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# Board Review

## Neurology

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# ARS Question

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A 48 y.o. male presents with gradual progressive weakness of his proximal legs and arms without sensory loss. The exam confirms the weakness, reflexes are reduced, and no fasciculations are seen.

The best anatomical localization is?

- A. Brain
- B. Muscle
- C. Motor neuron
- D. Spinal cord
- E. Peripheral nerve

# ARS Question

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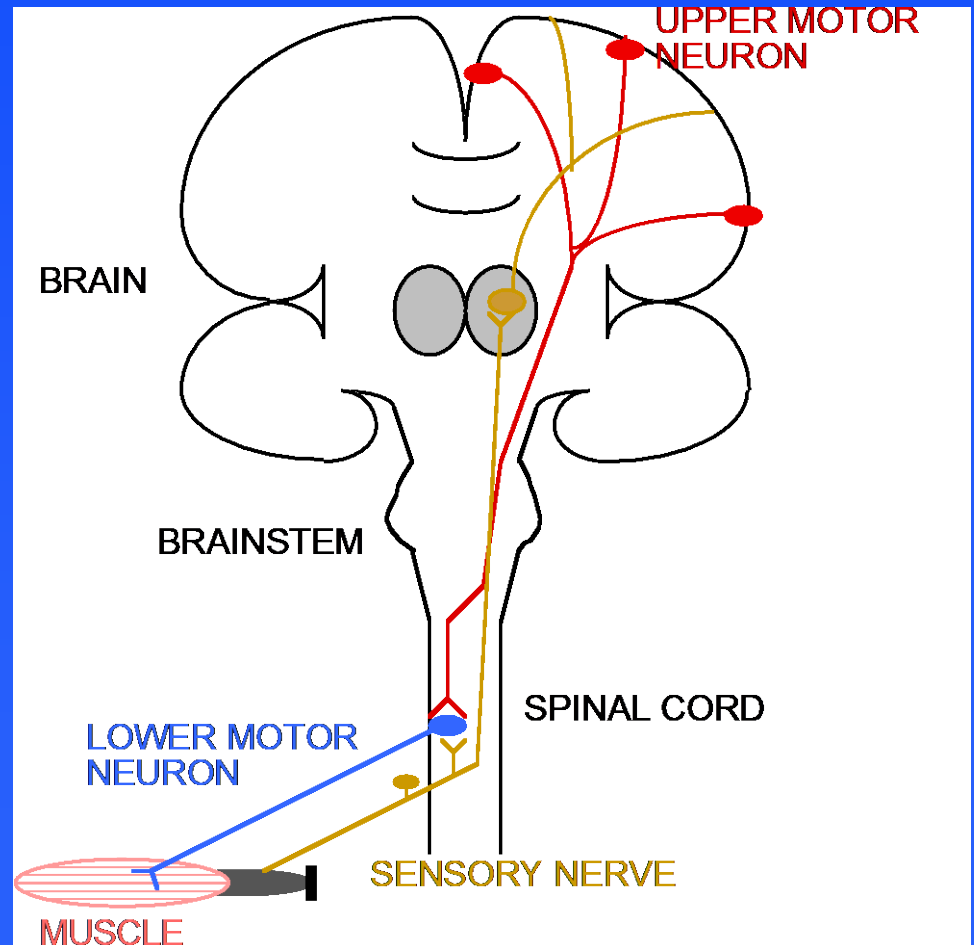
A 52 y.o. male presents with subacute sensory loss in the feet that ascended to his chest. Arm sensation is fine. Weakness is present in both legs. Urinary urgency has developed. The exam confirms leg weakness without atrophy or fasciculations. There is a sensory level at T6.

The best anatomical localization is?

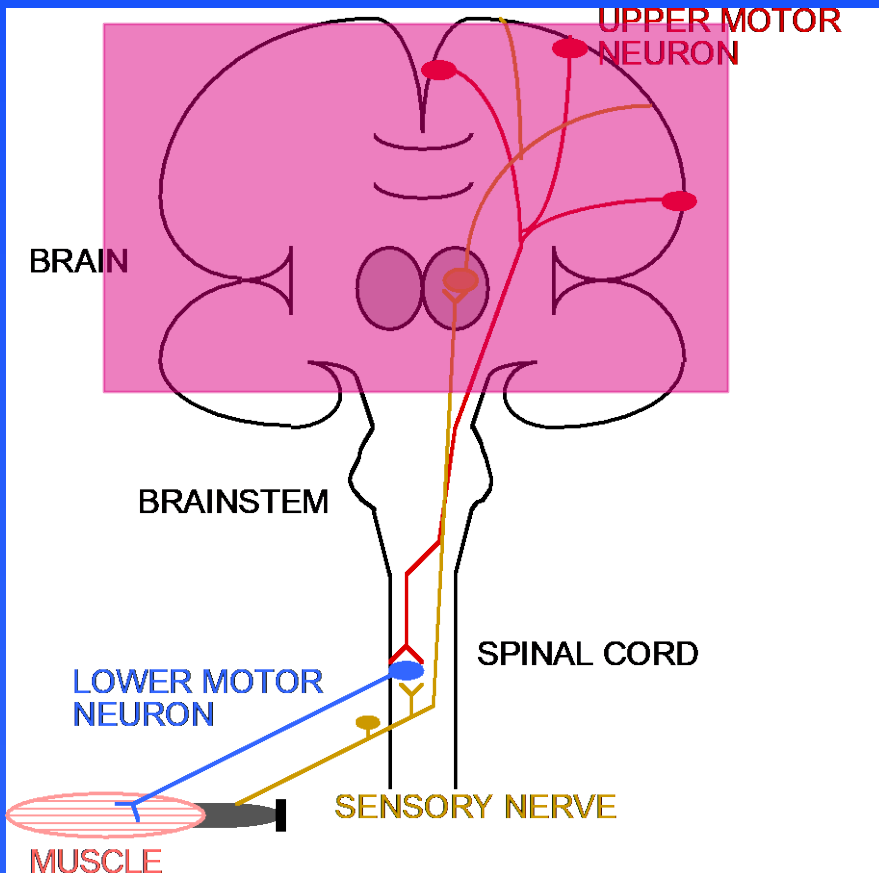
- A. Brain
- B. Muscle
- C. Motor neuron
- D. Spinal cord
- E. Peripheral nerve

# Neurological “Levels”

- Brain
- Brainstem
- Spinal cord
- Motor neuron
- Peripheral nerve
- Neuromuscular junction
- Muscle

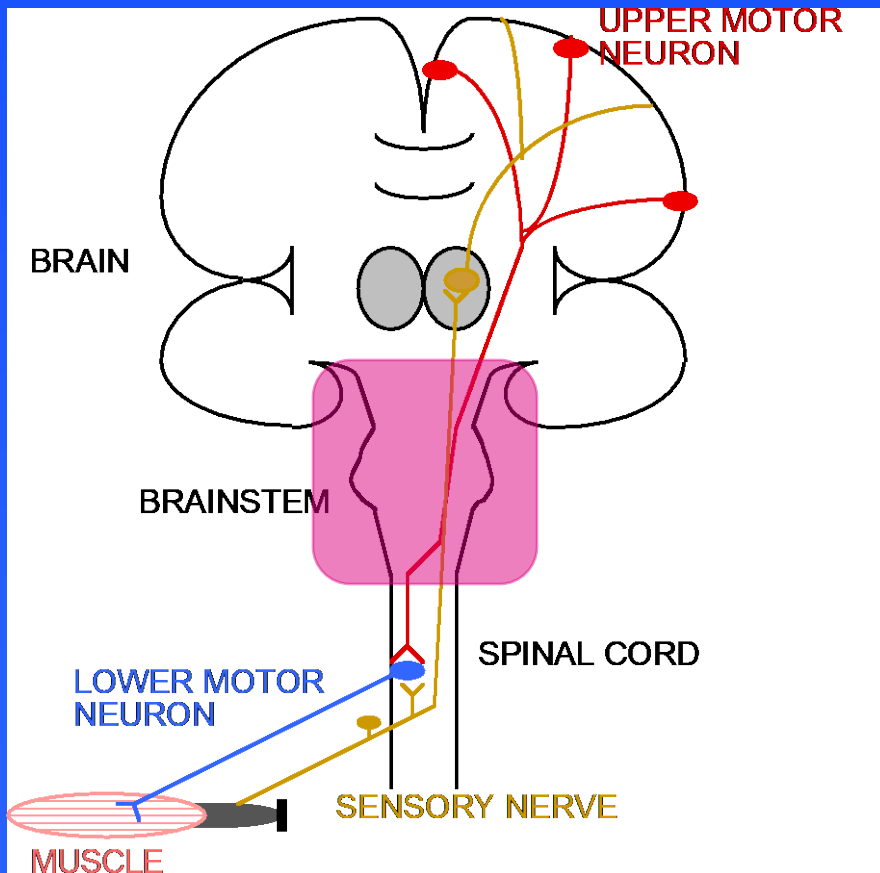


# Level: Brain



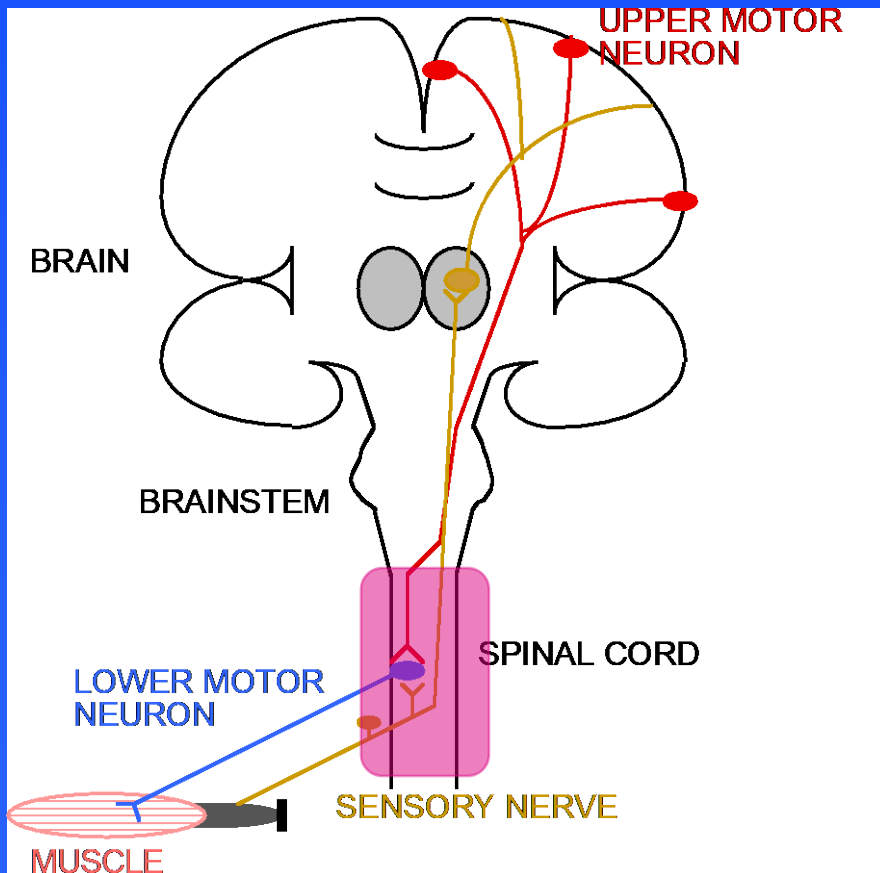
- Often unilateral
- Motor and/or sensory
- Language
- Consciousness
- Memory
- Behavior
- Vision
- Seizures
- Movement d/o

# Level: Brainstem



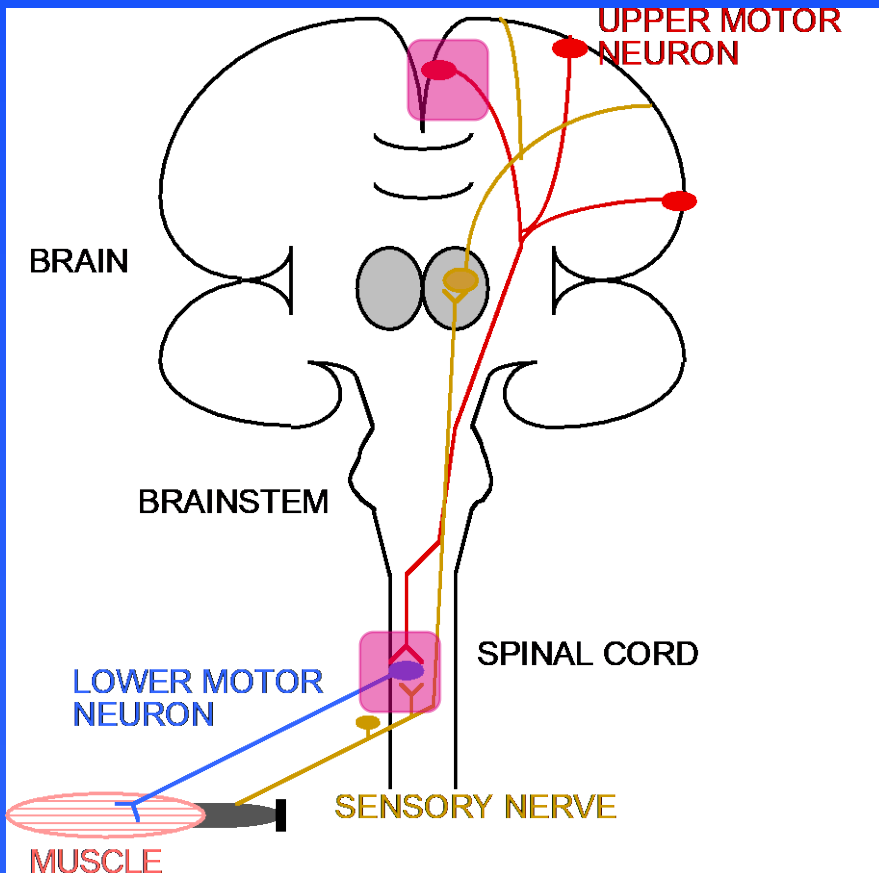
- Often unilateral
- Motor and/or sensory
- Consciousness
- Cerebellar
- Movement d/o
- Cranial nerves
  - » Diplopia
  - » Vertigo
  - » Face
  - » Swallow
  - » Tongue

# Level: Spinal Cord



- Often bilateral
- Motor and/or sensory
- Head OK
- Bowel, bladder and erectile

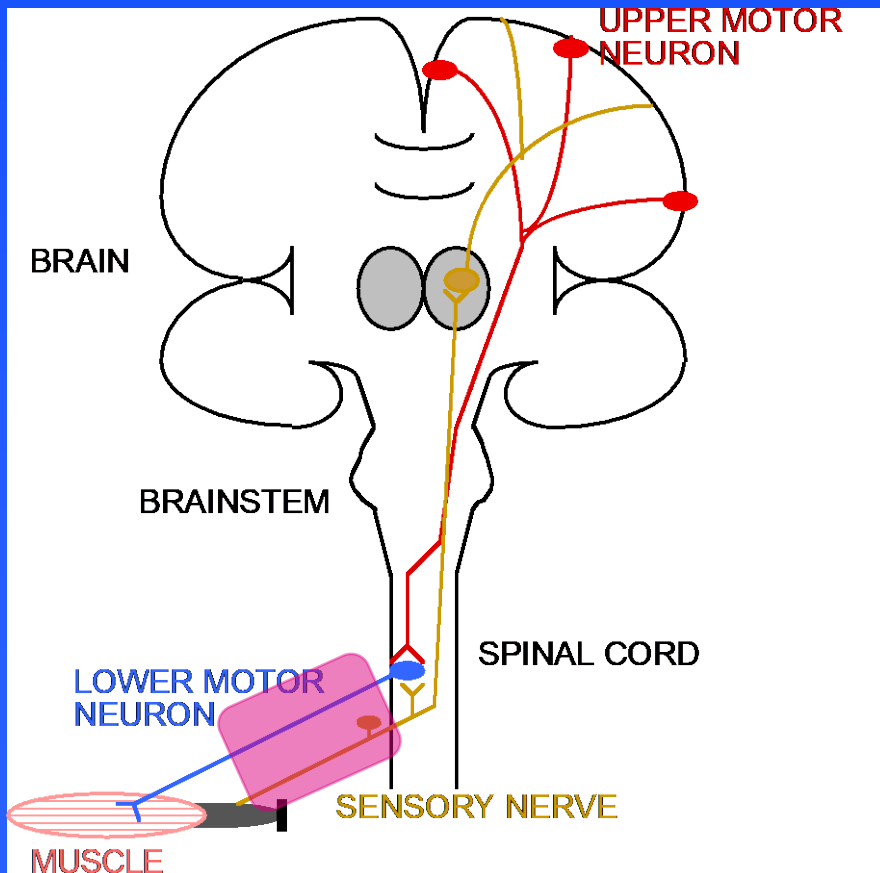
# Level: Motor Neuron



- Asymmetric bilateral
- Motor
- Proximal and distal
- Insidious onset
- Fasciculations

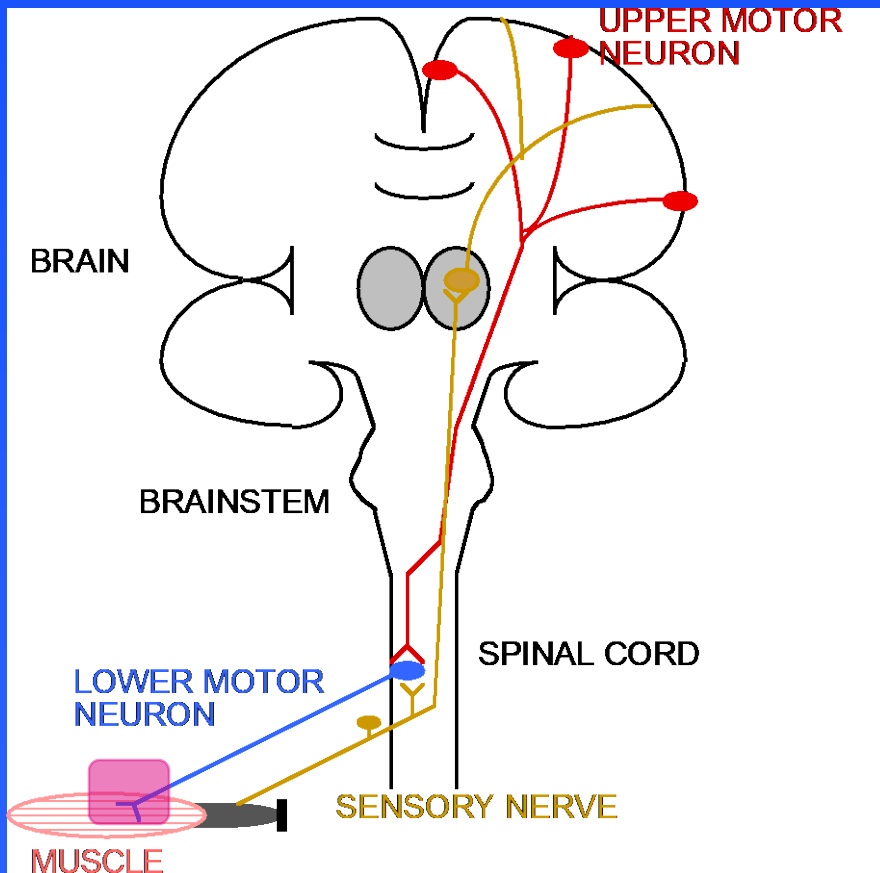


# Level: Peripheral Nerve



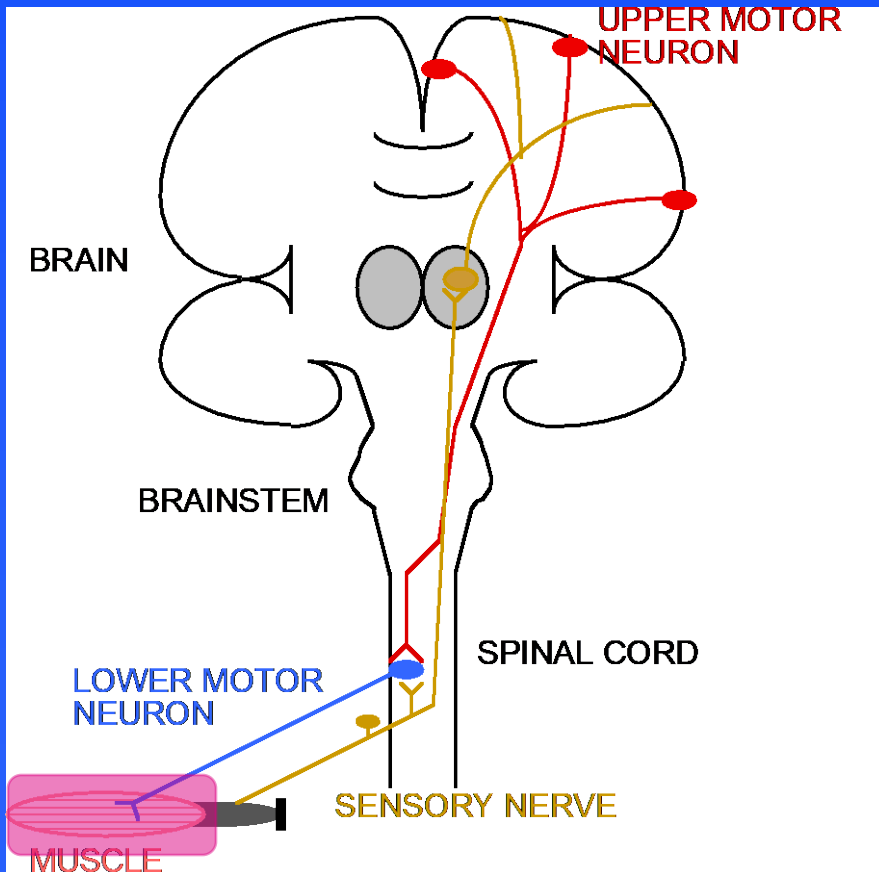
- Symmetric or focal
- Sensory > motor
- Cranial nerves 3-12
- Often distal
  - » Stocking-glove
- If proximal think
  - » Demyelinating (UE + LE)
  - » Cauda equina (LE)

# Level: NMJ



- Asymmetric bilateral
- Motor only
- Proximal and distal
  - » Eyes involved in myasthenia gravis
- Fatigable weakness
  - » Myasthenia gravis
- Progressive weakness
  - » Lambert-Eaton myasthenic syndrome

# Level: Muscle



- Symmetric bilateral
- Motor only
- Usually proximal

# ARS Question

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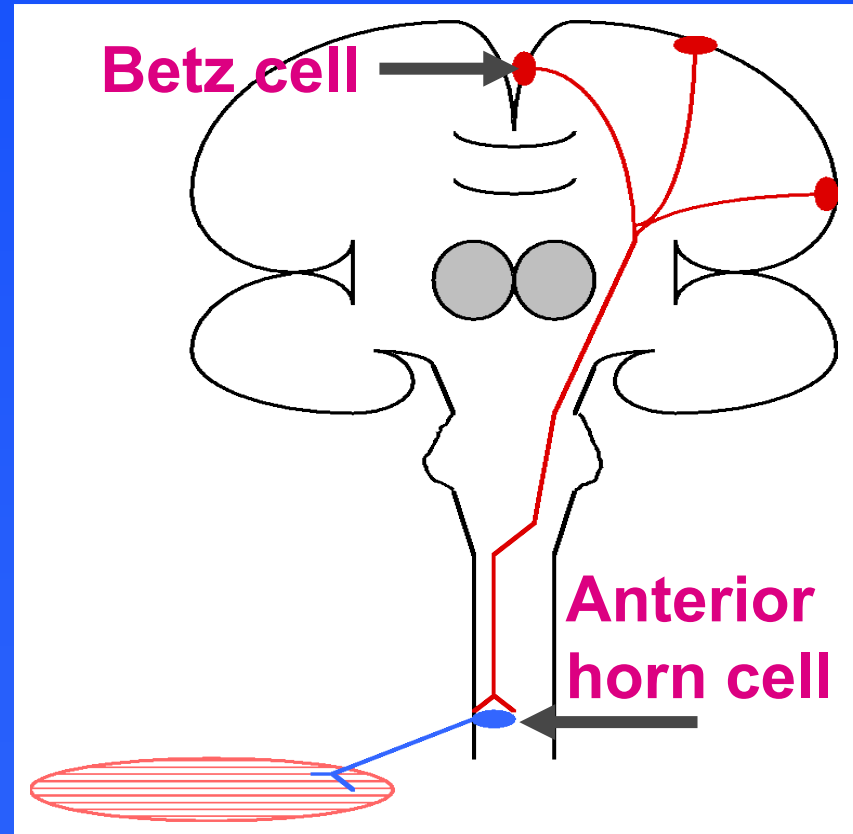
This is a 56 yo woman with gradual weakness of her right arm such that she can no longer open jars. She stopped playing tennis 3 months ago. Gait feels stiff. Her friends wonder why her speech is slurred. She denies sensory, cognitive, and bowel/bladder difficulties. Her exam is remarkable for diffuse weakness and hyperreflexia that is greater in the right arm, and diffuse fasciculations.

What test is likely to confirm your diagnostic suspicion?

- A. Brain MRI
- B. Spinal fluid for increased protein
- C. Cervical MRI
- D. Electrodiagnostic testing
- E. Tibial evoked potentials

# Amyotrophic Lateral Sclerosis

- Degeneration of motor neurons
- Etiology unknown
- No sensory, visual, or B/B involvement
- 20% have dementia
- Asymmetric weakness
- Bulbar or limb onset
- Survival usually 2-5 years
- Rx: Riluzole, Edaravone (2017)



# ALS: Diagnosis

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- H&P
- Electrodiagnostic testing = nerve conduction + EMG
  - » Denervation and reinnervation + fasciculations in three body segments
  - » Thoracic spine segment (look for this EMG info)

# ARS Question

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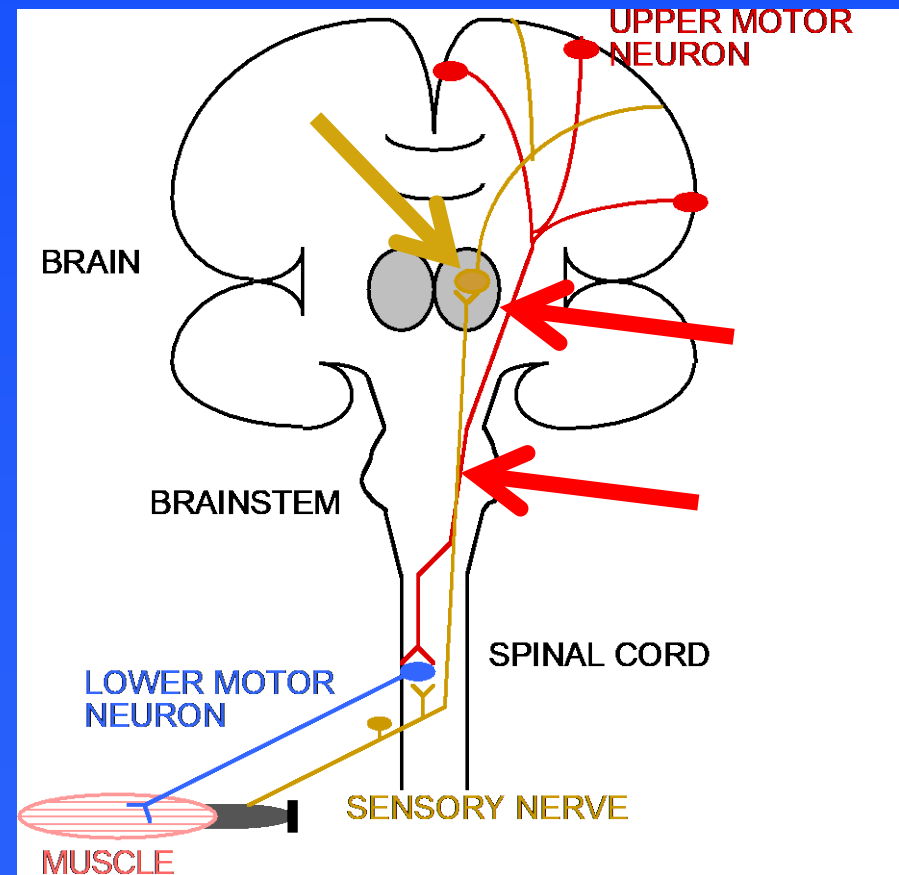
A 66 y.o. female presents with acute onset of sensory loss in the right face, arm and leg. No weakness, vision or cognitive change.

Reduction of which risk factor will have the greatest impact at preventing a second event?

- A. Hyperglycemia
- B. Hyperlipidemia
- C. Weight loss
- D. Hypertension
- E. Smoking

# Lacunar Syndromes

- Pure **motor** stroke
  - » Internal capsule
  - » Basis pontis
- Pure **sensory** stroke
  - » Thalamus





# HTN: Secondary Prevention

<b>Recommendations for Hypertension</b> Referenced studies that support recommendations are summarized in online Data Supplements 11 and 12.		
<b>COR</b>	<b>LOE</b>	<b>Recommendations</b>
<b>1</b>	<b>A</b>	1. In patients with hypertension who experience a stroke or TIA, treatment with a thiazide diuretic, angiotensin-converting enzyme inhibitor, or angiotensin II receptor blockers is useful for lowering BP and reducing recurrent stroke risk. <sup>185-189</sup>
<b>1</b>	<b>B-R</b>	2. In patients with hypertension who experience a stroke or TIA, an office BP goal of <130/80 mmHg is recommended for most patients to reduce the risk of recurrent stroke and vascular events. <sup>185,190-194</sup>
<b>1</b>	<b>B-NR</b>	3. In patients with hypertension who experience a stroke or TIA, individualized drug regimens that take into account patient comorbidities, agent pharmacological class, and patient preference are recommended to maximize drug efficacy. <sup>188,189,195,196</sup>

# Dyslipidemia: Secondary Prevention

Recommendations for Treating and Monitoring Hyperlipidemia Referenced studies that support recommendations are summarized in online Data Supplement 13.		
COR	LOE	Recommendations
		<b>Treatment</b>
1	A	1. In patients with ischemic stroke with no known coronary heart disease, no major cardiac sources of embolism, and LDL cholesterol (LDL-C) >100 mg/dL, atorvastatin 80 mg daily is indicated to reduce risk of stroke recurrence. <sup>208,209</sup>
1	A	2. In patients with ischemic stroke or TIA and atherosclerotic disease (intracranial, carotid, aortic, or coronary), lipid-lowering therapy with a statin and also ezetimibe, if needed, to a goal LDL-C of <70 mg/dL is recommended to reduce the risk of major cardiovascular events. <sup>210</sup>

# ARS Question

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A 63 y.o. male presents with right hemiparesis, hemisensory loss and nonfluent aphasia. Onset 1 day ago. The work-up ascribes the cause to emboli from 50% stenosis of the left carotid artery.

What treatment do you recommend to prevent another stroke?

- A. Anti-platelet therapy
- B. Anti-coagulation
- C. Carotid endarterectomy

# Antiplatelet Summary

MEDICATION	COMMENTS
ASPIRIN	325mg (81mg). Inexpensive. Often first choice.
CLOPIDOGREL	Once daily. More effective than ASA. Best first-line choice. Expensive, but generic soon. Rash
ASA + ER DIPYRIDAMOLE (AGGRENOX)	Twice daily. More effective than ASA. Equal to clopidogrel (PRoFESS <sup>1</sup> ). Expensive. HA, GI upset. Second line.
ASA + CLOPIDOGREL	Potent antiplatelet effect. Consider for 3 weeks after minor stroke or TIA. (POINT <sup>2</sup> ) Higher bleeding risk longterm and when major stroke (MATCH <sup>3</sup> , CHARISMA <sup>4</sup> ).

Adapted from AAN, Continuum: Secondary Stroke Prevention. 2011;17(6):1260

<sup>1</sup> N Engl J Med. 2008;359(12):1238-51

<sup>2</sup> N Engl J Med 2018;379:215-25

<sup>3</sup> Lancet 2004;364:331

<sup>4</sup> N Engl J Med. 2006;354(16):1706-17

# 2018 Antiplatelet Guidelines

**Dual antiplatelet therapy (DAPT)**, when started within 24 hours of symptom onset of minor stroke (NIHSS  $\leq 3$ ) or high risk TIA (ABCD2  $\geq 4$ ) and used for 10-21 days:

--Reduces non-fatal recurrent stroke (ischaemic and haemorrhagic) in the first 90 days by 1.9% (high quality evidence)

DAPT has little or no impact on:

--All-cause mortality (moderate quality evidence)

--Incidence of myocardial infarction or recurrent transient ischaemic attack (moderate quality evidence).

DAPT has some harms:

--A small (0.2%), possibly important increase in moderate to major extracranial bleeding events (moderate quality evidence)

--A small increase in the risk of minor extracranial bleeding events by 0.7% (high quality evidence).

# ARS Question

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A 63 y.o. male presents with right hemiparesis, hemisensory loss and nonfluent aphasia. Time of onset was 3 hours. The work-up reveals a left proximal MCA occlusion.

What treatment do you recommend?

- A. Anti-platelet therapy
- B. T-PA alone
- C. Thrombectomy alone
- D. T-PA followed by thrombectomy
- E. Heparin SQ

# t-PA Trials

## □ NINDS

- » <3 hours
- » Good outcome
  - No or minimal disability at 3 months
  - 29% placebo
  - 41% t-PA
  - NNT = 7
- » Bleeding risk
  - 0.6% placebo
  - 6.4% t-PA
  - NNH = 17 or more

## □ ECASS-III

- » <4.5 hours (Not FDA approved)
- » Exclusions from NINDS
  - NIHSS > 25 (severe)
  - Previous stroke and diabetes
  - Coumadin regardless of INR
  - Age > 80
- » Good outcome
  - 45% placebo
  - 52% t-PA
  - NNT = 14
- » Bleeding risk unchanged

# MR. CLEAN Trial

- 500 adults
- Confirmed anterior circulation occlusion
- 2<sup>nd</sup> generation thrombectomy device
- 90% received t-PA
- <6 hours to groin puncture
- Good outcome
  - » No or minimal disability at 3 months
  - » 19% t-PA
  - » 33% thrombectomy
  - » NNT = 7
- No difference death or intracranial hemorrhage
- New clinical ischemic strokes in different vascular territories at 3 months
  - » 5.6% thrombectomy
  - » 0.4% t-PA



# DAWN and DEFUSE-3 Trials

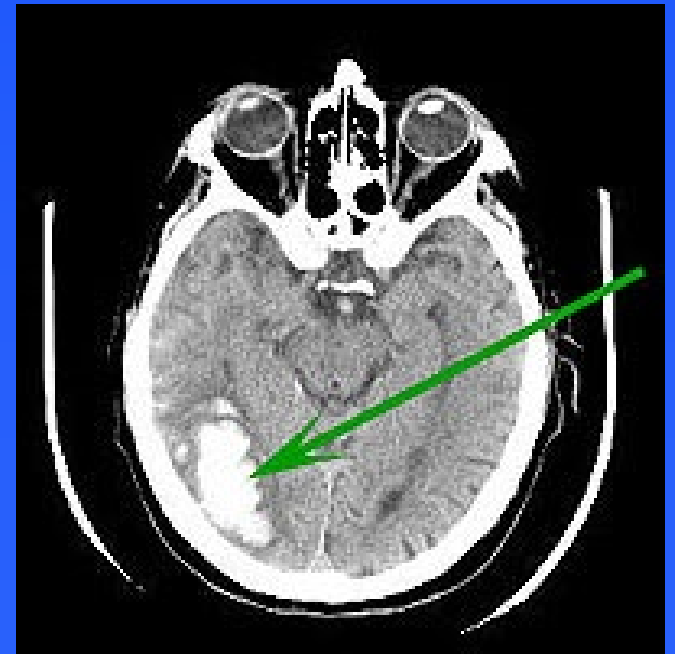
- 206 adults
- Confirmed anterior circulation occlusion
- 2<sup>nd</sup> generation thrombectomy device
- 6-24 hours to groin puncture
- Either
  - » Mismatch between the severity of the clinical deficit and the infarct volume.
  - » Assessed by diffusion-weighted MRI or perfusion CT.
- Good outcome
  - » No or minimal disability at 3 months
  - » 13% placebo
  - » 46% thrombectomy
  - » NNT = 3
- No difference death or intracranial hemorrhage

# ARS Question

A 82 y.o. woman has diabetes mellitus, and ASCVD. On the day prior to admission she had acute onset of left vision loss and headache. The work-up shows a right temporal-occipital lobar hemorrhagic stroke. She is on ASA 81mg.

What is the most likely cause of lobar hemorrhage in this age group?

- A. Amyloid angiopathy
- B. Hypertension
- C. Endocarditis, septic emboli
- D. Vascular malformation
- E. Brain tumor

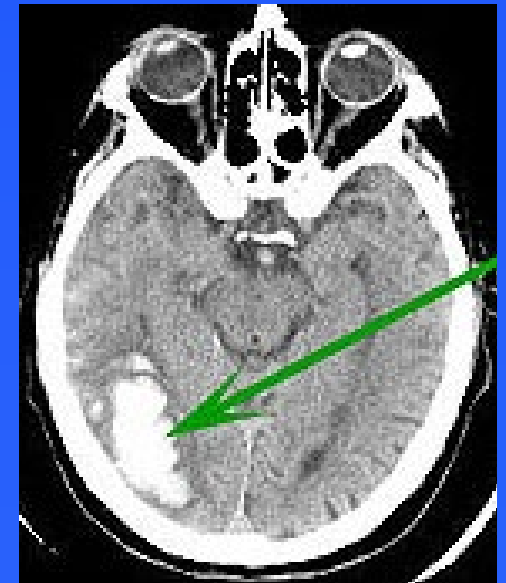


# Stroke: Hemorrhagic

- HTN
  - » Most common cause of hemorrhage
  - » Deep location
- Amyloid angiopathy
  - » Lobar location
- Vascular malformations
  - » Any location
- Treatment
  - » Control HTN
  - » Avoid blood thinning
  - » Surgery



HTN



Amyloid angiopathy

# ARS Question

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This 24 yo female presents with blurry vision and pain in the right eye for 3 days. Last year she had numbness in her left leg for 2 weeks?

You make a presumptive diagnosis and prescribe what to treat her acute symptoms?

- A. Betaseron
- B. IV steroids
- C. Ocular steroids
- D. Amantadine
- E. Nataluzimab

# MS: Glucocorticoids

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## □ Level 1 evidence

- » Methylprednisolone 1000mg IV qd x 3-5d
- » Short-term benefit to hasten recovery
- » No long-term influence on course
- » Cochrane Database Syst Rev 2000;(4):CD001331

## □ Level 2 evidence

- » Following IV methylpred with PO prednisone (ex. 60mg x 10d)
- » Using a PO equivalent dose instead of IV. Cochrane Database Syst Rev. 2009 Jul 8;(3):CD006921

# ARS Question

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A 56 y.o. man presents with trouble walking, small hand writing, and a resting tremor.

You make a presumptive diagnosis and suggest:

- A. Sinemet trial
- B. Head MRI
- C. Propranolol for essential tremor
- D. Stroke work-up
- E. SSRI for anxiety

# PD: Clinical

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- Slow progressive course
- 3 cardinal features
  - » Resting tremor
  - » Bradykinesia
  - » Rigidity (cogwheel)
- These features lead to gait trouble
- Other: dementia, depression, autonomic

# PD: Treatment

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- Sinemet = carbidopa/levodopa
  - » Levodopa converts to dopamine
  - » Most potent drug for PD
- Dopamine agonists
  - » Act directly on dopamine receptors
  - » Synergy with levodopa
  - » Ex. Pramipexole, Ropinirole
- Other: COMT inhibitors, MAO-B inhibitor, anticholinergics



# ARS Question

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This is a 42 yo woman with an uncomfortable feeling in her legs 5 days per week in the evening. Walking will relieve the problem temporarily. Her neurological exam is normal.

What test should you order next?

- A. Serum copper and ceruloplasmin
- B. B12
- C. Potassium
- D. Iron studies
- E. TSH

# Restless Leg Syndrome

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- Common, F>M
- Etiology
  - » Primary: CNS dopamine deficiency
  - » Secondary: iron deficiency, pregnancy, uremia, medication, diabetes mellitus, venous insufficiency.
- Treatment
  - » Replace iron, dopamine agonists, others

# ARS Question

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This is a 26 yo man with diarrheal illness 2 weeks ago. Distal paresthesias started 2 days ago, followed by leg weakness yesterday and today he can't get out of bed. He denies cognitive or bowel/bladder difficulties. His exam is remarkable for poor smile, proximal and distal weakness of the arms and legs, and areflexia. Eye function and pupils are normal.

You make a diagnosis and start treatment with?

- A. No treatment is helpful
- B. IVIG or plasmapheresis
- C. Steroids or IVIG
- D. Steroids
- E. Botulism immune globulin

# Guillain-Barre Syndrome

## □ Clinical

- » Paresthesias
- » Ascending weakness
- » Face
- » Mostly symmetric
- » Absent DTRs
- » Autonomic
- » Diarrhea - Campylobacter

## □ Studies

- » High CSF protein
- » Slow nerves (AIDP type)

## □ Course

- » Max weakness < 4wks
- » Slow recovery
- » 85% do very well
- » 10% disabled
- » 5% die

## □ Treatment

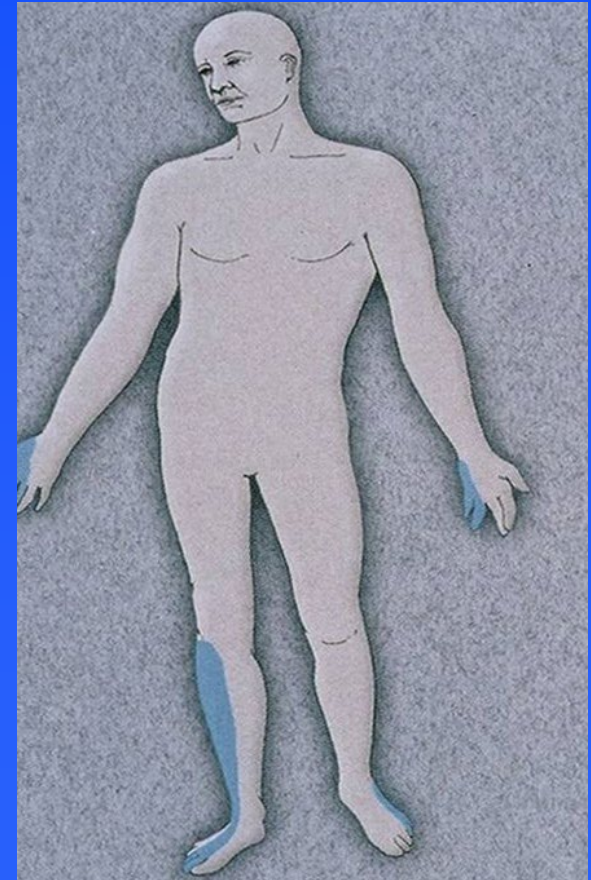
- » Avoid complications
- » Plasmapheresis
- » IVIG

# ARS Question

This is a 56 yo woman with acute painful sensorimotor symptoms over 6 days in all 4 extremities: R median, L ulnar, R peroneal, and L sural.

Your work-up should focus on what type of neuropathy?

- A. Monoclonal protein neuropathy
- B. B12 deficiency
- C. Diabetes mellitus
- D. Drug-induced neuropathy
- E. Vasculitic neuropathy



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Questions?

# Answers: ARS

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Slide 2: B

Slide 3: D

Slide 12: D

Slide 15: D

Slide 19: A

Slide 22: D

Slide 26: A

Slide 28: B

Slide 30: A

Slide 33: D

Slide 35: B

Slide 37: E