

ACETAMINOPHEN (N-ACETYL PARA- AMINIPHENOL) (APAP) TOXICITY

Practical Clinical Toxicology

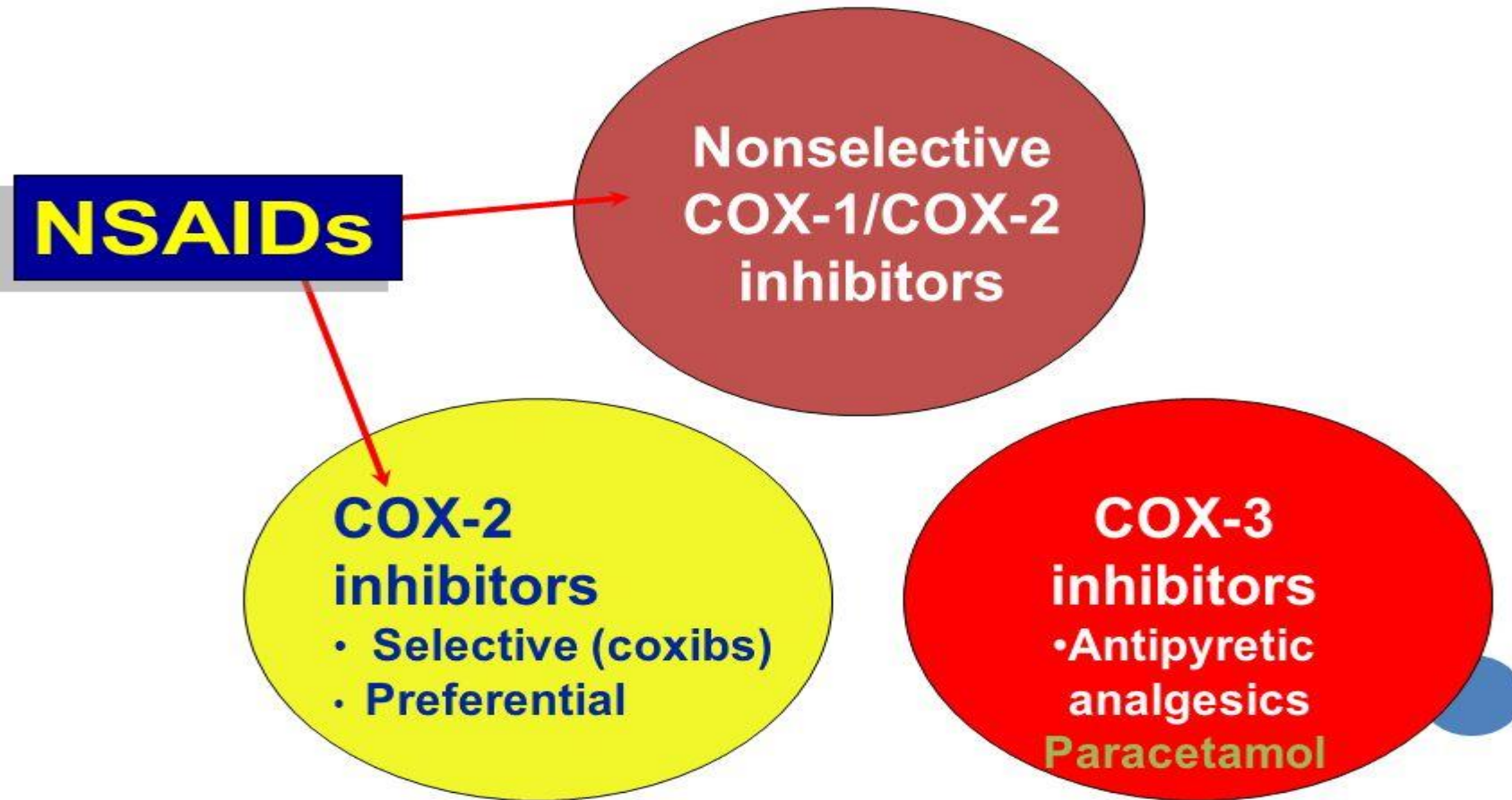
M.Sc. Nibras Jamal

M.Sc. Nada S. Rammahy

INTRODUCTION TO MECHANISM

- Acetaminophen, also known as Paracetamol, is N-Acetyl Para Amino Phenol (APAP).

Classification of COX inhibitors



EPIDEMIOLOGY

- It is contained in over 100 OTC preparations.

METABOLISM AND EXCRETION


- 90% of acetaminophen is hepatically metabolized to harmless glucuronide (60%) and sulfate (30%) metabolites excreted in urine .

TOXICITY

- NAPQI production exceed hepatic ability to detoxify NAPQI by glutathione conjugation.

ACUTE APAP POISONING

- Phase 1 (up to 24) hours
- Phase 2 (24-72) hours :

- 
- Phase 3 (72-96) hours:
 - Phase 4 (4 days-2weeks):



- **Chronic APAP poisoning**

- (Analgesic nephropathy is injury to the kidney caused by analgesic medications such as aspirin, phenacetin, and paracetamol.)



- Risk factors

- Increased NAPQI production as a result of cytoP450 enzyme induction

TOXIC DOSE

- Acute overdose is usually considered to be a single ingestion



- Because of

- 1. Age-dependent rate of glutathione turn over that can younger tolerate higher doses.

REFERENCES

- 1- Gossel TA, Bricker TD, (Eds.); Principles of Clinical Toxicology; latest edition.*
- 2- Viccellio P, (Ed.); Handbook of Medicinal Toxicology; latest edition.*
- 3- journals of pharmacology and toxicology*