


**FERNE / MEMC IV Brain Illness and Injury Course:
Diplopia & Edrophonium Testing
Jonathan A. Edlow, MD, FACEP**

**Evaluation of Patients
with Diplopia and the
Use of the
Edrophonium Test**

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**FERNE Brain Illness
and Injury Course**

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Disclosures

- ACEP Clinical Policies Committee
- Member, FERNE
 - FERNE support by Abbott, Eisai, Pfizer, UCB


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Clinical Case

- 48 year-old woman with episodic diplopia for 1 month, worse and more frequent over the prior 4 days. She also noted intermittent bilateral ptosis.
- Physical exam showed normal vital signs and general exam. Neurological exam showed . . .

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Key findings

- Normal pupils
- Right lid ptosis worse with prolonged upward gaze, better with rest
- Vertical diplopia is worse with prolonged upward, right-ward gaze
- Both lateral and medial rectus muscles tested normal

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Learning Objectives and Key Clinical Questions

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Session Objectives

- Discuss the differential diagnosis of non-traumatic binocular diplopia
- Review the anatomy and function of cranial nerves 3, 4 and 6
- Know how (and when) to perform the edrophonium test to help diagnose myasthenia gravis

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Key Clinical Questions

- Is the diplopia isolated (or are there other neurological findings)?
- Is there evidence of a cranial nerve palsy?
- When should I consider performing a edrophonium test?

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Key Learning Points

- Consider myasthenia gravis when
- Neurological findings that fluctuate
- Findings that are hard to localize
- Prominent cranial nerve symptoms

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Differential Diagnosis

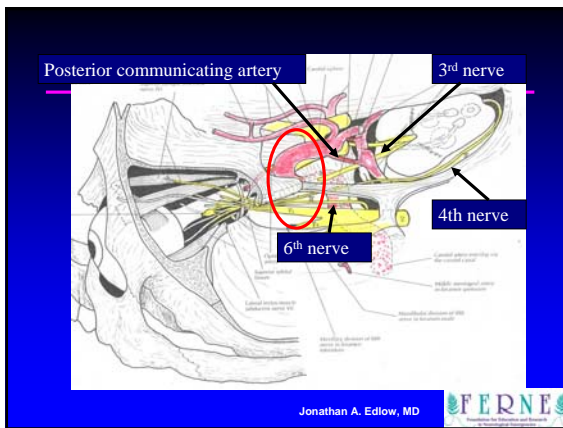
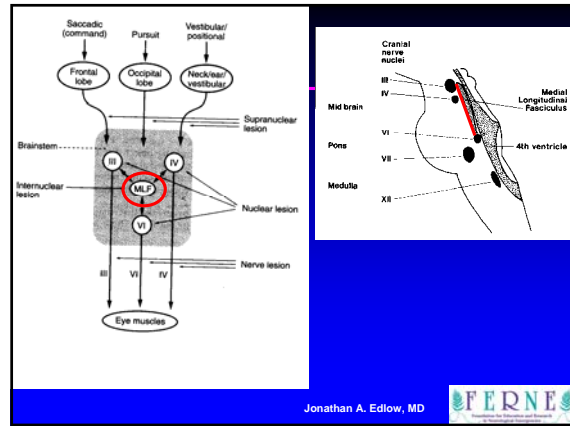
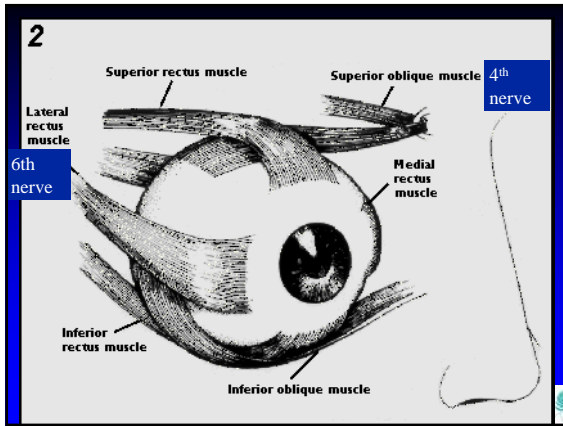
(Binocular, non-traumatic) Diplopia

- Most commonly a cranial neuropathy (either peripheral or nuclear)
- Internuclear - INO
- Supranuclear cranial nerve problems
 - Wernicke's, complex migraine, others
- Myasthenia gravis, thyroid disease
- Others
 - Botulism
 - Orbital pathology (tumor, infection or inflammation)

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Right 3rd nerve palsy

- Ptosis
- Dilated pupil
- In neutral gaze, the eye is "down and out"
- Eye will not track medially, upward

Is the pupil involved?

2.2a Enlarged Section of Third Nerve

"diabetic" 3rd nerve – pupil is spared

"surgical" 3rd nerve – pupil is involved

Left 4th nerve palsy

Head tilts TOWARDS the side of the lesion

Right 6th nerve palsy

6th nerve palsy is not of localizing value;
 It is very sensitive to ↑ ICP, meningeal inflammation

Neutral gaze
A

Right gaze
B

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Key clinical findings

- Fluctuating diplopia
- Bilateral ptosis
- Normal 3rd, 4th and 6th nerve function
 - Normal pupils
 - Normal horizontal & vertical gaze
- No other neurological findings

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Edrophonium Test

- Need a “testable” muscle
- Ideally have a blinded observer & use saline to “double-blind” the test
- Have atropine available in case of adverse reaction
 - bradycardia
 - respiratory distress
 - syncope

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Edrophonium Test

- Prepare 2 syringes
 - 10 mg of edrophonium
 - Saline (equal volume)
- Inject 2 mg of drug over 15”
- Inject the other 8mg (if no response)
- Results visible ~ 45” (and last ~ 5’)

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Edrophonium test

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What does it mean?

- Edrophonium test *
- Positive test – LR 15 (7.5-31)
- Negative test – LR 0.11 (0.06-0.21)

- Ice test *
- Positive test – LR 24 (8.5-67)
- Negative test – LR 0.16 (0.09-0.27)

* Sherer K; JAMA; 2005

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Medication triggers for myasthenia

- **Antiarrhythmic agents**
 - Quinidine
 - Procainamide
- **Antibiotics**
 - Aminoglycosides
 - Quinolones**
- **Antihypertensive agents**
 - Beta blockers
 - Calcium channel blockers
- **Magnesium-containing compounds**
 - Magnesium sulfate and citrate
- **Neuromuscular blocking agents**
 - Succinylcholine**
 - Curare derivatives

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Conclusions

- Myasthenia gravis often presents
 - with odd assortments of neurological symptoms, often **cranial neuropathy**
 - but that often **don't localize** into a neat neuro-anatomic territory
 - and which **fluctuate** over time
- Hardest part of diagnosing myasthenia is thinking of it (only ~ 60% patients diagnosed <1 year of onset)

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Recommendations

- Look for cranial nerve 3, 4 or 6 palsy
- Think of myasthenia gravis in patients with fluctuating or non-localizing symptoms
- Consider doing an edrophonium test (or ice test) to increase diagnostic accuracy

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Case resolution

- Myasthenia gravis is diagnosed
- Chest CT negative for thymoma
- Started on oral pyridostigmine
- Excellent response and has remained asymptomatic for 2 years

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Questions?

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