

### 3) Fulminant Hepatic Failure – Dr. Muhammad Omar

FHF is defined as severe hepatic failure with encephalopathy developing in a patient with a previously normal liver. When it occurs after 4 wk but less than 6 months it is called late-onset hepatic failure (Subacute).

#### Classification of acute liver failure

Type	Time: jaundice to encephalopathy	Cerebral edema	Common causes
Hyper acute	< 7 days	Common	Viral, paracetamol
Acute	8-28 days	Common	Cryptogenic, drugs
Subacute	29 days-12 weeks	Uncommon	Cryptogenic, drugs

The causes vary throughout the world, the majority is due to viral hepatitis (60-70%), but paracetamol overdose is common in the U.K (50%). Hep. A, B, D, E, Paracetamol and Halothane usually cause FHF. While Hep. C, other drugs cause sub-acute. 25% of FHF are drug-induced.

Rare causes; Wilson disease, Autoimmune CAH, Budd-Chiari syndrome, acute fatty liver of pregnancy.

Histologically there is massive necrosis of the whole liver lobule.

#### Clinical feature

Examination shows a jaundiced patient with the signs of hepatic encephalopathy developing within 2 wks. The condition is called sub-fulminant if the encephalopathy develops 2-8 wks. The mental state varies from slight drowsiness, confusion to coma with convulsion. Fetor hepaticus is common, but signs of chronic liver disease are usually small. Fever, vomiting, hypertension, & hypoglycemia occur, spasticity & extension of the arm & leg. Cerebral edema is found in 80% of patients who die.

#### Clinical assessment

- Toxicology screen of blood and urine
- IgM anti-HBc
- IgM anti-HAV
- Anti-HEV, HCV, cytomegalovirus, herpes simplex, Epstein-Barr virus
- Ceruloplasmin, serum copper, urinary copper, slit-lamp eye examination
- Autoantibodies: ANF, ASMA, LKM
- Immunoglobulins
- Ultrasound of liver and Doppler of hepatic veins

**Complication:** 1/ Bacterial infection. 2/ hypotension. 3/ GI bleeding. 4/ Respiratory arrest. 5/ renal failure (hepato-renal syndrome). 6/ pancreatitis. 7/ Increase ICP. 8/ Hypoglycemia. 9/ Lactic acidosis.

**Investigation:** as for acute liver disease, PT is the most useful index of severity & when very high indicates a poor prognosis. AST is high initially & it falls. With progressive liver damage, the serum albumin falls.

## Treatment

- There is no specific treatment, patient should be managed in intensive care unit;
- Supportive therapy as for hepatic encephalopathy is necessary.
- Head elevation (15-20 degree). CVP monitor.
- Cerebral edema is the major cause of death & treatment with 20% mannitol 1gm/ kg i.v., Hyperventilation, Dexamethasone is no value.
- Identify & remove any precipitating factor like drug with cerebral depressant properties.
- Hypoglycemia & hypokalemia should be corrected with 10% glucose + K + Ca. regular monitoring.
- Coagulopathy treated with i.v. vit. K, platelet, FFP.
- H<sub>2</sub> antagonist (PPI) to prevent G.I bleeding.
- Infection should be treated with suitable Antibiotics. 80% develop infection (80% bacterial, 20% fungal).
- Paracetamol induced FHF should be treated with N-acetylcysteine even after 10 (but < 36) hours.
- Flumazenil; a benzodiazepine receptor antagonist may give a transient improvement of encephalopathy.
- Restriction of protein intake with adequate calories.
- Sterilization of bowel.
  - a) Mech. Sterilization: by purgation (oral & enema) using lactulose 10- 30 ml 3\*/d is an osmotic purgative, reduce colonic pH & limit ammonia absorption. Also stimulate bacterial fixation of ammonia.
  - b) Bacteriological sterilization: using broad spectrum- non-absorbable AB e.g./ Neomycin 1gm/ 6hr. Metronidazole 200mg 3\*/ d also effective.
- Antibiotics, and Anti-fungal drugs.
- Arterio venous hemofiltration for uncorrected acidosis, fluid overload, hyperkalemia, renal failure.
- ICP monitoring in stage III or IV encephalopathy. (Keep between 15-20 mm Hg). By using epidural transducers.
- Liver transplantation has been a major advance for patient with FHF. (1 year survival 80-90%).
- ? Trial of PGE1...
- Steroids; No benefit.

## Course & Prognosis

Poor prognostic parameter in Non – acetaminophen FHF includes:

- A. PT > 100 sec. or
- B. Three of the following;
  1. Etiology (HCV & drug).
  2. Age (< 10 yr. & > 40 yr.).
  3. Interval from onset of J. to encephalopathy less than 7 days
  4. S. bilirubin > 18 mg/ dl.
  5. PT > 50 sec. (INR.>3.5).or
- C. Factor V level less 15% and encephalopathy grade 3 or 4.

Poor prognosis in Acetaminophen –induced FHF:

- A. Arterial pH <7.3 or
- B. PT >100s.plus S.Creatinine >3.4mg /dl.plus, Stage III or IV encephalopathy.

The overall mortality rate of FHF without Liver transplant is about 70-80 %. Pt who survive without transplantation have excellent prognosis.