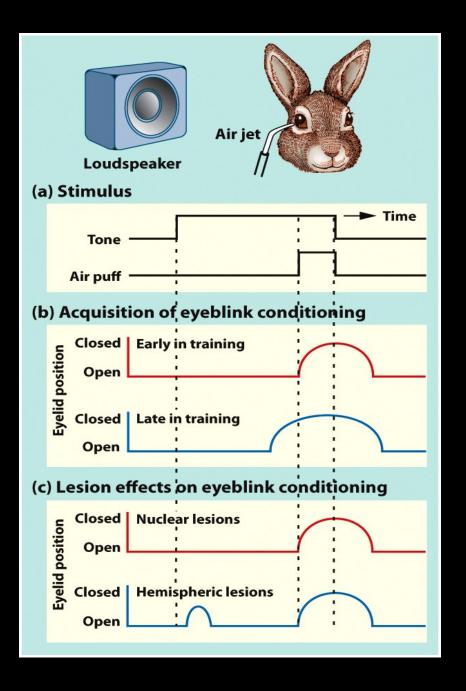


#### Cerebellum and Motor Learning

Vestibulo-ocular reflex

**Coordination of movements** 

**Pavlovian Classical Conditioning** 

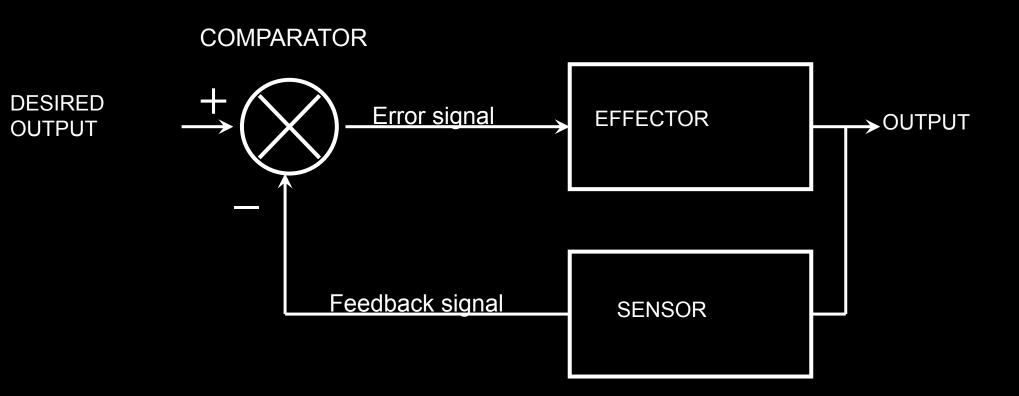


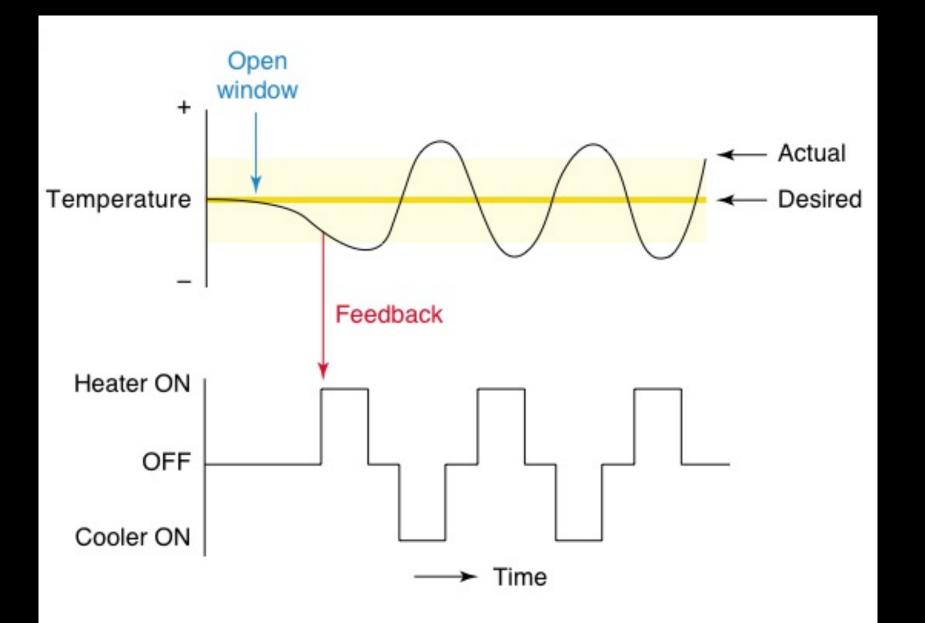
#### Cerebellum as a Control System

Feedback Controller for Slow movements (e.g., posture)

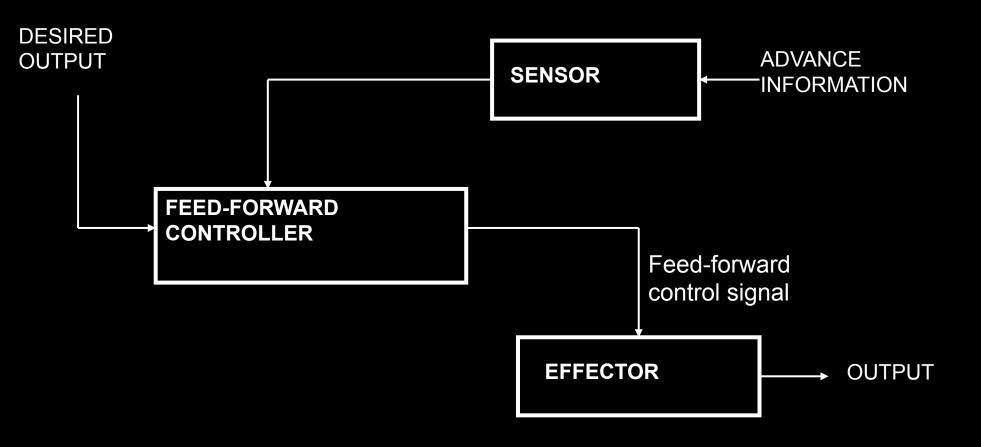
Feedforward Controller for Fast movements (e.g., most voluntary movements)

## Feedback control system

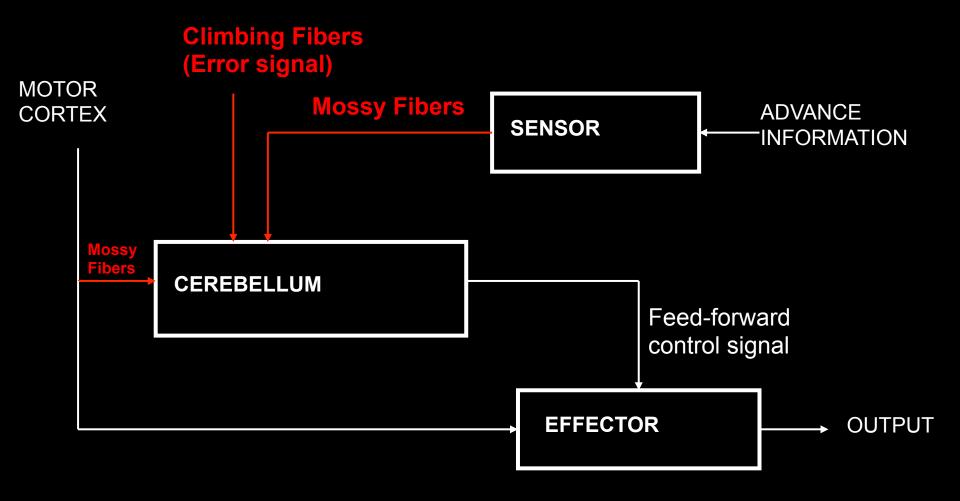




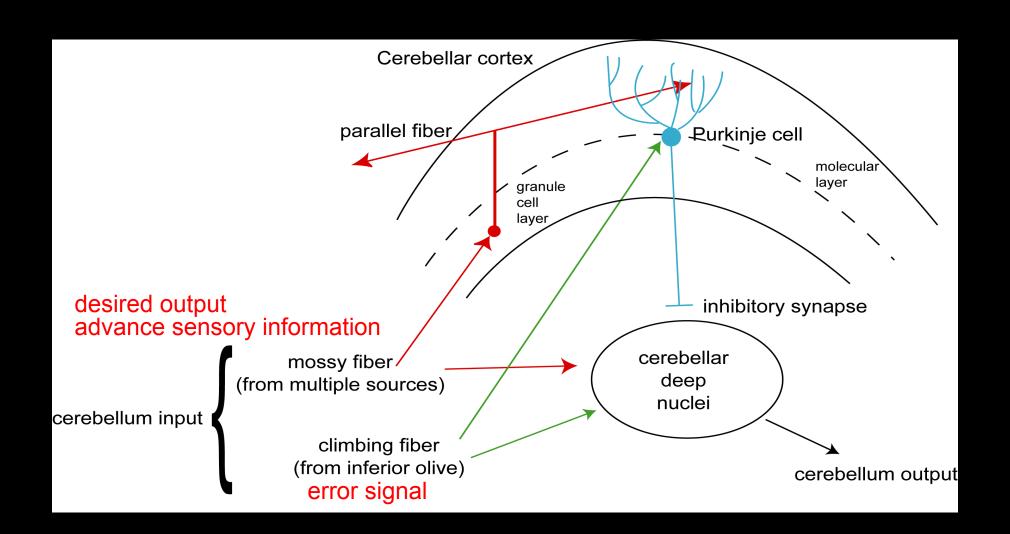
#### Feedforward control system



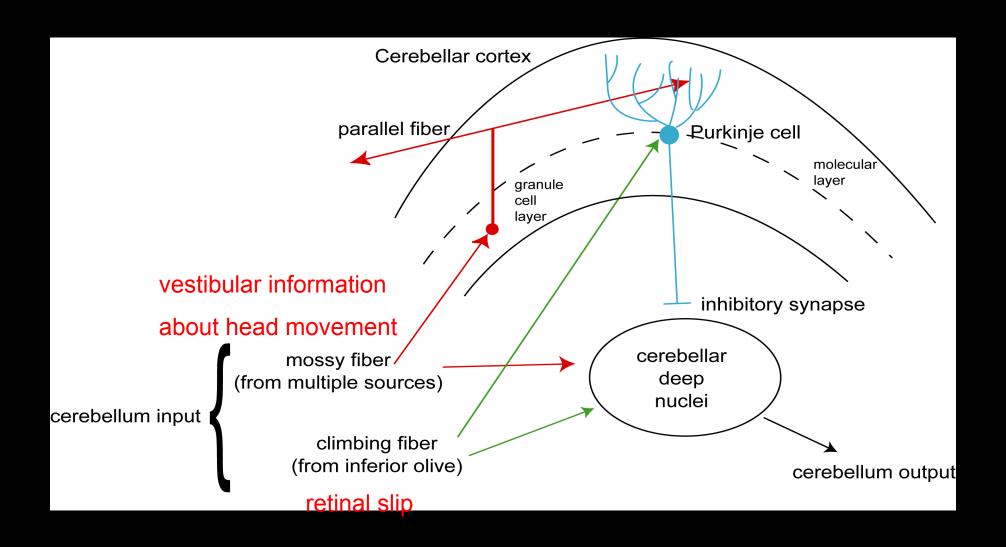
#### Cerebellum as a feedforward control system



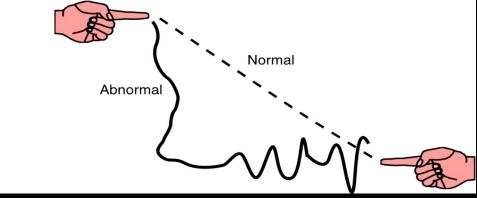
#### **Internal Circuitry of Cerebellum**



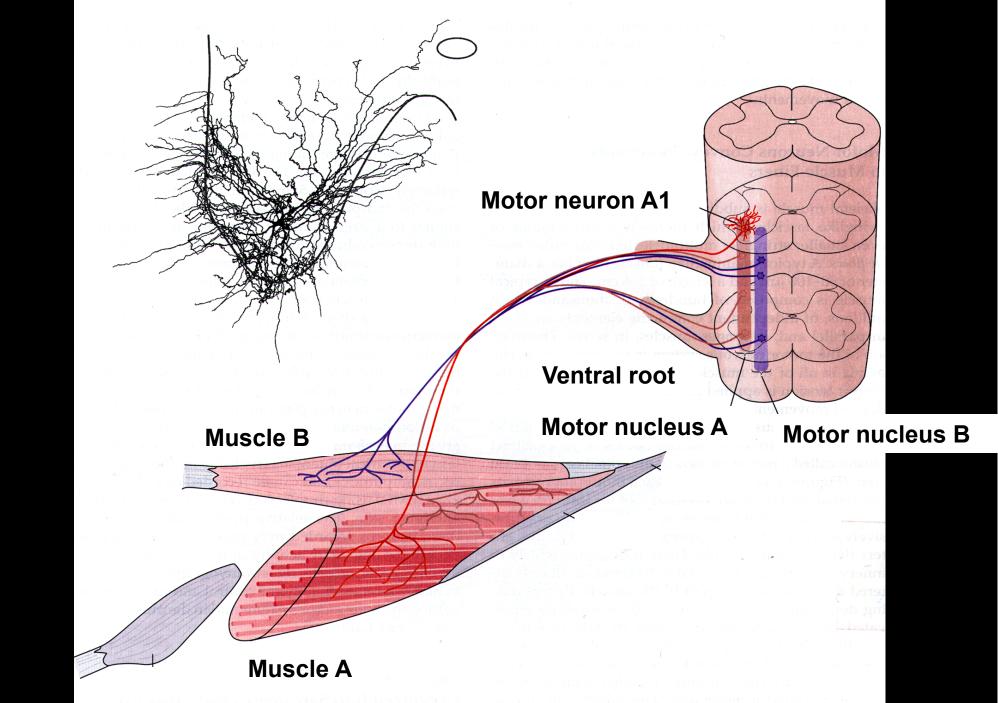
#### **VOR and feedforward control**



## Delay in initiating responses Normal Abnormal Dysdiadochokinesia **Abnormal** Decomposition of movement / intention tremor Abnormal



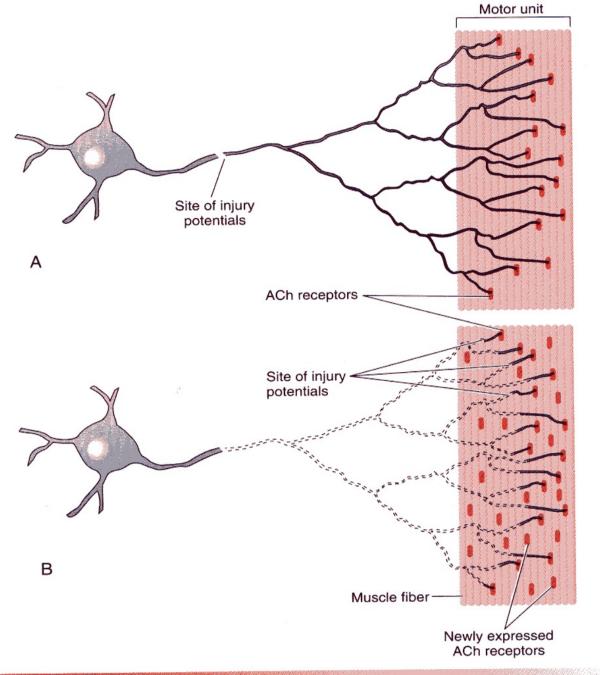
- Effects limited to small groups of muscles
- Muscle atrophy
- Weakness
- Fasciculation/fibrillation
- Decrease in muscle tone (hypotonia)
- Absent reflexes (hyporeflexia)



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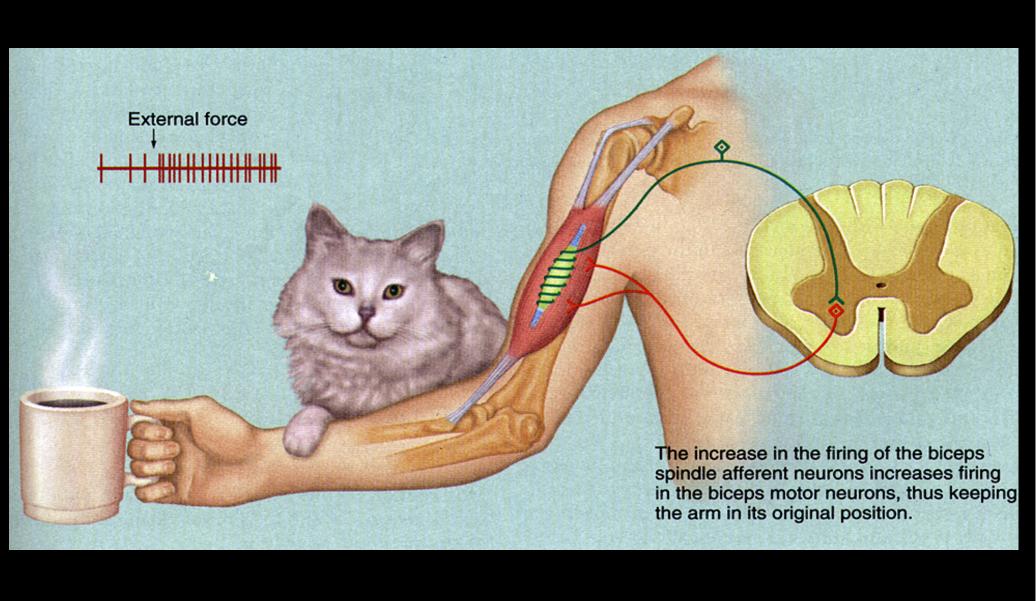
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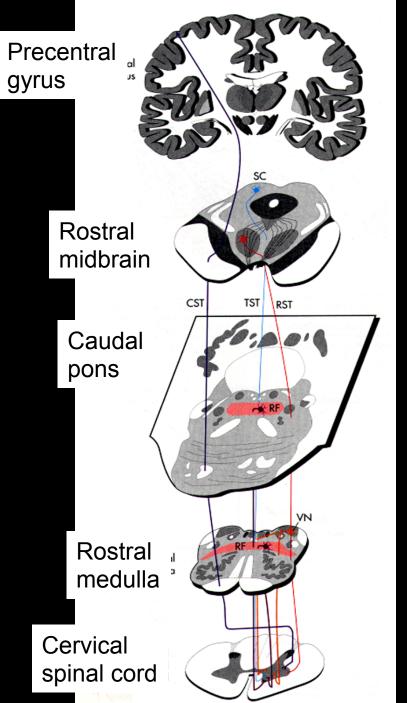
- Effects limited to small groups of muscles
- Muscle atrophy
- Weakness
- Fasciculation/fibrillation
- Decrease in muscle tone (hypotonia)
- Absent reflexes (hyporeflexia)



# Descending Spinal Pathways

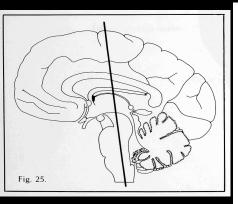
(from Nolte, p. 438)

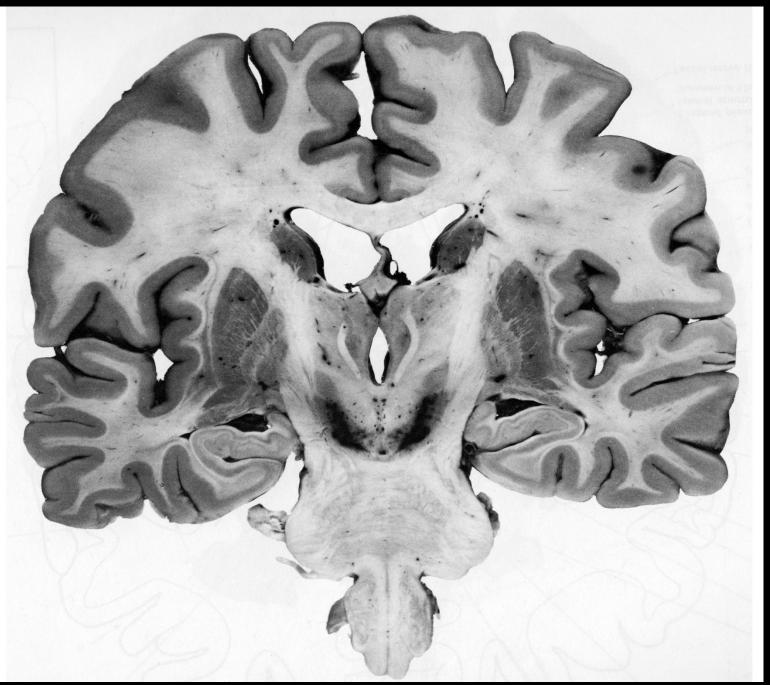
**Upper Motor Neurons** 

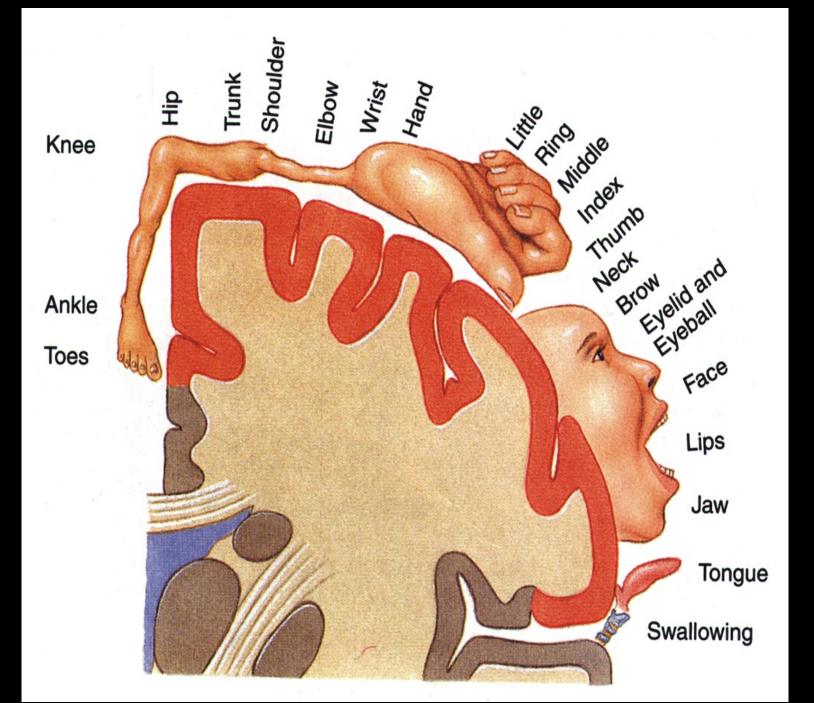


#### **Upper Motor Neuron Syndrome**

- Effects extend to large groups of muscles
- Atrophy is rare
- Weakness (paresis, not plegia)
- No fasciculation/fibrillation
- Increase in muscle tone (hypertonia)
- Hyperreflexia
- Clonus
- Initial contralateral flaccid paralysis
- Spasticity
- Babinski sign







#### **Upper Motor Neuron Syndrome**

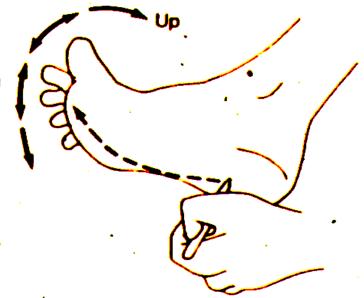
- Effects extend to large groups of muscles
- Atrophy is rare
- Moderate weakness (paresis, not plegia)
- No fasciculation/fibrillation
- Increase in muscle tone (hypertonia)
- Hyperreflexia
- Clonus
- Initial contralateral flaccid paralysis (eventual recovery of some movement)
- Spasticity ("clasp-knife reflex")
- Babinski sign

#### Normal plantar response

Down

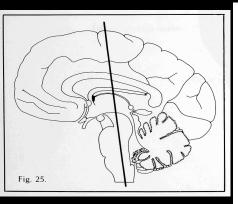
## Extensor plantar response (Babinski sign)

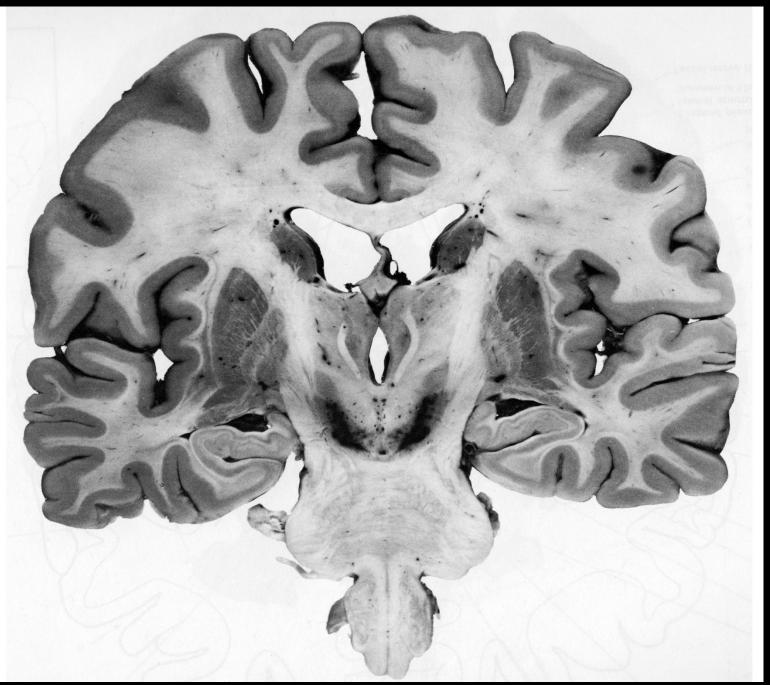
Fanning of toes



# Lesions of Higher Motor Cortical Areas (premotor, supplementary)

- Impairments in motor planning and strategies
- Unable to perform complex motor tasks (apraxia)



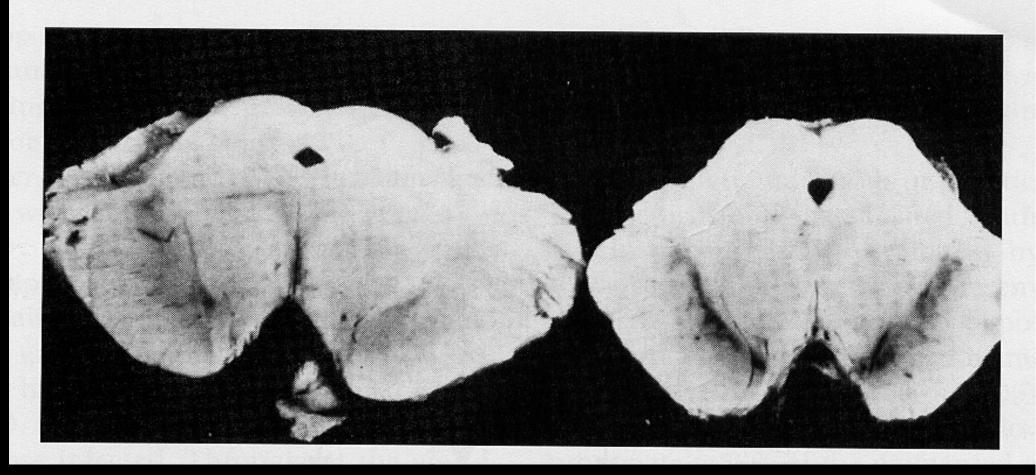


#### Disorders of the Basal Ganglia

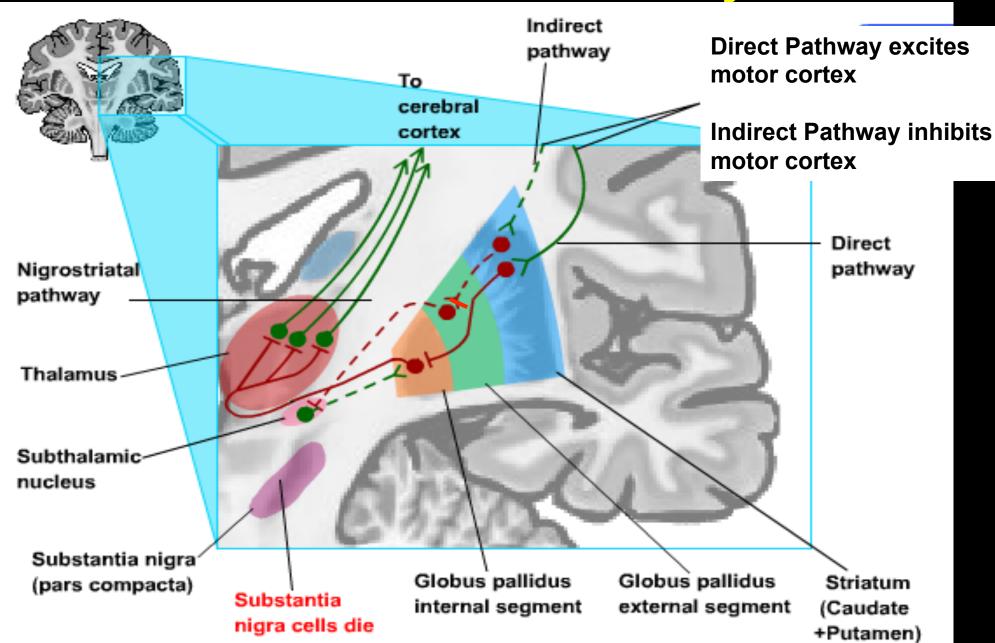
Dyskinesia: abnormal involuntary movements:

- Resting tremor ("pill-rolling tremor")
- Athetosis (slow, writhing movements of extremities)
- Chorea (jerky, random movements of limbs and face)
- Ballismus (violent, large movements of proximal limbs)
- Tardive dyskinesia
- Akinesia: abnormal involuntary posture
  - Rigidity (resistance to passive movement; "lead-pipe rigidity, cog-wheel rigidity")
  - Dystonia (abnormal postures from contraction of agonist & antagonist muscles)
  - Bradykinesia

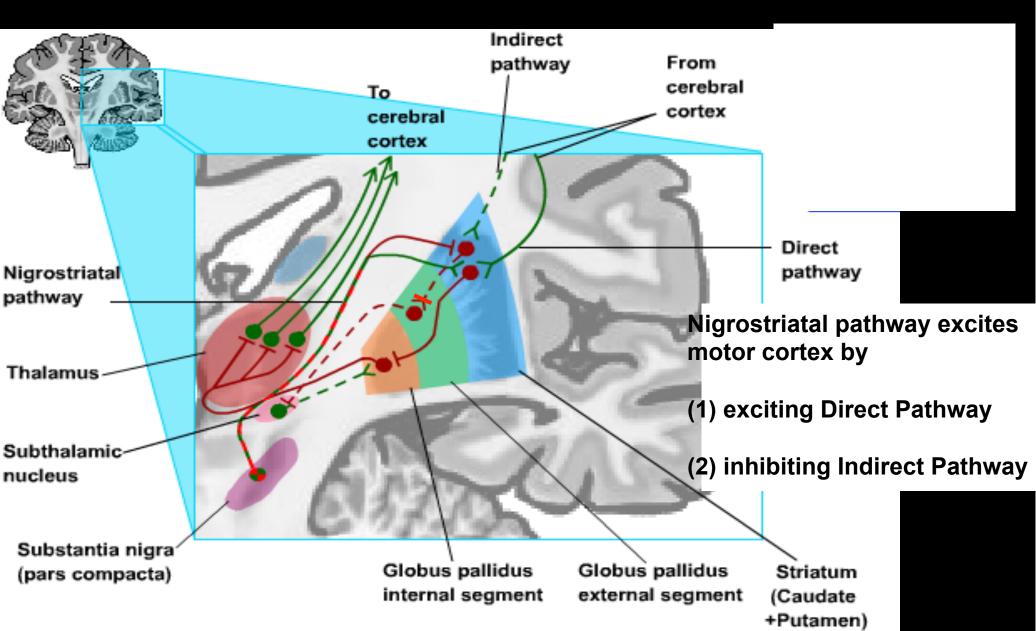
#### Parkinson's Disease



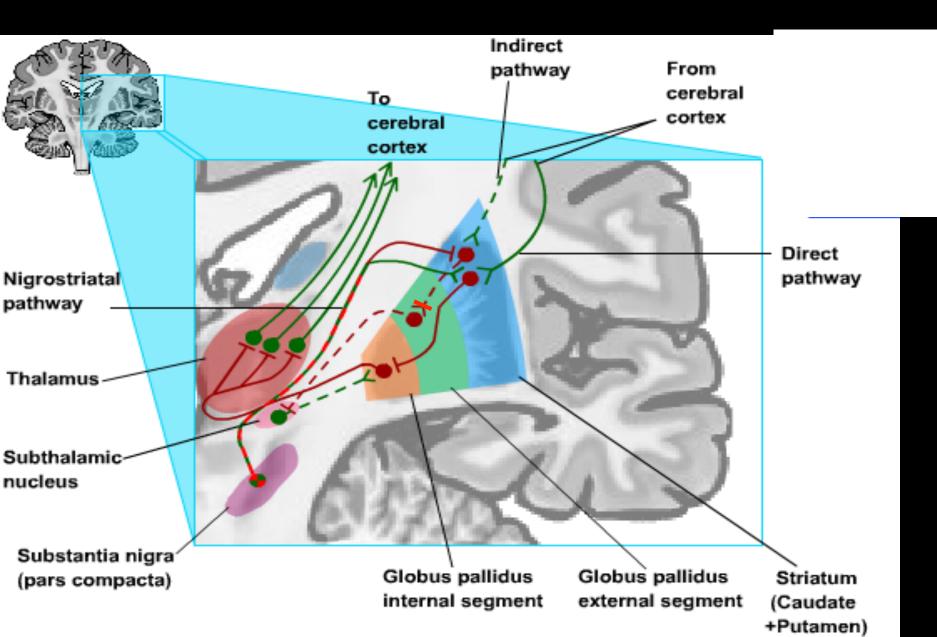
### **Direct and Indirect Pathways**



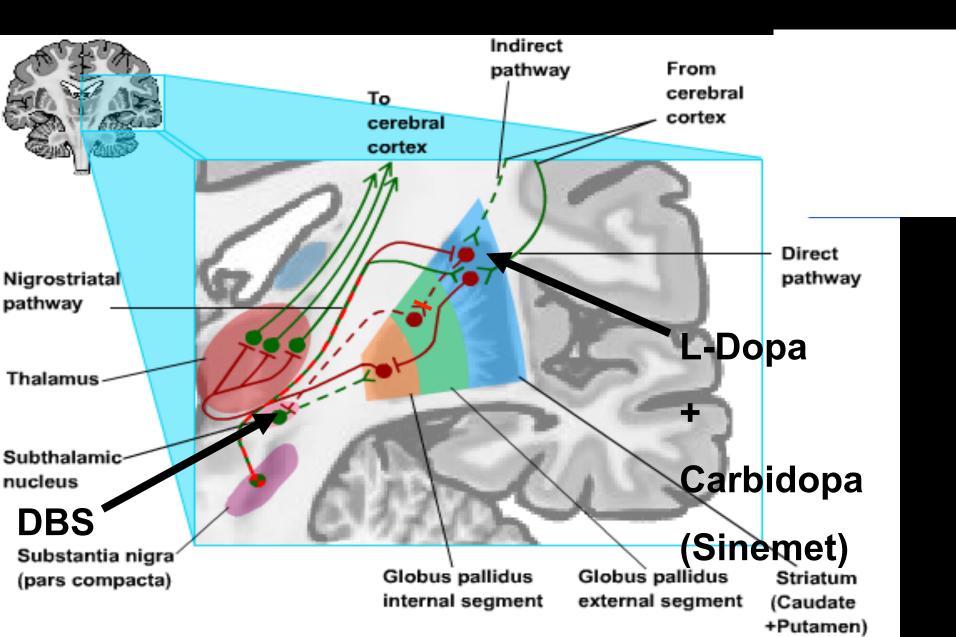
#### Dopaminergic Input from Substantia Nigra



#### Parkinson's Disease



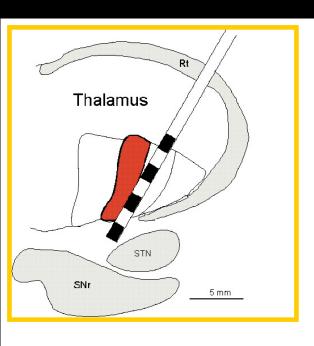
#### **Parkinson's Disease Treatments**

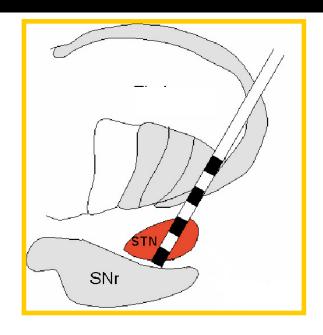


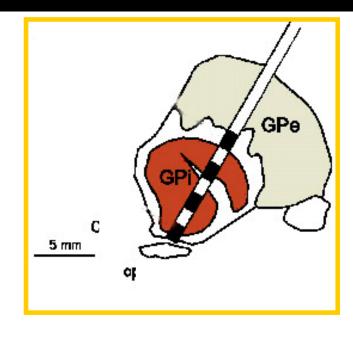
# Deep brain stimulation for movement disorders

- Mechanics:
- Pre-implantation MRI
- Sterotaxy (frameless or not)
- Recording to ensure correct placement
- Post-surgery: Programming: pulse width, rate, amplitude

#### **DBS** targets



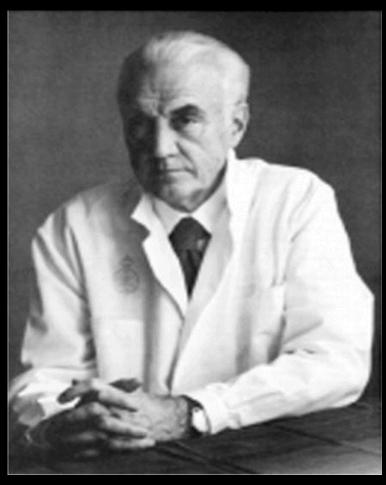




Vim Thalamus: Essential tremor & other tremor disorders Subthalamic nucleus: Parkinson's disease

Globus pallidus: Parkinson's disease & dystonia

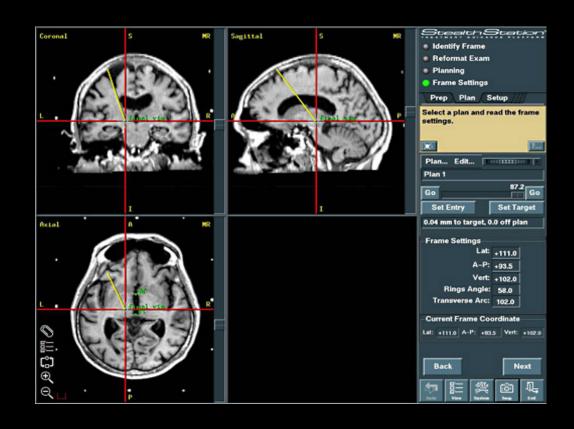
## Lars Leksell



"I was born under the sign of the 'archer' and look forward to sharpshoot into the brain."

### Surgical Technique: Targeting

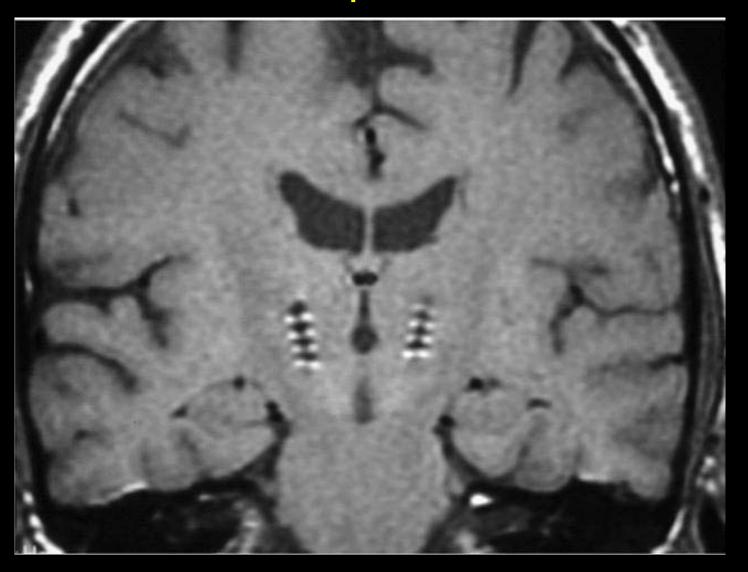
- Sophisticated imaging and software enables precise targeting for optimal outcomes and minimal risk
- Microelectrode recording (MER) offers additional levels of verification of lead location



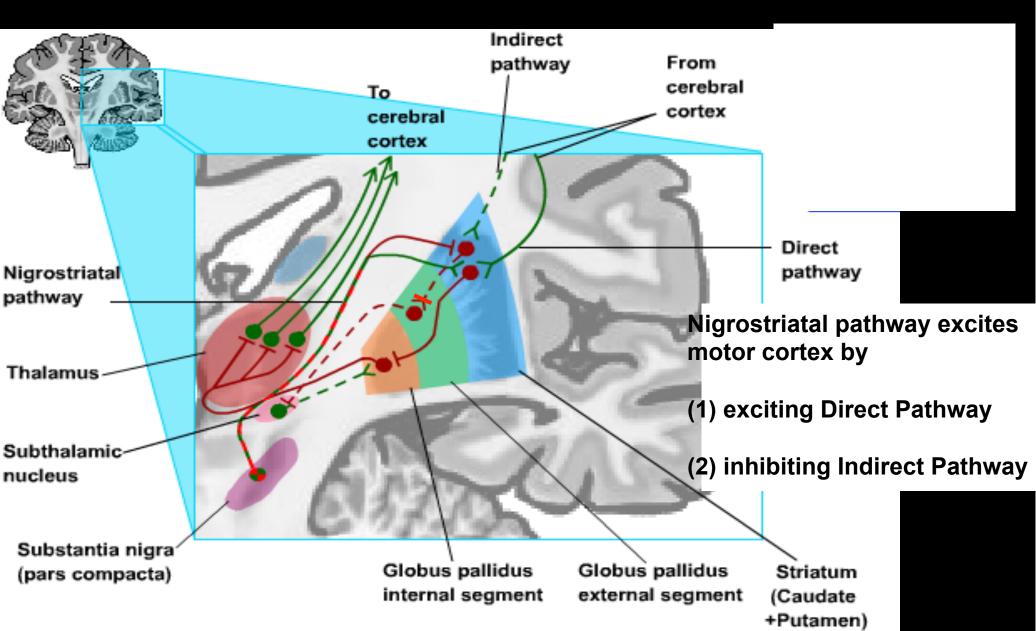
#### Complications:

- Stroke/ hemorrhage (2%)
- Infection (4%)
- Hardware complications leads breaking,
   electrode malfunction, stimulator failure, battery
   failure

## MRI of electrode placement



## Dopaminergic Input from Substantia Nigra



# Deep brain stimulation for movement disorders

- Post-surgery: Programming: pulse width, rate (135 185 Hz), amplitude
- Don't really understand how it works

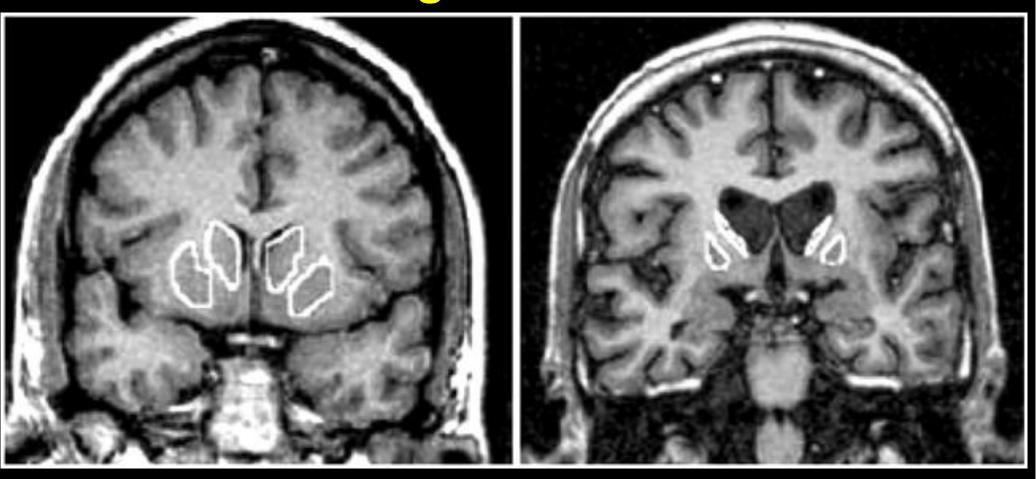
#### DBS and ...

- "No neuron is safe from a neurosurgeon"
- Andres Lozano, MD,PhD at the 62<sup>nd</sup> Annual Meeting of the AAN

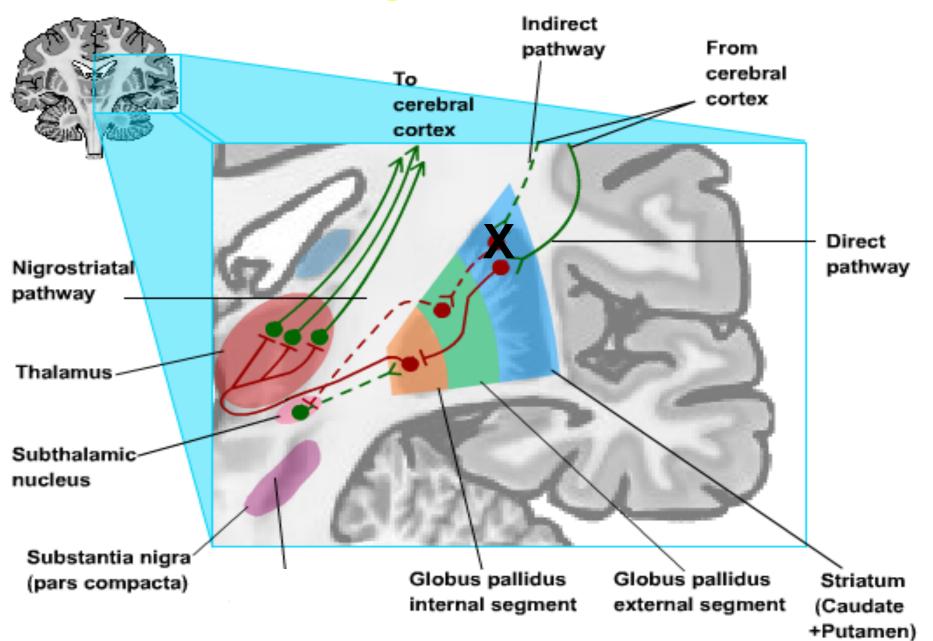
### **Huntington's Disease**

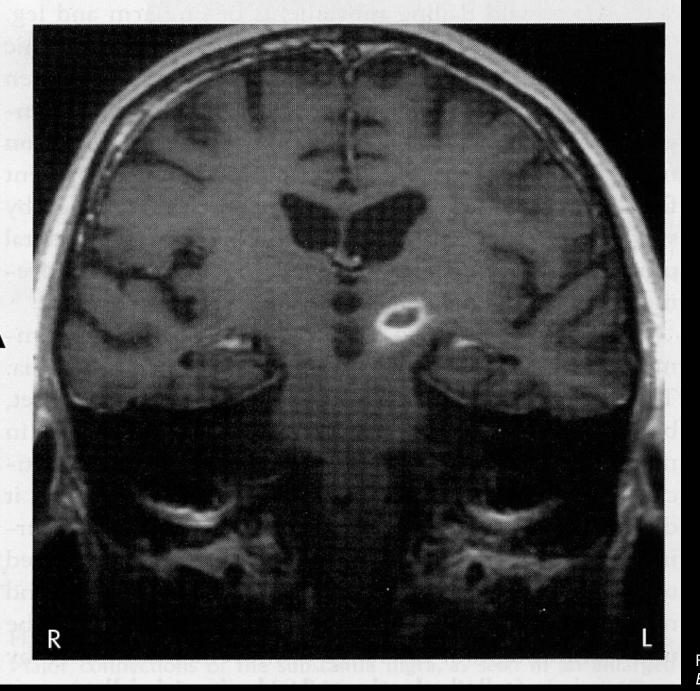
- Genetic disorder caused by an abnormally large number of repeats of nucleotide sequence CAG on chromosome 4
- Autosomal dominant mutation (50% chance of offspring inheriting the mutation and developing the disease)
- Mutation of huntingtin gene, causing destruction of indirect pathways neurons of striatum
- Characterized by continuous, choreiform movements of body (esp. hands and face)
- Dementia in advanced cases (reflecting loss of cognitive functions of basal ganglia as well as neocortical degeneration)
- No cure, treatment or prophylaxis

# **Huntington's Disease**



## **Huntington's Disease**

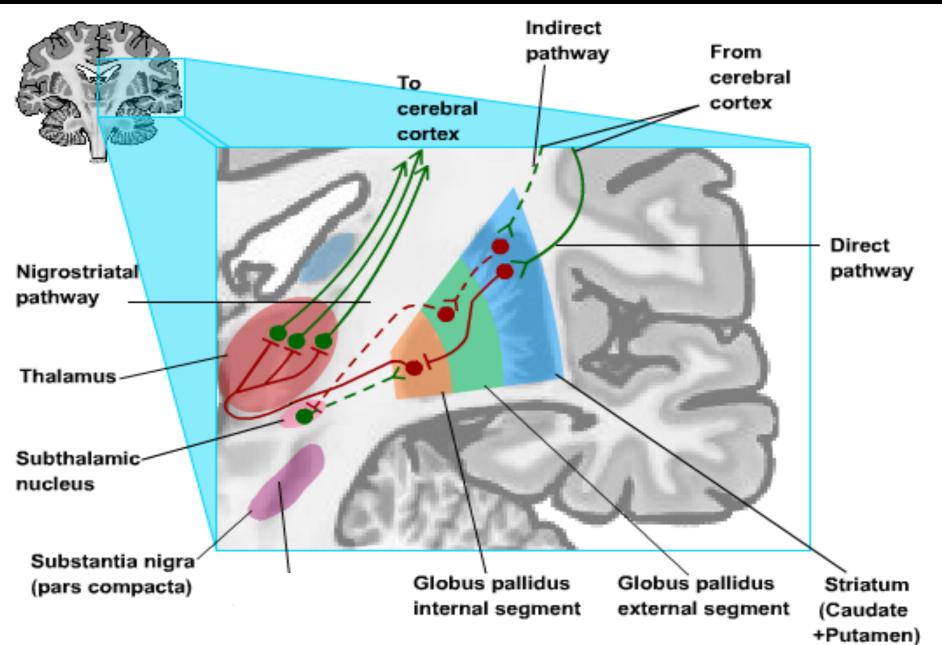


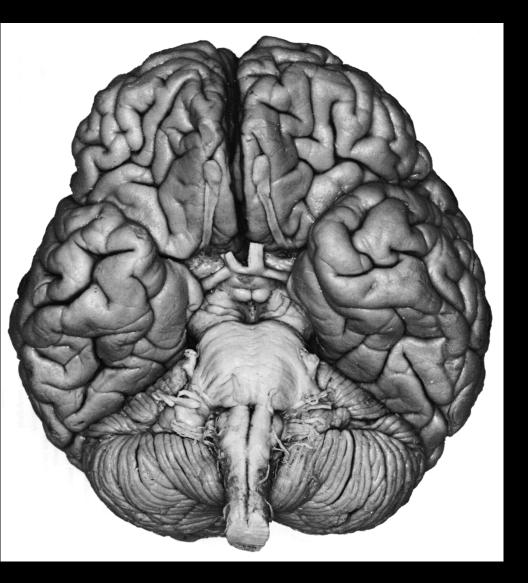


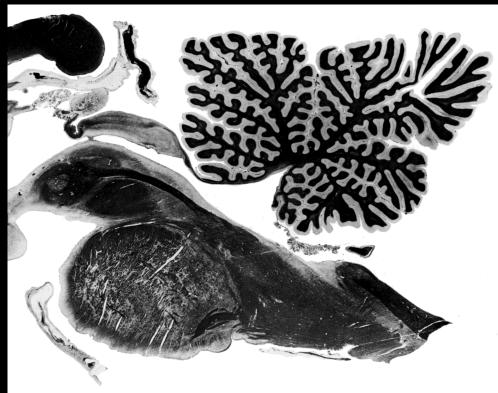
Lesion of the subthalamic nucleus produces hemiballismus

From J. Nolte (2002) *The Human Brain, 5<sup>th</sup> Edition* 

## Hemiballismus





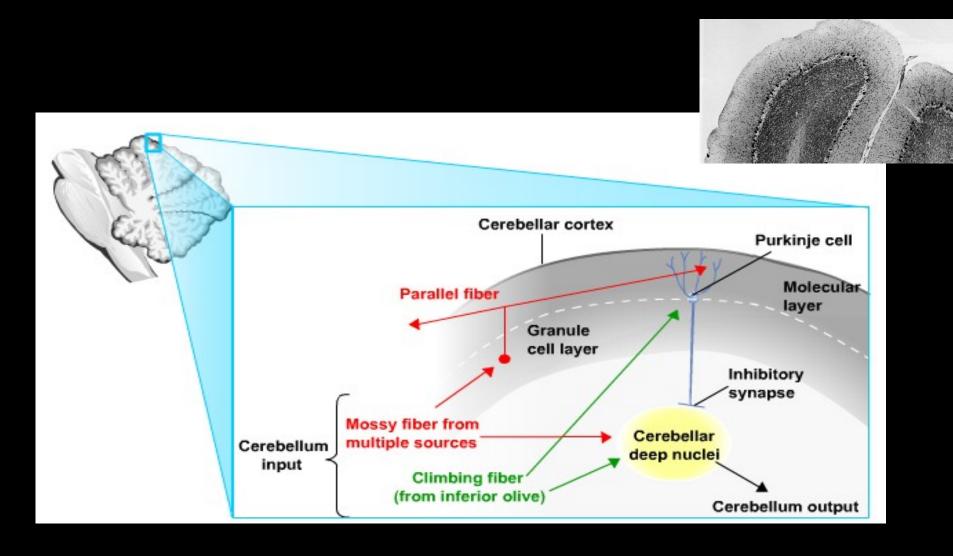


#### Disorders of the Cerebellum

- Ataxia
  - Disturbance of posture or gait
  - Decomposition of movement
- Dysmetria
- Dysdiadochokinesia
- Scanning speech
- Hypotonia
  - Pendular reflexes
- Tremor (intention tremor)
- Nystagmus
- Delay in initiating movements
- Subtle cognitive deficits

Cerebellum as a feedforward control system

#### **Internal Circuitry of Cerebellum**

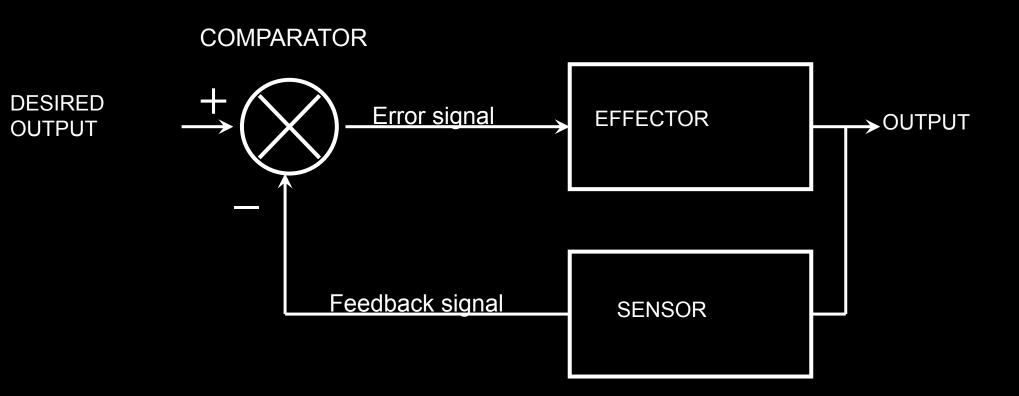


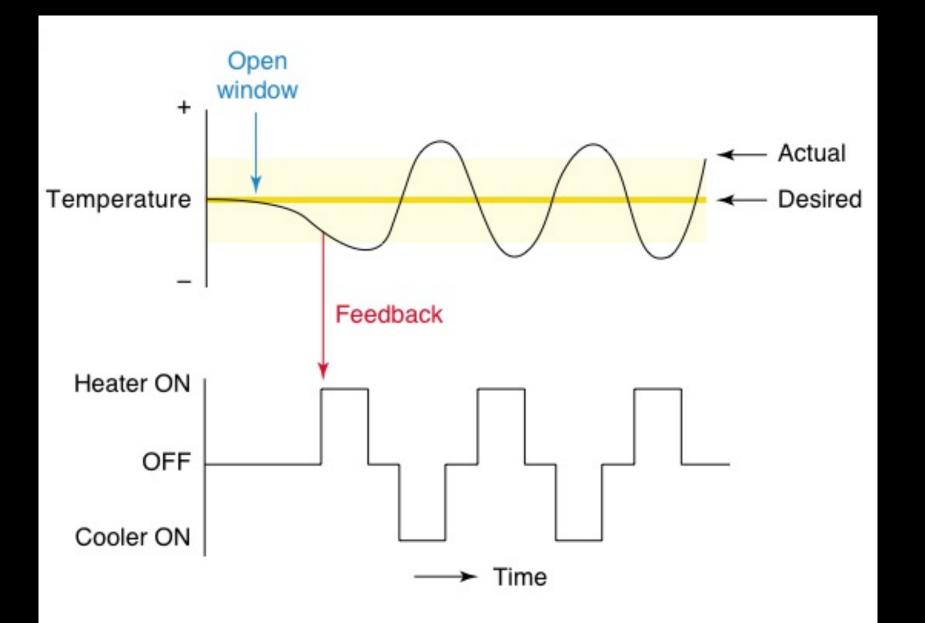
### Cerebellum as a Control System

Feedback Controller for Slow movements (e.g., posture)

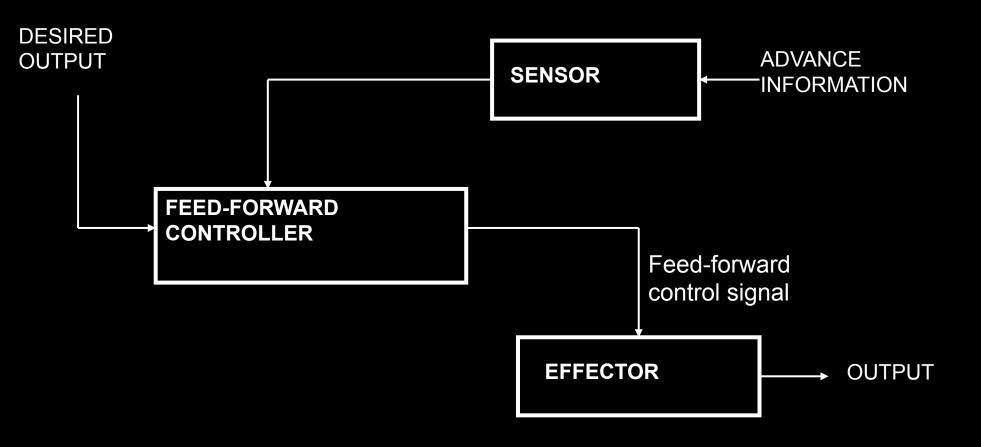
Feedforward Controller for Fast movements (e.g., most voluntary movements)

## Feedback control system

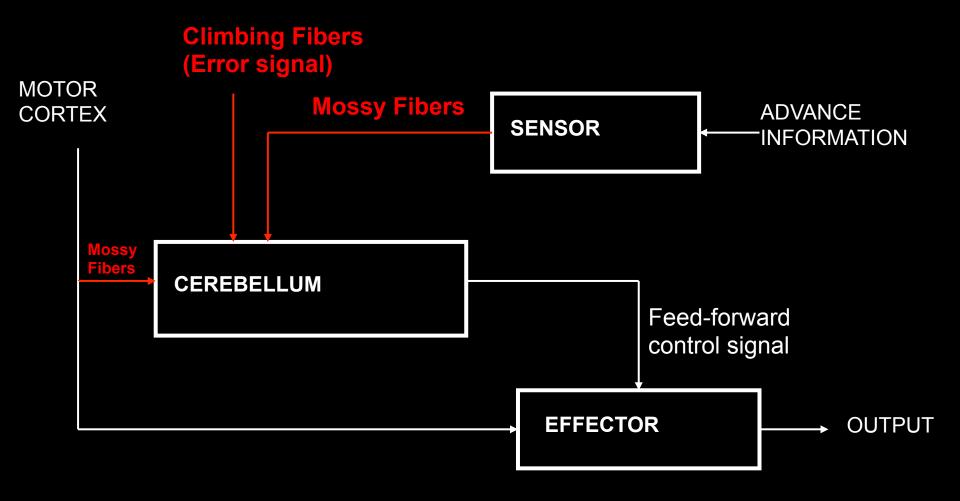




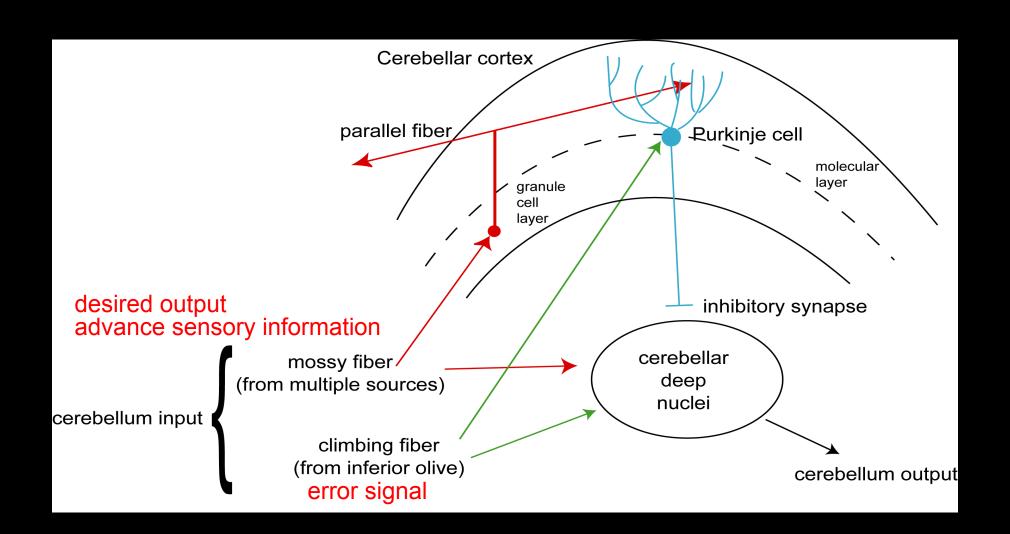
## Feedforward control system



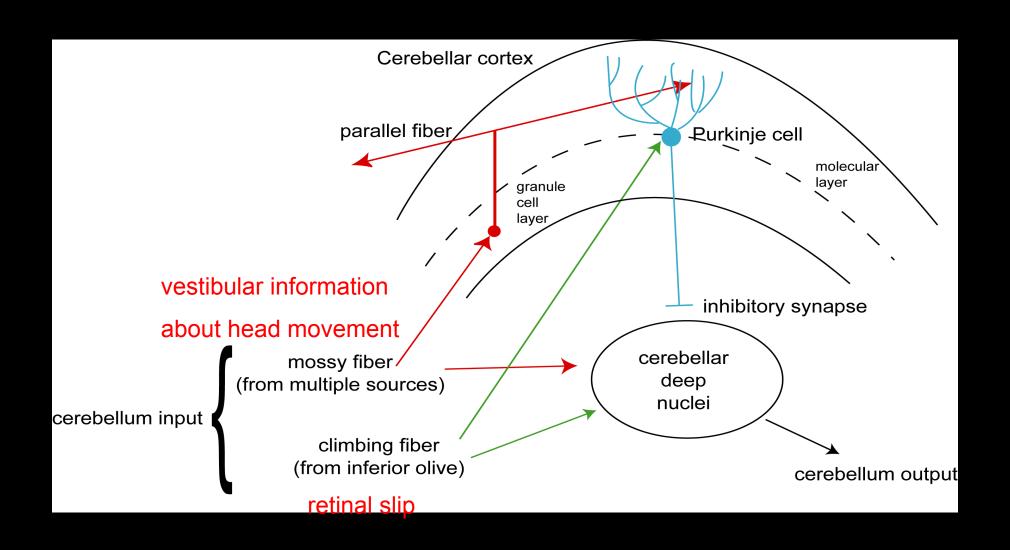
### Cerebellum as a feedforward control system



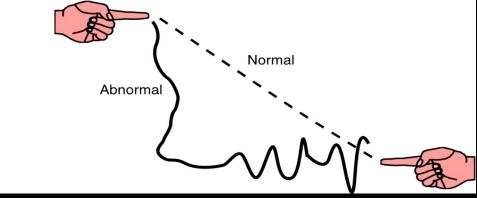
#### **Internal Circuitry of Cerebellum**



#### **VOR and feedforward control**



# Delay in initiating responses Normal Abnormal Dysdiadochokinesia **Abnormal** Decomposition of movement / intention tremor Abnormal



## Necessary Components of Proper Motor Control

- Volition
- Coordination of Signals to many muscle groups
- Proprioception
- Postural adjustments
- Sensory feedback
- Compensation for body and muscles
- Unconscious processing
- Adaptability