Introduction to Toxicology

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Topics

- Definition
- Terminology
- Classification of Toxic agents
- assessment
 - history
 - Examination
 - investigation
- Management
- Disposition
- Poison center No.

Definition

a science that deals with the adverse effects of chemicals on living organisms and assesses the probability of their occurrence

Why people get toxic?

intentional i.e. suicide

wrong dose (i.e. Insulin)

symptoms control (i.e. paracetamol for pain)

exposure i.e. radiation, organophosphate

bite i.e. snake bite

what are the routes of exposure?

inhalation (i.e. Nitrous oxide, CO)

skin or eye absorption (i.e. organophosphate)

ingestion: major one (i.e. paracetamol....etc)

injection (i.e. Opioids, insulin)

Assessment

History

may be unclear

substance

dose

rout of exposure

collateral Hx (i.e. family, friends, medical records)

Prehospital medical staff (i..e empty containers)

other (i..e hobbies, occupation, suicide note, change in behaviour recently)

Examination

Organ system	example of finding
General appearance	Malnurished (IV drug user, HIV infection)
CNS	Miosis (Opioids, organophsophate) Nystagmus/ataxia (ethanol)
CVS	Murmur (Endocarditis/IV drug user)
Respiratory system	Bronchorrhea/crepitations/hypoxia (Organophosphate)

Examination

Organ system	Example of finding
GIT	oral cavity burns (corrosive ingestion hyper salivation (cholinergic toxidrome
Urology	urinary retention (anticholinergic toxicity)
Peripheral nerves	tremor (Lithium) Lead pipe rigidity (NMS) clonus/hyperreflexia (serotonin toxicity)
Dermal	bruising (anticoagulant) flush, dry skin(anticholinergic toxicity) warm, moist skin(sympathomimetic toxicity)

Do not forget ...!

examine skin folds, clothes and bags for retained tablets or substances



Toxidrome

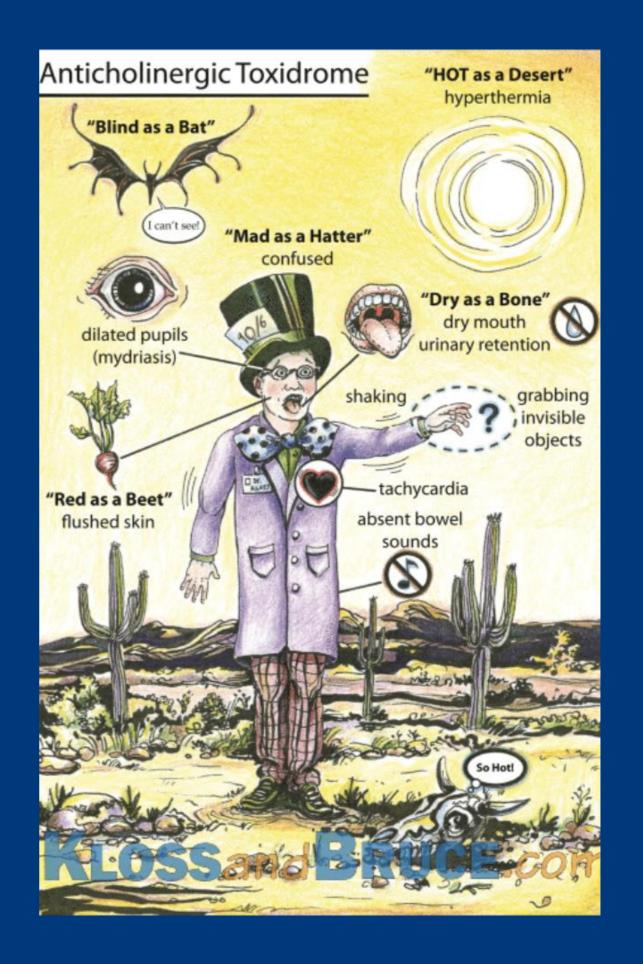
Cluster of symptoms and signs

enabling the identification of potential toxins when a clear history is unavailable

Anticholinergic = Antimuscarinic

clinical features	agents	potential interventions
 altered mental status mydriasis dry flushed skin urinary retention decreased bowel sounds hyperthermia (cause of death) dry mucus membrane 	Atropine scopolamine TCA Olanzepine antihistamine diphenhydramine	physostigmine benzodiazepine for sedation (MCQs) cooling supportive management
other • seizure • rhabdomyolysis • arryhythmia		

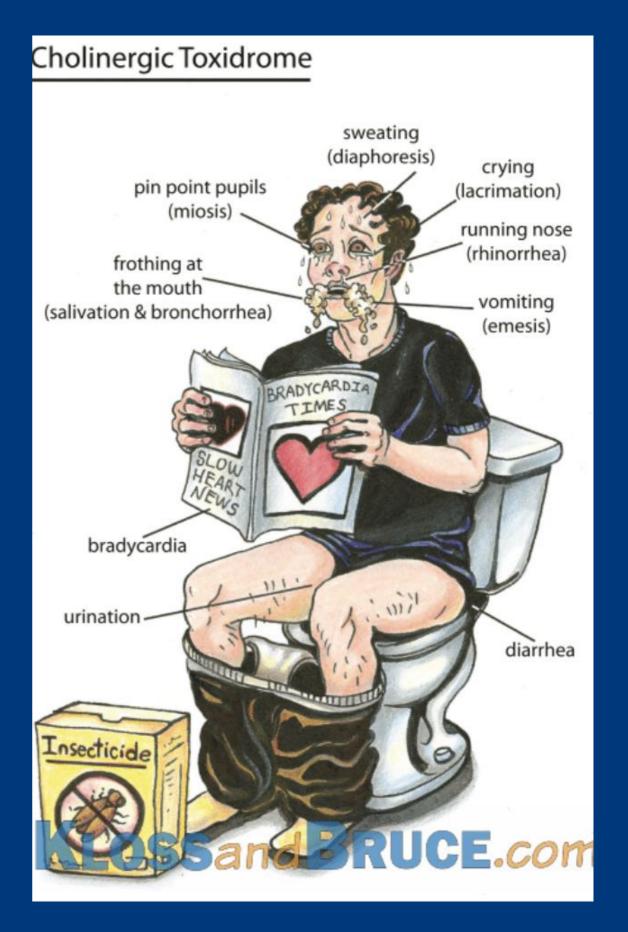
Anti-cholinergic



Cholinergic = Muscarinic

clinical features	agents	potential interventions
muscarinic effect salivation lacrimation diaphoresis nausea vomiting urination defecation bronchorrhea nicotinic effect muscle fasciculations weakness	organophosphate insecticides carbamate insecticides	Airway protection + ventilation atropine pralidoxime
other • bradycardia • miosis/mydraisis		
death—> respiratory arrest from muscle paralysis		

Cholinergic



Sympathomimetics

clinical features	agents	potential interventions
 psychomotor agitation mydriasis diaphoresis tachycardia hypertension hyperthermia 	Amphetamine cocaine	cooling sedation with benzodiazepine hydration
other • seizure • rehabdomyolysis • MI		
Death—> seizure, cardiac arrest, hyperthermia		
NB / very close to anticholinergics but the difference in Diaphoresis		

Opioid

clinical features	agents	potential intervention
CNS depression respiratory depression miosis	Heroin morphine oxycodone	Naloxone +/- airway support and ventilation
others - hypothermia - bradycardia		
death from respiratory depression		

Sedative-hypnotic

clinical features	agents	potential interventions
depressed LOC ataxia slurred speech respiratory depression bradycardia	benzodiazepines barbiturate	ventilatory support

Hallucinogenic

clinical features	agents	potential intervention
 hallucinations dysphoria anxiety hyperthermia mydriasis nausea +/- sympathomimetics 	phenocyclidine Lysergic acid diethyl amide psilocybin mescaline	supportive

Other toxidromes

Toxidrome	Examination finding
Hypoglycemic(i.e.insulin)	altered mental status, diaphoresis, tachycardia, HT
Serotonin (i.e.SSRIs)	altered mental status, hyperreflexia, hypertonia(LL>UL), clonus, tachycardia
Neuromuscular Malignant(i.e.antipsychotics)	sever muscle rigidity, hyperpyrexia, altered mental status
Extrapyramidal (i.e.haloperidol)	Dystonia, torticollis, muscle rigidity
Ethanol	CNS depression, ataxia, dysartheria, smell of ethanol
Salicylate(i.e. Aspirin)	AMS, Resp Alkalosis, Metabolic Acidosis, Tinnitus, Tachypnoea, Tachycardia, diaphoresis, nausea vomiting

Bedside:

Blood Glucose level: hypoglycaemia

ECG: Arrhythmias

VBG: i.e. metabolic acidosis —> paracetamol

Laboratory:

blood / urine drug level

TABLE 176-5	Drug Concentrations That May Assist Patient Assessment or Management	
Acetaminophen	Methanol	
Carbamazepine	Methotrexate	
Carbon monoxide	Paraquat	
Digoxin	Phenobarbital	
Ethanol	Phenytoin	
Ethylene glycol	Salicylate	
Iron	Theophylline	
Lithium	Valproic acid	
Methemoglobin		

what are the line it ations of Drug Screening Assays Says?

TABLE 1/0-0	Limitations of Toxicologic Drug Screening Assays	
Nonspecific	Most tests use enzyme-immunoassays that only detect typical drugs within a class: opioids, amphetamines, benzodiazepines, cannabinoids, cocaine, barbiturates. Amphetamine screens do not detect methylenedioxymethamphetamine. Opioid screens do not detect meperidine. Benzodiazepine screens do not detect flunitrazepam.	
Time frame	Drugs may be detected days to weeks after exposure. A positive test may not account for current clinical findings.	
Cross-reactivity	Carbamazepine, cyproheptadine, and chlorpromazine test positive for tricyclic antidepressants. Selegiline, methylphenidate, and pseudoephedrine test positive for amphetamines.	
Noninclusive	A negative drug screen does not exclude a rare exposure.	
Sampling error	Assay may be negative if dilute urine is tested.	

Electrolytes:

K level: i.e. hyperkalemia in digoxin overdose

LFT:

elevated liver enzymes in Paracetamol toxicity

Management:

Resuscitation

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Airway:
 intubation: if compromised
Breathing:
 O2 administration, if hypoxic (i.e. O2sat <94%)
 mechanical ventilation if intubated
Circulation:
 hypotension
  IV fluid (10-20ml /Kg), avoid excess fluid administration
  specific antidote
  inotropic support (i.e.Adrenaline infusion)
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Resuscitation

Antidotes list

Poison	Antidote
Acetaminophen	N-acetylcysteine
Anticholinergics	Physostigmine
Anticoagulants	Vitamin K, FFP
Aspirin	Sodium bicarbonate
Beta blockers	Glucagon, insulin
Benzodiazepines	Flumazenil
Calcium channel blockers	Calcium, glucagon, insulin
Carbon monoxide	Oxygen
Cholinergics	Atropine, pralidoxime (2-PAM)
Cyanide	Hydroxycobalamin, amyl nitrite, sodium thiosulfate
Digoxin	Digoxin FAB
Heparin	Protamine
Heavy metals Arsenic Copper Lead Mercury	Dimercaprol EDTA Penicillamine Succimer (DMSA)
Hydrofluoric acid	Calcium gluconate
Insulin	Glucose
Iron	Desferoxamine
Isoniazid	Pyridoxine
Methanol	Ethanol
Ethylene glycol	Fomepizole, ethanol
Methemoglobin	Methylene blue
Opioids	Naloxone
Serotonin repute inhibitors	Cyproheptadine
Sulfonylurea	Octreotide, glucose
Tricyclic antidepressant	Sodium bicarbonate

Resuscitation

some specific presentations

Hypoglycemia

- BGL : < 4mmol
- give IV dextrose (Glucose)

Cardiac Arrhythmias

Anti-arrythmic drugs are not first line treatment in toxin induced arrhythmias

treatment:

O₂ sat

antidote (i.e. digoxin Fab in digoxin overdose)

Seizure

treatment

1st: IV benzodiazepine (except in Isoniazed toxicity —> Pyridoxine)

2nd: Barbiturates

treat hypoglycaemia and hyponatremia

No rule for Phenytoin in toxin induced seizure

Agitation

1st line treatment : benzodiazepine

2nd line treatment : antipsychotic agents

Hyperthermia and hypothermia

core temperature > 39* —> aggressive cooling

core temperature <32* —> aggressive rewarming

My brain is like The Bermuda Triangle...
Information goes in and then it's never found again.

Decontamination

two ways

GIT Decontamination

Enhanced Elimination

GIT decontamination

Activated Charcoal

whole bowel irrigation (WBI)

Gastric lavage

Induced emesis (Syrup or Ipecac)

Activated Charcoal (single dose)



indications	contraindications	complications	technique
 preferred method < 1 hour from ingestion charcoal sensitive substances: (MCQs) paracetamol benzodiazepines barbiturates TCA phenothiazines most anticonvulsants aspirin theophylline digoxin dextropropoxyph en amphetamines quinine morphine ciclosporin most NSAIDs beta blockers 	 incomplete initial resuscitation non toxic ingestion subtonic dose risk assessment —> good outcome with supportive care & antidote risk assessment —> potential for seizure of decrease LOC decrease LOC decrease LOC , seizure (unless Intubated) charcoal resistance agents (see below) corrosive ingestion ileus is not a contraindication	 vomiting 30% messy aspiration direct admisntration into lung if NG tube placed in lung impaired absorption of subsequent oral antidote, therapeutic agents corneal abrasion staff distraction from resuscitation and supportive priorities 	adult 50 gm children 1gm/Kg mix with water self administration if GCS 15 via OG / Ng tube if intubated (first confirm tube position with chest X-ray) no difference between mixing AC with water or other (sorbitol)

Activated Charcoal (single dose)

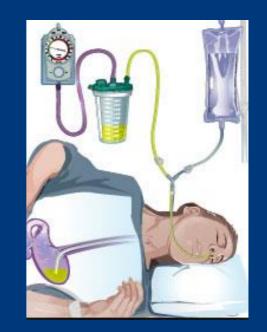
Q: what are the charcoal resistance substances?

hydrocarbons and alcohol	metals	corrosive
ethanol isopropyl alcohol ethylene glycol methanol	Ithium Iron K lead arsenic mercury	acids alkalis

whole bowel irrigation

indications	contraindications	complications	technique
iron overdose >60mg/kg lead ingestion arsenic ingestion body packers slow release preparations: lithium verapamil / diltiazem potassium formulations	 risk assessment —> good outcome with supportive care and antidote risk assessment —> potential for seizure or decrease LOC uncooperative patient inability to place NG tube uncontrolled vomiting ielus intestinal obstruction intubated and ventilated patient (relative) 	 nausea /vomiting abdominal bloating non anion gap metabolic acidosis pulmonary aspiration staff distraction from resuscitation and supportive care priorities delayed retrial to a hospital offering definitive care 	 Polyethylene glycol electrolytes solution (PEG-ELS) single nurse assigned enough supply of PEG-ELS NG tube inserted and confirmed AC charcoal administer PEG adult 2L/hr child 25ml/kg/hr give metoclopromide -> decrease vomiting and enhance motility explosive diarrhoea continue irrigation until it clear stop if abdominal distension or bowel sound lost

Gastric Lavage



indications	contraindications	complications	technique
rare in ED serious poisonings <1hr other methods are unavailable mercury ingestion arsenic ingestion	 incomplete initial resuscitation risk assessment —> good outcome with supportive care and antidote decrease LOC risk assessment —> potential for Decrease LOC during the procedure small children corrosive ingestion hydrocarbon ingestions 	 pulmonary aspiration hypoxia laryngospasm mechanical injury to GIT water intoxication (children) hypothermia staff distraction form resuscitation and supportive priorities 	 resuscitation area GCS 15 / intubated Pt left decubitus position, head down 20" pass gastric lavage tube (36-40 G) (OG route) confirm tube position (aspiration and auscultation) administer 200 ml aliquot of warm tab water or NS drain the fluid into dependent bucket repeat until its clear give AC 50 G via the lavage tube once lavage is completed

Induced emesis (Syrup or Ipecac)

indications	contraindications	complications	technique
 limited charcoal resistant poison serious risk of toxicities < 1 hour after ingestion large fragments in stomach (WBI is better) Fe sustained release lithium enteric coated tab poisonous mushrooms 	 non toxic ingestions sub toxic doses seizures Decrease LOC risk assessment —> potential for seizure / decrease LOC within the next few hours activated charcoal available within 1 hour and know to bind to the substance infant < 12 months corrosive ingestion hydrocarbon ingestion 	 prolong vomiting > 1 hr in 10-20% diarrhoea 20% lethargy 10% pulmonary aspiration if (seizure / Decrease LOC) mallory weiss tear pneumomediastinum gastric perforation 	 children —> 15 ml Adult —> 15-30 ml with glass of water usually vomit after 18 min repeat the dose if no vomit after 30 min

Enhanced Elimination

Multiple dose activated charcoal

urine alkalisation

extracorporeal technique of elimination

harm-dialysis and haemofiltration

charcoal haemoperfusion

Multiple doses of AC

indications	contraindication	complication	technique
 carbamazepine coma (most common indication) phenobarbitone coma dapsone overdose —> methaemoglobinaemia Quinine overdose Theophylline overdose phenytoin 	Decrease LOC anticipate decrease of LOC bowel obstruction	 vomiting 30% pulmonary aspiration constipation bowel obstruction bowel perforation corneal abrasion staff distraction from resuscitation and supportive care 	 give the atoll dose adult 50 g kids 1gm/kg repeat doses of adult 25gm kids 0.5g/kg every 2 hours route oral if GCS 15 NG/OG tube after position confirmed by chest X-ray check bowel sound before each dose if no bowel sound stop doses reconsider indication and endpoints every 6 hours very rare therapy continue > 6 hours

Urinary Alkalinisation

Mechanism

make urine PH alkaline —> ionisation of highly acidic drug —> decrease renal absorption & increase renal excretion

indications	contraindication	complications	technique
 Salicylate overdose phenobarbitone coma (not first line) cyanide isoniazid toxic alcohol TCA propranolol felcainide quinidine methotrexate 	fluid overload	alkalemia hypokalaemia hypocalcaemia volume overload	Sodium bicarbonate 1-2 mmol/kg IV bolus infusion @ 250ml /hr 100 mmol NaHCO3 in 1000ml 5% dextrose add 20 mol of KCL to the infusion to maintain the normokalaemia follow serum HCO3 and K every 4 hr aim urine PH >7.5 continue till the lab and clinical evidence of toxicity is resolved.

Extracorporeal technique of elimination

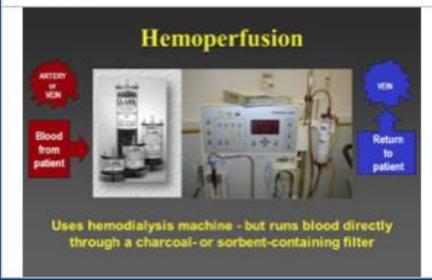
Hemodialysis

Movement of solute down a concentration gradient across a semipermeable membrane



Hemoperfusion

Movement of toxin from blood, plasma or plasma proteins onto a bed of activated charcoal (or other adsorbent)



Indications

- · sever life threatning
- deteriration despite full supportive care
 - Carbamazepine
 - Potassium overdose
 - Sodium valproate
 - metformin
 - Phenobarbitone chronic lithium
 - salycilate
 - toxic alcohol
 - methanol
 - ethylen glycol
 - theophylline

Contraindications

- · Hemodynamic instability
- Poor vascular access
- · Significant coagulopathy

Complications

- Hypotension (most common)
- bleeding from vascclar access
- air emboli
- blood loss
- · systemic heparinisation
- thromobcytopenia
- neutropenia

Technique

- invasive
- special staff
- special equipment
- monitoring

Disposition

if asymptomatic for 6 hours in ED —> discharge

otherwise admission to hospital is required

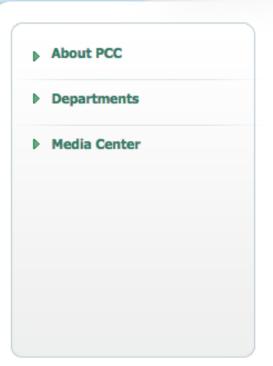






Poison Control Center - Riyadh General Directorate of Health Affairs - Medial Province

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Thank You

All the best!

