Drugs in the liver (lec 1)



Drugs	features
propranolol, lignocaine opiates (like fentanyl, and morphine)	high hepatic extraction ratios >0.7
lorazepam	low hepatic extraction ratios <0.3 less affected in liver disease does not require a phase 1 reaction
diazepam	low hepatic extraction ratios < 0.3 ,highly protein-bound drugs, (benzodiazepines),
methadone.	low hepatic extraction ratios < 0.3
benzodiazepi nes, tricyclic antidepressan ts and antipsychotic.	metabolized into active metabolites (phase I)
Olanzapine, oxazepam	less affected in liver disease does not require a phase 1 reaction (Phase II)

Drugs	features	Contraindication
Digoxin, Warfarin., Antibiotics (ceftriaxone, Cefoperazone, macrolides rifampicin, and others), Mycophenolate mofetil. ,Spironolactone, Steroid hormones (e.g., estrogen), Opioids, NSAIDs (e.g., diclofenac & indomethacin), The anticancer doxorubicin	molecular weight of > 300 g/mol with both polar and lipophilic groups excreted in bile and subject to enterohepatic recycling.	NSAIDs are contraindicated for systemic use in most liver disease patients,
dexamethasone	safe in patients with chronic stable liver disease.	
methyprednisolone	in high doses reactivate HBV Increase spontaneous bacterial peritonitis in severe cases.	

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Drugs in the liver (lec 1)



Drugs	features
Carbon tetrachloride	block the secretion of triglycerides, causing fatty liver. metabolized by cytochrome P450 to form free radicals, damage macromolecules
	causing necrosis hepatic carcinogens for laboratory animals.(Carcinogenesis)
tetracycline	block the secretion of triglycerides, causing fatty liver.
chronic ethanol	block the secretion of triglycerides, causing fatty liver. damage the hepatocytes & increase the activity of fibroblasts resulting in cirrhosis and fibrosis. cuasing cirrhosis
chloroform, bromobenzene, halogenated hydrocarbons,	metabolized by cytochrome P450 to form free radicals, damage macromolecules cuasing necrosis
Troglitazone	trigger apoptosis in hepatocytes.

Drugs	features	Uses
Paracetamol (acetaminophen)	overdose (>5-6 gm/day in adults) and a glutathione depletion (in liver cirrhosis); lead to significant necrosis and hepatotoxicity	
N-acetylcysteine		used for treating paracetamol induced liver toxicity.
steroids (£ contraceptives), phenothiazines, tricyclic antidepressants.	Can cuase Cholestasis and gall stones	
Clofibrate	increase the risk for gallstone formation.	

Drugs in the liver (lec 1)



Drugs	features
anesthetic halothane	cause a condition resembling viral hepatitis.
Valproic acid	cause fulminant hepatitis in children
chloroform	hepatic carcinogens for laboratory animals.(Carcinogenesis)
Aflatoxins	potential hepatic carcinogens.
vinyl chloride	causes a rare type of liver cancer known as angiosarcoma