

Stony Brook University Medical

#### Pre-Operative Services Teaching Rounds 13 April 2011

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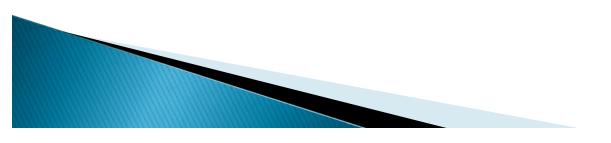
Home of the best ideas in medicine

## Pseudo-cholinesterase deficiency:

- Patho-physiology
- Clinical presentation
- Diagnosis
- Management
- Contra-indicated drugs

## Knee arthroscopy procedures:

- Anesthesia
- Positioning
- Post op analgesia



## Case

63 yr old lady for knee arthroscopy in ASC PMH:

HTN, Obesity, GERD, OSA dx 4 months ago, not using CPAP <u>PSH:</u>

Lap chole 2 years ago "difficulty waking up"

<u>FH</u>:

Difficulty breathing after anesthesia, 'allergy to drug' <u>Meds:</u>

Omeprazole, Atenolol, thiazide

<u>Exam:</u>

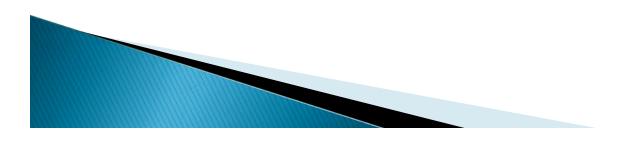
BMI 37, BP 140/90 P 65. No cardiac failure. Nonremarkable.

Airway: MP 3, FROM, 'thick neck', no dental work/loose teeth.

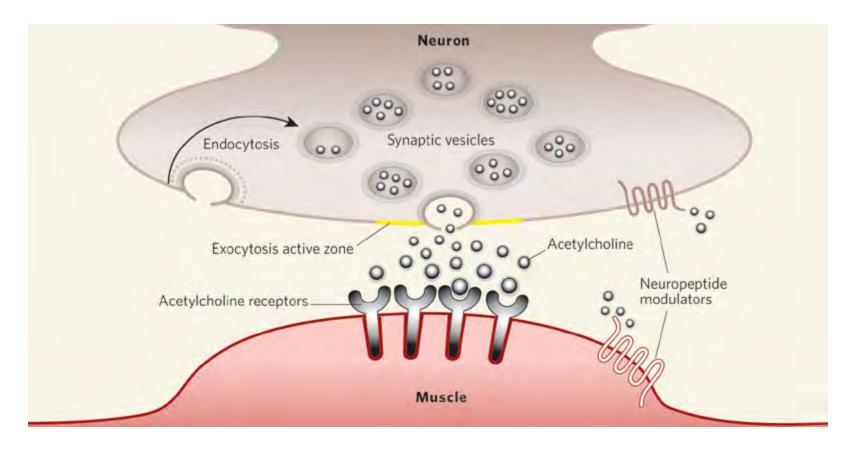
## Case discussion

Differential diagnosis:

- OSA
- Heavy handed anesthesiologist
- "Naïve liver"
- Pseudocholinesterase deficiency



## Physiology of the neuromuscular junction

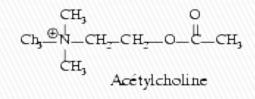


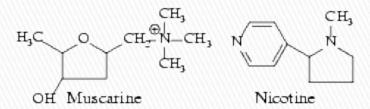
Cholinergic receptors Nicotinic type



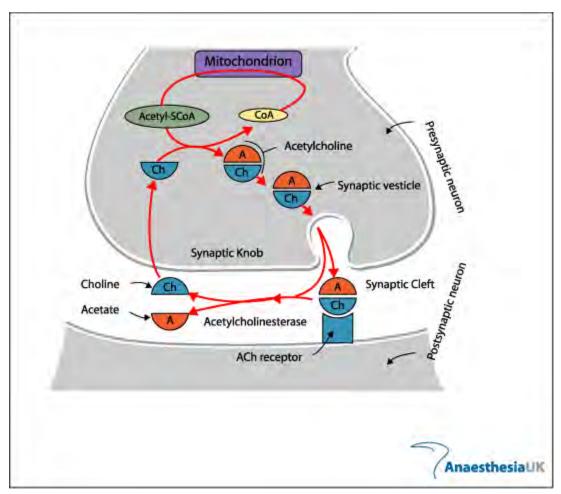
## Neurotransmitter

#### Acetylcholine





## Neuromuscular transmission

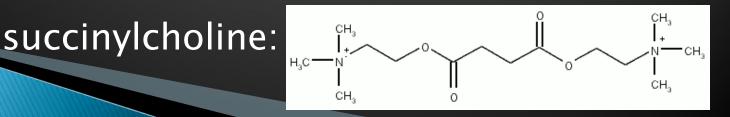


Acetylcholine is catalysed by hydrolysis to choline and acetic acid by acetylcholinesterase in neural synapse

## Muscle relaxants

Mimic acetylcholine

- Act by binding Ach receptors
  - Depolarising
    - Succinylcholine/suxamethonium
      - Metabolized by pseudocholinesterase
  - Non-depolarising
    - Pancuronium/vecuronium/rocuronium/atracurium
      - Metabolized liver/kidney/Hoffman degradation
    - Mivacurium
      - Metabolized by pseudocholinesterase



## Muscle relaxants

Indications:

- Facilitate intubation
- Facilitate surgery
  - Abdominal wall relaxation
- Facilitate ventilation
  - Laparoscopic/trendelenberg
  - ICU
- Prevent movement
  - Ophthal/neurosurgery
- Succinylcholine
  - Rapid acting
    - Rapid sequence induction
    - Difficult mask
    - Obesity
    - Short procedure

## Pseudocholinesterase enzyme

(plasma or butrylcholinesterase)

- glycoprotein enzyme
- produced by the liver
- circulates in plasma
- hydrolyzes exogenous choline esters
  - Succinylcholine
  - Ester local anesthetics
  - mivacurium
- no known physiologic function
- clinically significant if >75% reduction in enzyme activity

Acetylcholinesterase (True cholinesterase)

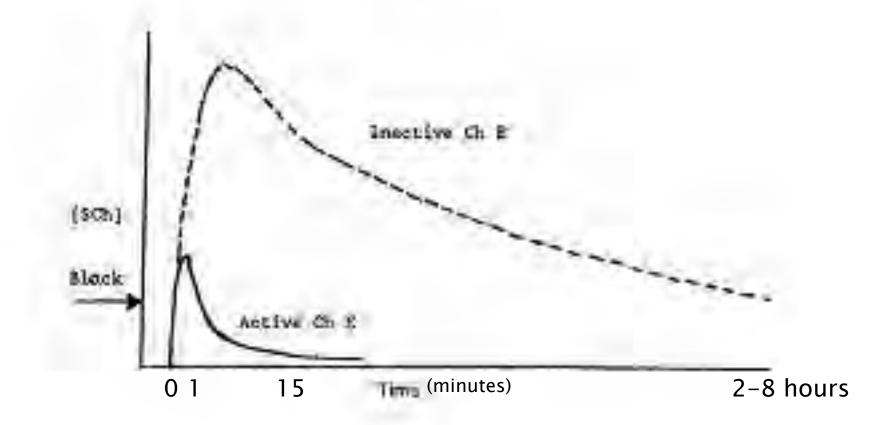
- found in neuro-musc junction
- acetylcholine metabolism and nerve function



## Succinylcholine metabolism in normal cholinesterase activity

- 90% metab in blood before reaching NMJ
- <10% clinical effect</p>
- Agonist at post junctional endplate
- Reversible
- in equilibrium across junction/plasma

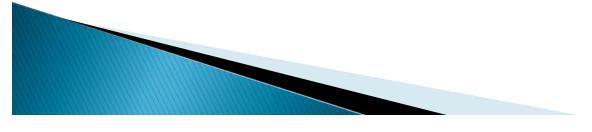






# <u>Acquired</u> abnormal pseudocholinesterase activity

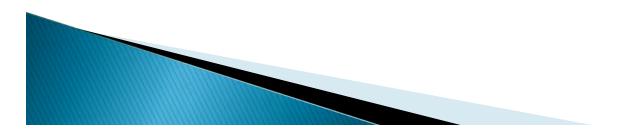
- Liver disease
- Pregnancy
- Neonates/elderly
- Malnutrition
- Chronic infections
- Extensive burns
- Organophosphate poisoning
- Uremia
- Medications
  - Echothiopate
  - neostigmine



## <u>Inherited</u> plasma enzyme deficiency or decreased activity

## 'BCHE' gene that codes for the pseudocholinesterase enzyme

- $\circ~$  located on E1 locus on the long arm of chromosome 3  $\,$
- 96% of the population is homozygous for the normal pseudocholinesterase genotype – EuEu.
- Remaining 4% of the population carries one or more of the atypical gene alleles for the pseudocholinesterase gene in either a heterozygous or homozygous fashion.
- $\circ$  <0.1% are homozygous for the abnormal gene



- Eu normal
- Ea Atypical dibucaine-resistant variant
- Ef Fluoride-resistant variant
- Es Silent variant (absent enzyme)

These alleles may occur either in the homozygous form or in any heterozygous combination with each other, with the normal Eu allele, or with a number of additional rare variant abnormal alleles

C5 variant - increased activity - shortened succinylcholine time

More common in Europeans than Asians Hindu Arya Vysya community in India

EuEu – normal activity

Heterozygous EuEa succinylcholine activity 50–100% < 1 hour EaEa succinylcholine activity 4–6 hours EsEs succinylcholine activity 8 hours (no activity)

## Clinical presentation of pseudocholinesterase deficiency

- Family/personal history "scoline apnea"
- No physical exam findings
- Prolonged paralysis after administration of succinylcholine
- Decreased or absent twitch height on nerve stimulator



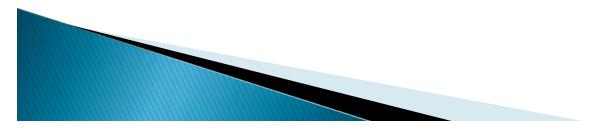
## Paralyzed and awake

#### Signs and symptoms of awareness

- Movement
- Tachycardia
- Hypertension
- Sweating
- $\circ$  Lacrimation

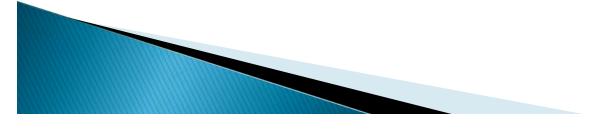
Movement

- Diaphragm
- Limb twitching 'floppy fish'



## Awareness

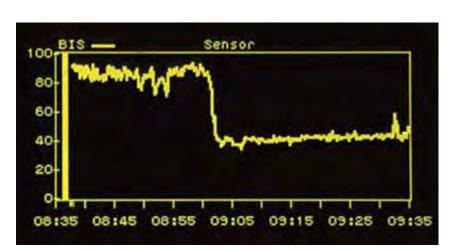
- Hearing most common remembered sense
- Pain most frightening
- ETT unable to speak also common
- Big concern
- Can recur
- Recently in the press "Awake" movie
- Redheads
- PTSD
- Reassure patients
- Communicate with anesthesia personnel
- BIS monitoring



## **BIS** monitoring

- Modified EEG
- Not a standard monitor
- Readily available
- 90 awake
- 40 anesthetized



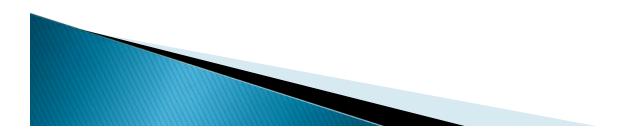




Diagnosis of pseudocholinesterase deficiency

Clinical setting

Nerve stimulator

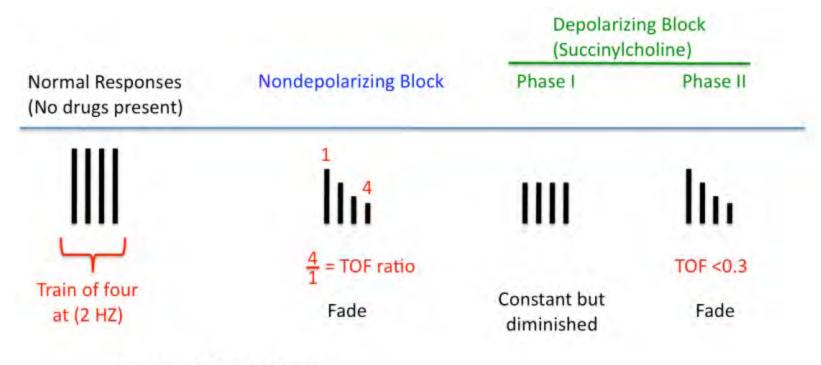


## Nerve stimulator



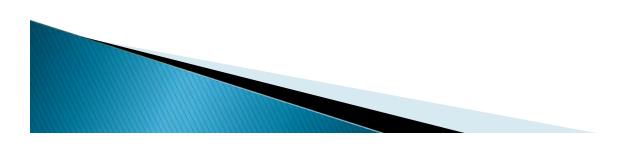






Common TOF Guidelines:

TOF 0.15-0.25: indicates adequate surgical relaxation TOF >0.9: needed for safe extubation & recovery after surgery



## Diagnosis

Fig 1: Act	holest test paper
Reaction Time	Pseudocholinesterase Enzyme Activity
<5 min	Above normal
5–20 min	Normal
20-30 min	Borderline low
>30 min	Below normal

- a)Pseudocholinesterase level
  - Not accurate after succinylcholine
  - Decreased in organo-phospate poisoning
- <u>b)Dibucaine number</u> (Salt Lake City batches 10 days)
  - Dibucaine inhibits Pseudocholinesterase activity
  - 80% inhibition = normal genes
  - 40-60% heterozygous
  - 20% inhibition homozygous

- $\circ$  Varying penetrance
- (Also fluoride inhibition 60%=normal /36%= defic)

### Treatment

#### <u>Acute event</u>

- Supportive
  - Ventilate
  - Anesthetize
  - Consider FFP
  - Usually reverse by 8 hrs

#### <u>Preoperative</u>

- Alert anesthesia
  staff (OR booking)
- Avoid specific drugs
- Reassure patient
- Dibucaine number

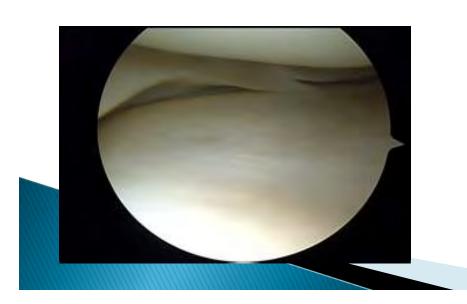
## Drugs to avoid

- Succinylcholine
- Mivacurium
- Ester local anesthetics
  - $\circ$  procaine
  - $\circ$  cocaine



## Arthroscopy

- Supine
- Low risk procedure
- Tourniquet
- Hip movement
- DVT risk





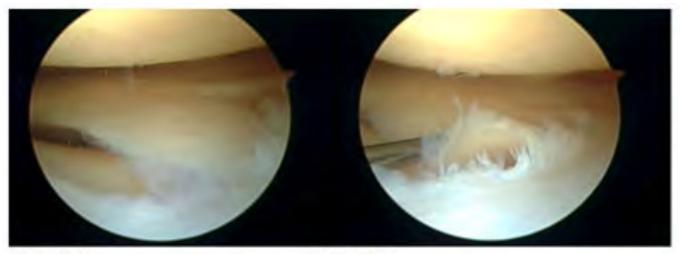
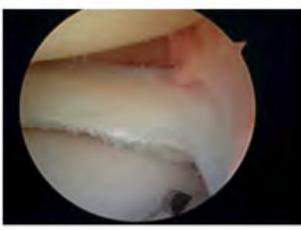


Figure 3A.

Figure 3B.







Anterior cruciate ligament Creative Commons- wikimediacommons

## **Open procedures**

- Supine
- Tourniquet
- Hip movement
- Intermediate risk procedures
- DVT risk



## <u>Anesthesia</u>:

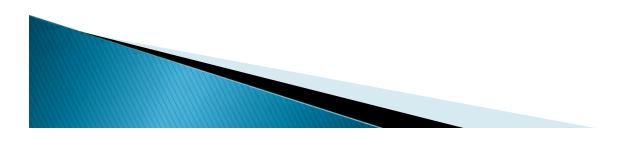
- General earlier discharge
   Spinal
  - ?better in OSA
- 3. Intra-articular
  - sedation for:
  - hip movement
  - tourniquet pain

operator dependent

(Segreto in ASC only one using intra-articular anesthesia with sedation)

## Post op analgesia

- Femoral nerve block/catheter for analgesia if big repair/graft
- Ice packs patients to fill prescription to bring their own



## Take home points

#### Pseudo- cholinesterase deficiency

- Test patient, carry on with surgery
- Notify OR booking
- Avoid succinylcholine, mivacurium
- Alert patient and PCP with result

#### Knee surgery

- GA
- Book early for spinal
- Bring icepack