Paracetamol Naloxone Opkast Kul

- HVAD VED VI?

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.....nothing to disclaim

- Paracetamol
 NAC treatment is not indicated when s-paracetamol
 is zero
- Naloxone

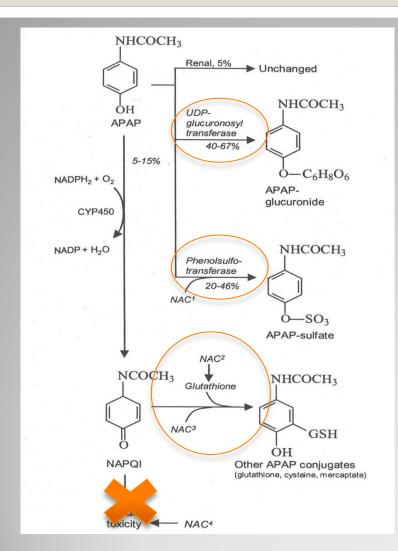
After reversal of an opioid overdose the patient MUST always be admitted for observation for at least 4-6 hours

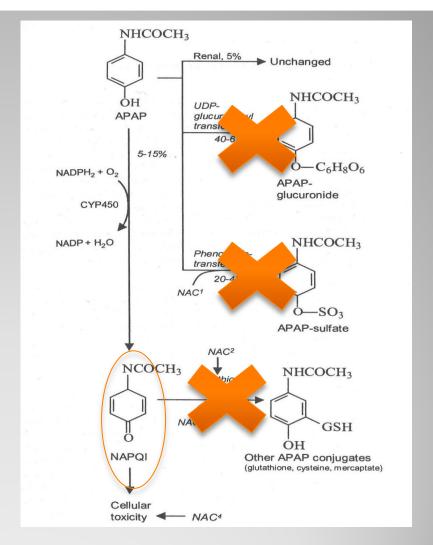
- Activated charcoal
 Is only effective if administred within 1 hour of drug ingestion
- Induced emesis (vomiting)
 Syrup of ipecac is an effective way of gastric decontamination

...what we know.....myths?



Paracetamol - initial assesment





Paracetamol poisoning

Toxic hepatitis

- Symptoms developes with a latency of 1-3 days
- Day 1 nausea and vomiting may occur, but otherwise the patient is unaffected and normal has biochemistry
- Day 2 increases in transaminase levels, INR and s-creatinin
- Day 3 progressing liver and renal failure
- ANTIDOTE N-Acethylcystein (NAC) stimulates gluthation production

Paracetamol poisoning

- S-paracetamol = 0
- Transaminase = normal
- Parentdrug not metabolites



- NAPQI T½ unpredictable in overdose
- Substantial amounts of toxic metabolites
- Before hepatic damage is measurable

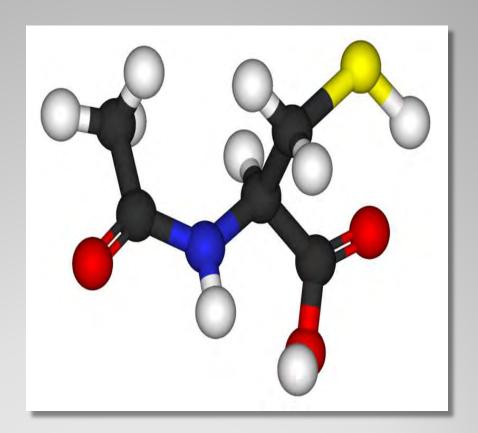
Suspicion!

- Unknown dose or large dose
- Start upon arrival regardless of time interval from ingestion
- Don't wait for S-paracetamol
- Lever function tests
 may be normal even in severe cases



NAC - indications

- Rash
- Nausea
- Vomiting
- Cramps
- Diarrhea



Angioedema (edema or swelling of the skin)

NAC is SAFE

 Don't wait for S-paracetamol – treat if suspected ingestion

NAC is SAFE

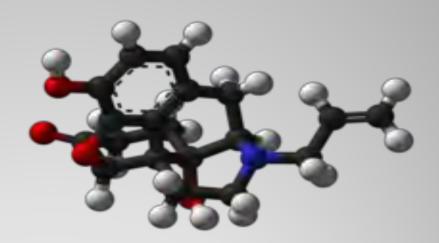
 If paracetamol poisoning is confirmed, highly likely or cannot be ruled out treat with NAC even if s-paracetamol is ZERO

Paracetamol - conclusion



Naloxone

- Primarily competitive µ-receptor antagonist
- Rapid reversal
- Duration 1-4 hours



 After reversal of opioid overdose the patient MUST always be admitted for observation for at least 4-6 hours

ref - Tintinalli J. Emergency medicine - a comprehensive study guide, 6the edition

Naloxone

Contact and death pattern among heroin users after initial contact with a MECU in Copenhagen

 To describe the development and pattern in the number of overdoses in Copenhagen during a 10 year period and to examine 48h and 1 year mortality rate



Naloxone

S. Rudolph, K. Nielsen and S.L. Nielsen. Contact and death pattern among heroin users after initial contact with a doctor staffed mobile emergency care unit (MECU) in Copenhagen Resuscitation; Volume 77, Supplement 1, May 2008, Page S69

- The unresponsive patient
 - 0,8 mg naloxone IV
 -in some cases 0,4 mg IM/SC supplement
- The reponsive but obtunded patient
 - naloxone is titrated to effect
- The patient is released on scene if a substantial and lasting improvement is obtained and mental status, hemodynamics, and pulse oximetry results are acceptable

Release-on-scene

- The MECU had 2854 contacts with 1731 patients for opioid overdose between 1994-98 and 2000-2003
- In 5.7 % (n=99) cases the patients had another MECU contact within 48 hours
- 14 patients (0.8 %) died within 48 hours of a MECU contact and a post treatment release on-scene

Naloxone

Autopsy reports

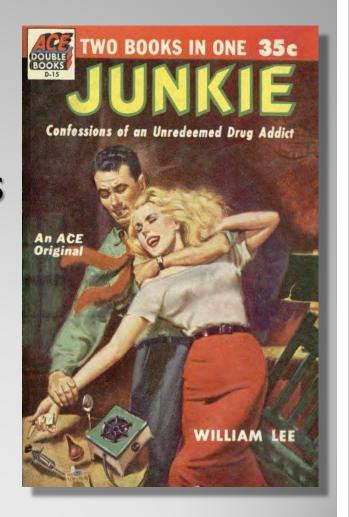
- 6 found with a needle in the arm
 - considered a new overdose

- 5 found with a single fresh needle mark 2 with multiple - but no needles or other drugs
 - inconclusive
 - likely new or possible rebound overdose
- 1 found with no signs of new overdose
 - considered possible rebound overdose

Naloxone - 14 deaths.....

TOX SCREENING

- in 5 of 7 inconclusive cases METHADONE was the suspected drug
- new overdose?
- rebound overdose?



Type of opioid

- Reversal of heroine/morfin overdose and release on-scene is safe
- Only in 0,4 % of cases a rebound overdose was considered possible
- Consider admittance if Methadone



Naloxone - conclusion

American Journal of Emergency Medicine (2006) 24, 567-572



The American Journal of Emergency Medicine

www.elsevier.com/locate/ajem

Original Contribution

Does naloxone alone increase resuscitation rate during cardiopulmonary resuscitation in a rat asphyxia model?

Meng-Hua Chen MD^{a,*}, Tang-Wei Liu MD^a, Lu Xie DPharm^b, Feng-Qing Song MM^a, Tao He MM^a

Naloxone alone can increase resuscitation rate following asphyxial cardiac arrest in rats.

Naloxone - cardiac arrest?



- Gastric decontamination
- Superb adsorptive properties
- Reduces systemic absorption
- Enhances elimination by interuption of the enterohepatic cycle for some drugs
-theoretically the single most useful treatment of acute oral overdose



Activated charcoal

- Lack of sound evidence of its benefits as defined by clinically meaningful endpoints
- " a single dose of activated charcoal should not be administered routinely"
- "the administration of activated charcoal may be considered if a patient has ingested a potentially toxic amount of a poison (which is known to be adsorbed to charcoal) up to one hour previously"

Position statement: single-dose activated charcoal. American Academy of Clinical Toxicology; European Association of Poisons Centres and Clinical Toxicologists. Clinical Toxicology, 43:61–87, 2005

Activated charcoal - time factor



- Most patients do not present to the ED within 1 hour
- In 63 patients
 - median arrival time 136 minutes
 - only 15 presented withinhour
 - 4 of 10 who qualified for treatment received charcoal within 1 hour

Problem

Karim A, Ivatts S, Dargan P, Jones A: How feasible is it to conform to the European guidelines on administration of activated charcoal within one hour of an overdose? *Emerg Med J. 2001;18:390-392.*

 Meta-analysis - data from 64 controlled studies

• Evaluate:

- the effect of activated charcoal on xenobiotic absorption during the first 6 hours after ingestion
- influence of physical and pharmacologic properties

Activated charcoal - time factor

Jürgens, G. et al. The Effect of Activated Charcoal on Drug. Exposure in Healthy Volunteers: A Meta-Analysis. *Clin Pharmacol Ther.* **85**, 501-5 (2009)

 Most effective when administered immediately

BUT....

 4 hours after ingestion, 25% of the participants achieved at least a 32% reduction in absorption - especially when activated charcoal was given with large charcoal-to-drug-ratios

Activated charcoal - metaanalyse

- An 18-month consecutive case series
 - activated charcoal can be administered successfully in the home by the lay public
- Significantly reduced the time to treatment

- ED: mean of 73 ± 18.1 min

- Home: mean of 38 ± 18.3 min



 However, many still consider this evidence insufficient to recommend that activated charcoal be stored in the home.

Activated charcoal - at home

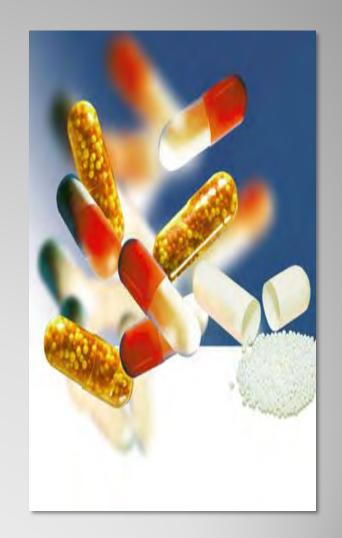
Spiller HA, Rodgers GC Jr: Evaluation of administration of activated charcoal in the home. *Pediatrics. 2001;108:E100.*

- Prospective follow-up study from Finland
- Activated charcoal by either the firstresponse unit or paramedics
- Activated charcoal was administred to 555 patients with a mean of 88 minutes after ingestion
- No adverse effects noted, although 72 patients refused to drink the charcoal slurry
- Feasible to administer activated charcoal in the prehsopital setting

Activated charcoal - prehospital

Alaspää AO, Kuisma MJ, Hoppu K, Neuvonen PJ: Out-of-hospital administration of activated charcoal by emergency medical services. *Ann Emerg Med.* 2005;45:207-212.

- Activated charcoal should be given routinely up til 4 hours after ingestion
- The 1 hour limit is a guideline more than a cut off point
- In life-threatning poisonings, activated charcoal should be given regardless of time of ingestion
- Consider prehospital use



Activated charcoal – conclusion



Syrup of ipecac

- Gastric decontamination
- Acts locally and centrally



• 1997 Position statement

American Academy of Clinical Toxicology European Association of Poisons Centres Clinical Toxicologists

 No additional useful materials were found



Syrup of IPECAC

Krenzelok EP, McGuigan M, Lheur P: Position statement: ipecac syrup. *J Toxicol Clin Toxicol*.1997;35:699-709.

 Time to performing GI decontamination is critical

Delaying factors

- Time delay from administration to onset of emesis – mean time 58 minutes
- Uncertainty of the effect of the administered dose
- Ipecac can cause sedation

Syrup of IPECAC – a delay?

Indications

- The patient meets criteria for gastric emptying.
- Orogastric lavage cannot be performed or is contraindicated because of the size of the formulation of xenobiotic
- Likely significant amount of xenobiotic in the stomach
- The benefits outweigh the risk from the contraindications

Contraindications

- The patient does not meet criteria for gastric emptying
- Activated charcoal is expected to be nessesary in the next several hours
- Airway maybe lost in the next hour
- Caustic ingestion
- Foreign body ingestion
- Drugs of high aspiration potential
- Infant < 6 months
- Premorbid conditions that be decompensated

Syrup of IPECAC

Goldfrank's Toxicologic Emergencies

Indications

- VERY FEW !!!!!
 - only recommended once in 2009!
- Xenobiotics that don't adsorb to charcoal
 - Iron and Lithium
- Only children
 - to small for NG tube for aspiration of pills

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Syrup of IPECAC

Giftliniens pratical guideline

- No clinical benefit vs the recognized benefit of activated charcoal
- Administration of syrup of ipecac delays the administration of activated charcoal
- Given the lack of evidence demonstrating a clinically meaningful benefit and the significant contraindications, the routine administration of syrup of ipecac should be abandoned
- Contact your local poison center

Syrup of IPECAC - conclusion

- Don't wait for S-paracetamol treat if suspected ingestion even in the face of normal transaminase levels
- A release on scene is safe in severe heroine/morfine overdose
- Activated charcoal should be given routinely uptil 4 hours after drug ingestion
- Induction of emesis has an extremely limited role in the contemporary management of poisoned patients and should probably be abandoned

so....in conclusion

