EVALUATION OF LIVER FUNCTION

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HEPETIC SYSTEMS

■ BIOCHEMICAL HEPATOCYTIC SYSTEM

HEPATOBILIARY SYSTEM

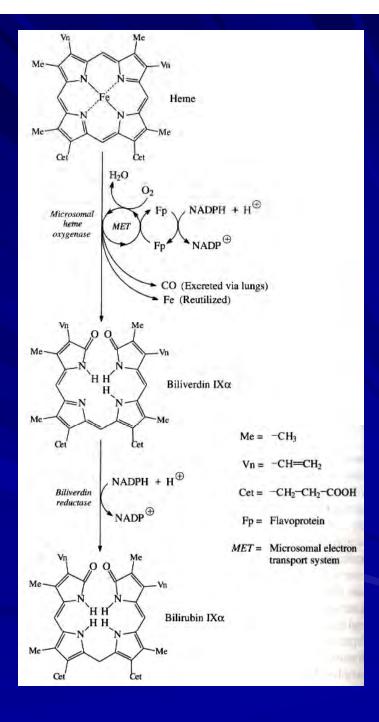
RETICULOENDOTHELIAL SYSTEM

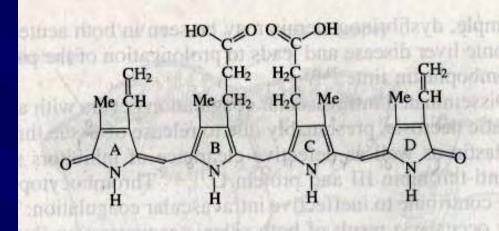
METABOLIC FUNCTION

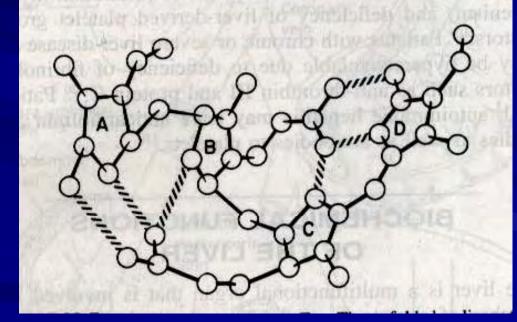
- CARBOHYDRATE METABOLISM
- LIPID METABOLISM
- AMONIA METABOLISM
- BILIRUBIN METABOLISM
- XENOBIOTIC METABOLISM

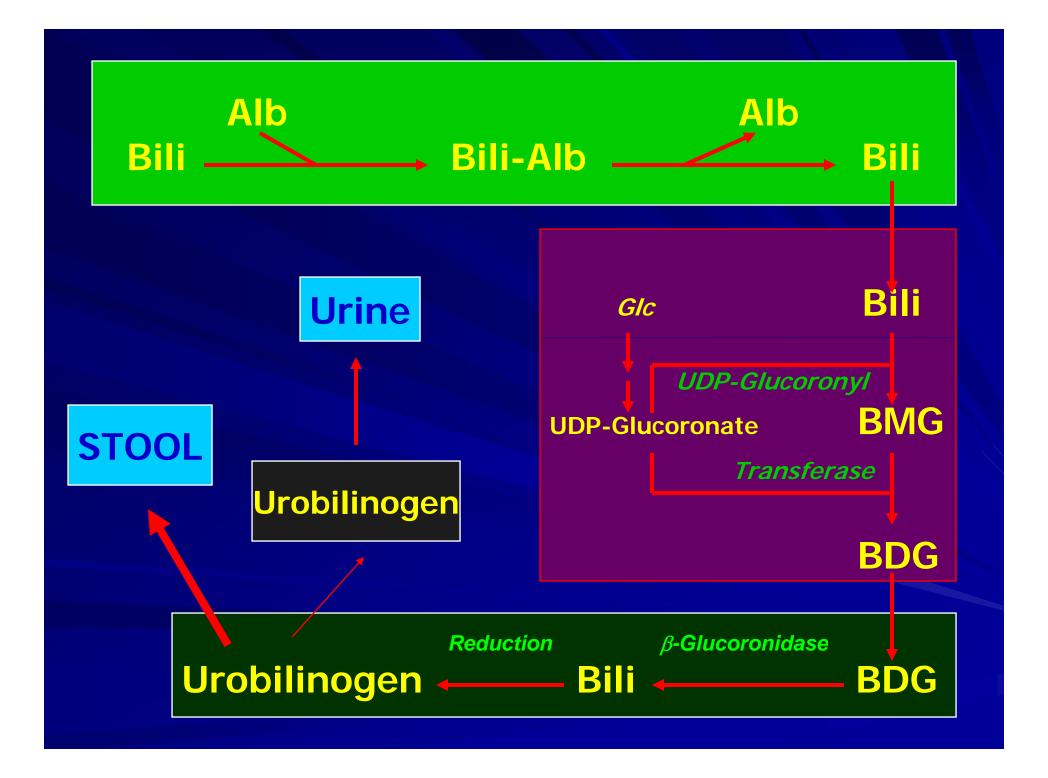
HEME CATABOLISM

- HEME
 OXYGENASE
- **BILIVERDIN**REDUCTASE









HYPERBILIRUBINEMIA

ICTERUS, JAUNDICE

■ DIRECT, CONJUGATED

■ INDIRECT, UNCONJUGATED

■ MIX

INDIRECT (UNCONJUGATED) HYPERBILIRUBINEMIA

- NEONATAL PHYSIOLOGIC JAUNDICE
- RBC ABNORMALITIES
- CRIGLER NAJJAR TYPE I SYNDROME UDP-Glucoronyl Transferase
- CRIGLER NAJJAR TYPE II SYNDROME
- GILBERT DISEASE

DIRECT (CONJUGATED) HYPERBILIRUBINEMIA

- CHOLSTASIS
- DUBIN JHONSON SYNDROME
- ROTOR SYNDROME

HEPATIC JAUNDICE

May Be:

- **DIRECT**
- **INDIRECT**
- **MIXED**

SYNTHETIC FUNCTION

- ALBUMIN
- ALPHA-1 ANTITRYPSIN
- CERULOPLASMIN
- CLOTTING FACTORS

TESTS OF LIVER INJURY

- LIVER FUNCTION TESTS
- **AUTOIMMUNE MARKERS**
- MARKERS OF VIRUS INFECTION
- **TUMOR MARKERS**

PLASMA ENZYME LEVELS

- Enzymes Primarily Reflecting Hepatocellular Injury
- Aspartate Aminotransferase (AST, GOT)
- Alanine Aminotransferase (ALT, GPT)
- Lactate Dehydrogenase (LD)
- Enzymes Primarily Reflecting Canalicular Injury
- Alkaline Phosphatase (ALP)
- Gamma-Glutamyl Transferase (GGT)
- 5'-Nucleotidase (5'-NP)
- Lucine Aminopeptidase (LAP)

Factors Affecting Enzyme Activities In Plasma

- Cellular and Subcellular Localization
- Rate of Enzyme Production
- Enzyme Gradient
- Plasmal Half-Life
- Assay Condition

ALANINE AMINOTRANSFERASE (ALT)

- Primarily Found in Liver, But Significant amounts Are Also Present in Kidney
- Hepatocyte : Plasma Ratio Is 3000 : 1
- Plasma Half-Life Is 47
- Decrease Synthesis with Pyridoxal deficiency and Fibrosis
- Is More Specific For detecting Liver Disease In Nonalcoholic, asymptomatic patients

ASPARTATE AMINOTRANSFERASE (AST)

- Distributed In ALL Body Tissues
- Has Two Isozymes

 CYTOPLASMIC

Hepatocyte: Plasma Ratio Is 7000: 1

Plasma Half-Life Is 17 h

MITOCHONDRIAL

Plasma Half-Life Is 87 h
Release By Severe Hepatocellular Damage

Is Used For Monitoring Therapy With Potentially Hepatotoxic Drugs

HEPATOCELLULAR INJURY

- In Acute Hepatocellular Injury Such as hepatitis Initially AST Increases More Than ALT After 24-48 h, ALT Increases More Than AST ALT / AST > 1.0 in Viral & Toxic Hepatitis
- In Acute Alcoholic Hepatitis

 AST Increases More Than ALT, Because of Severe Damage
 & Pyridoxal deficiency
- In Chronic Hepatocyte Injury such As Cirrhosis
 ALT Is Elevated More Than AST
 But As Fibrosiss Progresses, ALT Activity Declines
 So by The Time, AST Is Often More than ALT
- Mild & Prolonged Elevated ALT Suggest Viral Hepatitis C

ALKALINE PHOSPHATASE (ALP)

- ALP Is Found on Canalicular Surface and Is Therefore a Marker of Biliary Dysfunction
- Serum ALP Has Two Forms

Mainly in Unbound Form

To a Lesser Extent, Complexed with Lipoproteins
or Rarely Immunoglobulins

- In Cholestasis, a High-molecular-weight ALP Appears in Serum
- Normal Serum ALP, Rejects Cholestasis

Gamma-Glutamyl Transferase (GGT)

- Like ALP, Is Found on Canalicular Surface and Is Therefore a Marker of Biliary Dysfunction
- Also Is Found in Microsomes, Which Is luduced by Ethanol, Phenobarbital, Phenytoein
- So It Is a Marker of Alcoholism

Acute Injuries and / or Necrotic Lesions

- AST ——— Increased
- ALT ——— Increased
- LD ——— Increased
- Bilirubin —— Increased
- **Total Protein** → *Normal*
- Albumin → Normal
- Ammonia —— Normal

Hepatic Cirrhosis

- AST ———— Normal
- ALT ——— Normal
- LD Normal
- ALP Normal to Slightly Increased
- Bilirubin —— Increased
- Total Protein → Decreased
- Albumin —— Decreased
- Ammonia —— Increased

Acute Fulminant Hepatic Failure

- AST ———— Highly Increased
- ALT ——— Increased
- LD ——— Increased
- Bilirubin —— Increased
- Total Protein → Decreased
- Albumin ——— Decreased
- Ammonia —— Increased

Acute Biliary Obstruction

- AST ———— Normal
- ALT ———— Normal
- LD Normal
- Bilirubin —— Increased
- **Total Protein** → *Normal*
- Albumin ——— Normal
- Ammonia → Normal

Passive Congestion

AST Slightly Increased ALT **LD** ALP Normal to Slightly Increased Bilirubin ———— Normal to Slightly Increased ■ Total Protein → Normal ■ Albumin —— Normal ■ Ammonia → Normal

Space-Occuping Lesion

- AST → Normal to Increased
 ALT → Normal to Increased
- LD ——— Increased
- ALP Increased
- Bilirubin ———— Normal to Increased
- Total Protein → Normal
- Albumin —— Normal
- Ammonia → Normal

AUTOIMMUNE MARKERS

- Primary Biliary Cirrhosis (PBC)
 - Antimitochondrial Antibody (AMA)
- Primary Sclerosing Cholangitis (PSC)

Perinuclear Antineutrophil Cytoplasmic

Antibodies (p-ANCA)

Antinuclear Antibodies (ANAs)

Anti-smooth muscle Antibodies (ASAMs)

Autoimmune Hepatitis

Anti-smooth muscle Antibodies (ASAMs)

Antinuclear Antibodies (ANAs)

Antibodies to Liver-Kidney Microsomal Antigens (Anti-LKM)

TUMOR MARKERS FOR HEPATOCELLULAR CARCINOMA (HCC)

- Alpha-Fetoprotein (AFP)
 Increased in >90% Patients
 It Is Not Specific
- Ferritin

MARKERS OF HEPATITIS VIRUS INFECTION

- HBsAg
- HBsAb
- HBcAb
- HBeAg
- HBeAb
- HCV
- HAV

