#### Variceal Bleeding from Theory to Practice



#### Nir Hilzenrat, M.D.

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### CASE STUDY

- JP.T. is a 47 year-old man who was referred to the liver clinic due to the finding of anti HCV Ab+. Recently, he felt mild fatigue.
- **PMH:** IVDU between age 20-25 years old. Has been drinking 2 glasses of wine per day for many years.
- **Physical Exam:** No jaundice, but has palmar erythema & spider angioma. Unable to palpate the spleen. No ascites.





## CASE STUDY

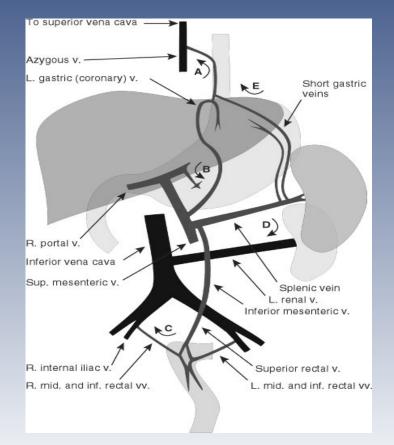
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- **Physical Exam:** No jaundice, but has palmar erythema & spider angioma. Unable to palpate the spleen. No ascites.
- Lab: 
  ALT: 65 iu/L

  T. Bilirubin: 17 mmol/L

  PLT: 130,000

  PT: 1.3
- Abdominal US: suggested liver cirrhosis, normal spleen size
- **Liver biopsy :** liver cirrhosis
- What would PUT his life at risk?
- What is the next step?

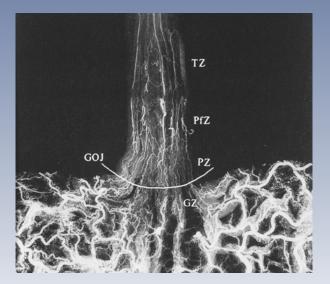
### VARICEAL BLEEDING



## DEFINITIONS

#### collateral vessels (pre-existing channels)

Portal blood is diverted to the systemic circulation bypassing the liver.



 $\triangleright$ 

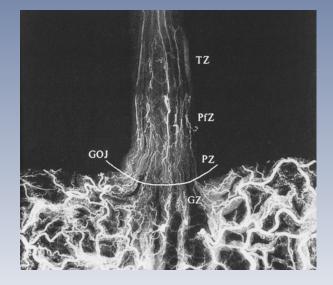
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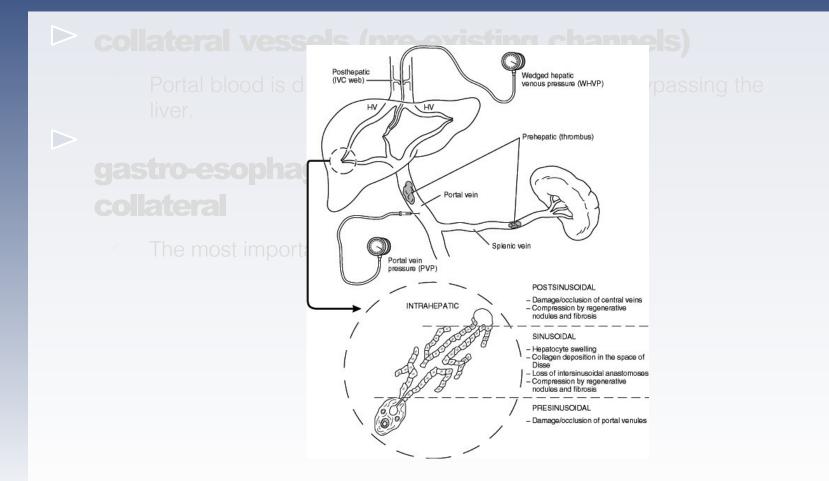
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#### gastro-esophageal collateral

The most important anastomotic system.



## DEFINITIONS



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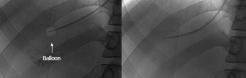
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gastro-esophageal collateral

The most important anastomotic system.

#### **Portal Hypertension in chronic liver disease**

- Normal Hepatic Venous Pressure
   Gradient (HVPG): > 5 mmHg.
- Mild portal HTN:
   HVPG > 5 mmHg, but < 10 mmH</li>
- Clinical Significant Portal HTN (CSPH):
   HVPG > 10 mmH



HVPG (18	mmHg) = Wi	HVP (22 mmHg) - FHVP (4 n	nmHg)	
	35	140		
	30	120		
14 0 0	25	100		
And my me me	20 m	M 80		
(1	15	60		
WHVP	10	40	FHVP	
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### PREVALENCE

#### Esophageal varices at presentation

- 30-40% in compensated cirrhosis.
- 60-85% in decompensated cirrhosis.

#### **30-40%**

**60-85**%

#### **Compensated Cirrhosis** Decompensated Cirrhosis

# PREVALENCE

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#### incidence of new varices

- 8-23% every year.
- Varices progress from small to large at a rate of 10-12% per year.





# PREVALENCE

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- 8-23% every year.
- Varices progress from small to large at a rate of 10-12% per year.

#### Risk of bleeding

- In 1st year after finding: 2% (no varices),
   5% (small varices), 15% (large varices).
- 6 weeks mortality : 15-25%.

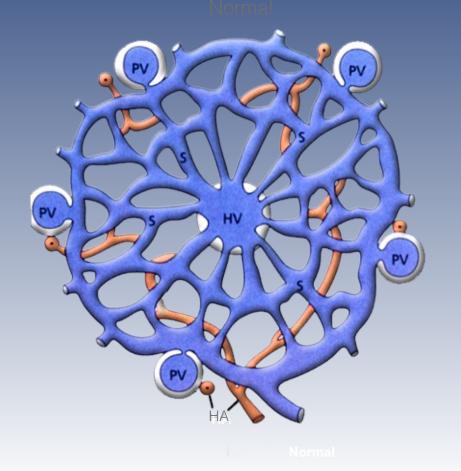


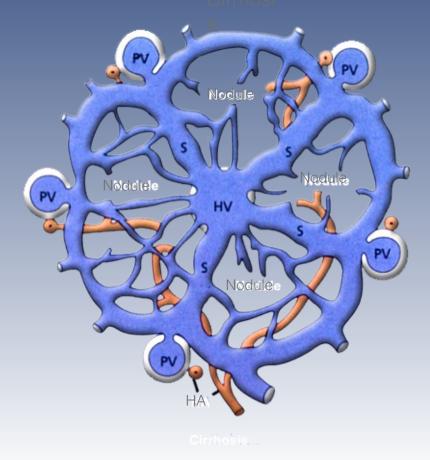


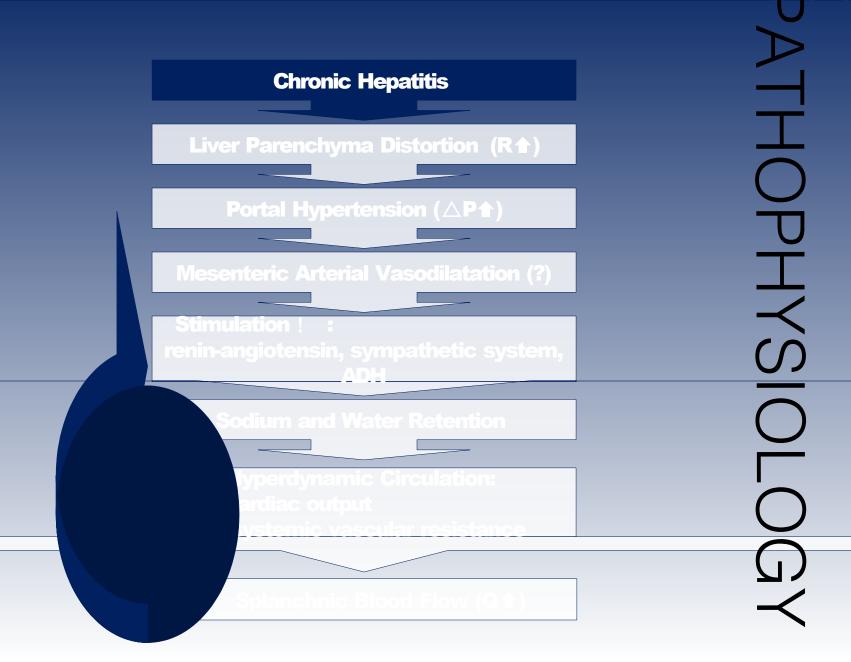
### PATHOPHYSIOLOGY

Ohm's law:  $\triangle P = Q \times R$ 

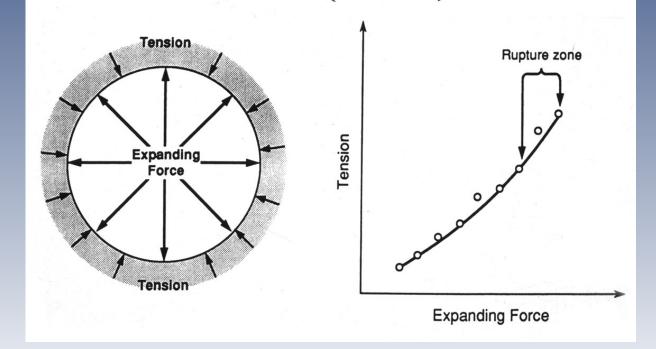
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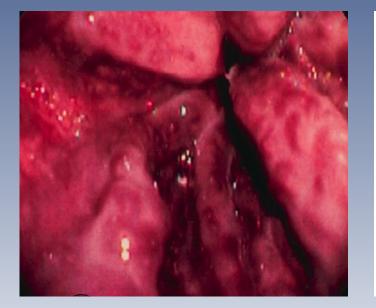


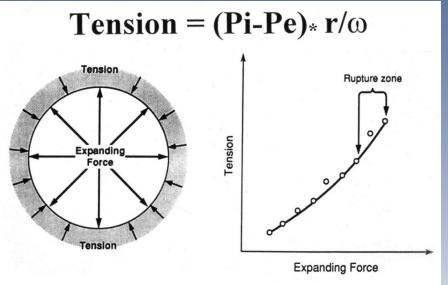


### WHY DO VARICES RUPTURE? Tension = (Pi-Pe)\* r/ω



# WHY DO VARICES RUPTURE?





### DIAGNOSIS

#### NON-INVASIVE Tests in the Diagnosis of Clinical Significant Portal HTN (CSPH)

Abdominal US: portocollateral circulation or reversal flow within the portal system.

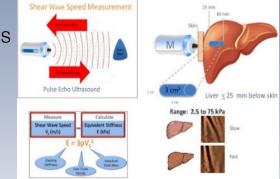




## DIAGNOSIS

#### NON-INVASIVE Tests in the Diagnosis of Clinical Significant Portal HTN (CSPH)

- Abdominal US: portocollateral circulation or reversal flow within the portal system.
- Fibroscan: Liver Stiffness > 20-25 kPa
   Robic MA et al. J Hepatol 2011;55:1017
  - Liver stiffness (kPa) X Spleen size (cm) / Platelets count (number/mm3) Score **(LSPS**) > 2.06 *Berzigotti A et al. Gastroenrerology 2013;144102*
- Spleen Stiffness > 55 kPa needs
   sonoelastographic methods.



## CASE STUDY -

- JP.T is a 52 year-old man with liver cirrhosis due to PCV and alcoho. UED
  - Child–Pugh score: 6 (A)
  - Abdominal US: spleen size 13 cm
  - Fibroscan: 17 kPa
  - LSPS: 1.8
- \* What would be your next step?
- Patient refused Tx with PEG Inf. / RBV and continued to drink alcohol.
  - Repeat abdominal US (one year later): mild ascites & splenomegaly of 14.5cm
- What should you do?

## PREVENTION

#### of first variceal hemorrhage (VH) in patients with medium/large esophageal varices

#### NON SELECTIVE BETA BLOCKERS (NSBB)

NSBBs (propranolol, nadolol) have benefit in preventing first VH. D'Amico G. et al. Semin Liver Dis 1999:19:475

In case of intolerance, try switching to Carvedilol:

- Propranolol: 20-40mg bid; Max: 320mg/d; Goals?: HR 55-60 beats/min, BP< 90 mmHg</li>
- Nadolol: 20-40mg qd; Max: 160mg/d; Goals?: as above
- Carvedilol: 6.25mg qd; Max:12.5mg/d; Goals?: BP< 90mmHg

Garcia-Tsao G et al. Hepatology2017;65:310-335



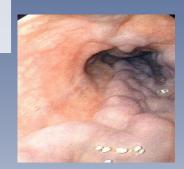
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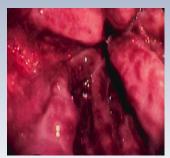
#### NON SELECTIVE BETA BLOCKERS (NSBBs)

#### **Esophageal Varices Ligation (EVL)**

- Esophageal Varices Ligation (EVL) may associate with lower rates of VH Compare to NSBBs but, no change in mortality.
   Gluud LL et al. Cochrane Database Syst rev 2012;(8): CD005-44
- Combination of NSBBs & EVL is not recommended due to high SE?
   Sarin SK et al. Am J Gastroenterol 2005;100:797







### PREVENTION

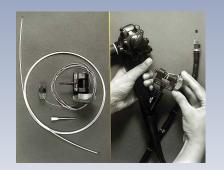
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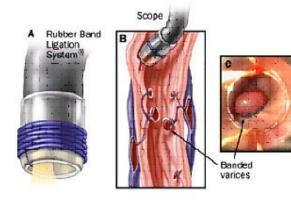
#### NON SELECTIVE BETA BLOCKERS

#### (NSBBs)

#### Esophageal Varices Ligation (EVI)









## CASE STUDY -

- JP.T is a 53 year-old man with liver cirrhosis due to HCV and alcoho. UED
- Patient refused Tx with PEG Inf. / RBV and continued to drink alcohol.
  - Repeat abdominal US (one year later): mild ascites & splenomegaly of 14.5cm.
  - Took NSAIDs for low back pain
  - Presented to ER with massive hematemesis.
- \* What are the possible causes of the bleeding?
- What should you do?

### MANAGEMENT Acute Esophageal Variceal Bleeding

#### Immediate goals

- Control bleeding.
- Prevent early recurrence in 5 days.
- Prevent 6-weeks mortality.



### MANAGEMENT Acute Esophageal Variceal Bleeding

#### Immediate goals

#### VOLUME RESTITUTION

- "Restrictive" packed red blood cell (PRBC) transfusion strategy: start transfusion when Hb < 70 gr/L and maintain Hb 70-90 gr/L...</li>
   *Villanueva C et al. N Engl J Med 2013;368:11-21*
- There is no evidence that correction of coagulopathy has any benefit.



# MANAGEMENT

Acute Esophageal Variceal Bleeding

#### Immediate goals

VOLUME RESTITUTION

#### Antibiotics

High risk to develop bacterial infections → antibiotics
 prophylaxis [IV ceftriaxone 1 gr/d for 5-7 days]

### **MANAGEMENT** Acute Esophageal Variceal Bleeding

#### Immediate goals

- VOLUME RESTITUTION
- Antibiotics

#### Vasoactive agents

- Lower 7-days mortality.
   Wells M et al. Aliment Pharmacol Ther 2012;35:1267-1278)
- Start as early as possible before gastros

Name	
Address	Date
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<b>-X</b>	
- Octreo	tide: IV bolus
50 mcg	Ir then
continu	
CONTINUE	ous IV
50mcgr	/h for 2-5
days.	
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Terlinrog	and the

 Terlipressin: IV 2mg every 4 h until control bleeding, then, IV 1
 MD mg every 4 h for 2-5 Signature days.

### MANAGEMENT Acute Esophageal Variceal Bleeding

#### Immediate goals

- > VOLUME RESTITUTION
- Antibiotics
- VASOACTIVE AGENTS

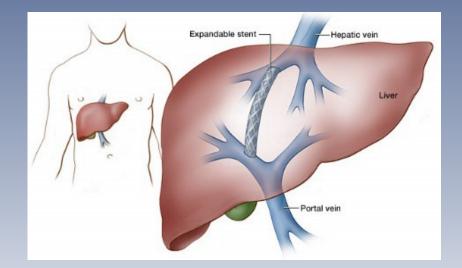
#### Endoscopy

- Within 12 hours after presentation.
- EVL for esophageal bleeding- active bleeding or sign of recent bleeding.



### MANAGEMENT Acute Esophageal Variceal Bleeding

- Immediate goals
- VOLUME RESTITUTION
- Antibiotics
- VASOACTIVE AGENTS
- Endoscopy



#### TIPS

For patients with early rebleeidng who failed vasoactive agents and/or EVL or high-risk patients (Child-Pugh B/C & active bleeding during endoscopy

### AFTER RECOVERY Patients who recovered from an episode

of acute esophageal varices bleeding



- Risk of Rebleeding: 60% in the first year.
- Mortality: up to 33%.

# AFTER RECOVERY

Patients who recovered from an episode of acute esophageal varices bleeding

#### **RISK**

#### NSBB + EVL (for eradication)

- Combination of **NSBB + EVL** is first line therapy.
- Propranolol 20-40 mg bid;
   Max: 320mg/d, with ascites- 160mg/d.
- Nadolol 20-40mg qd;
   Max: 160mg/d, *with ascites*-80mg/d.





# AFTER RECOVERY

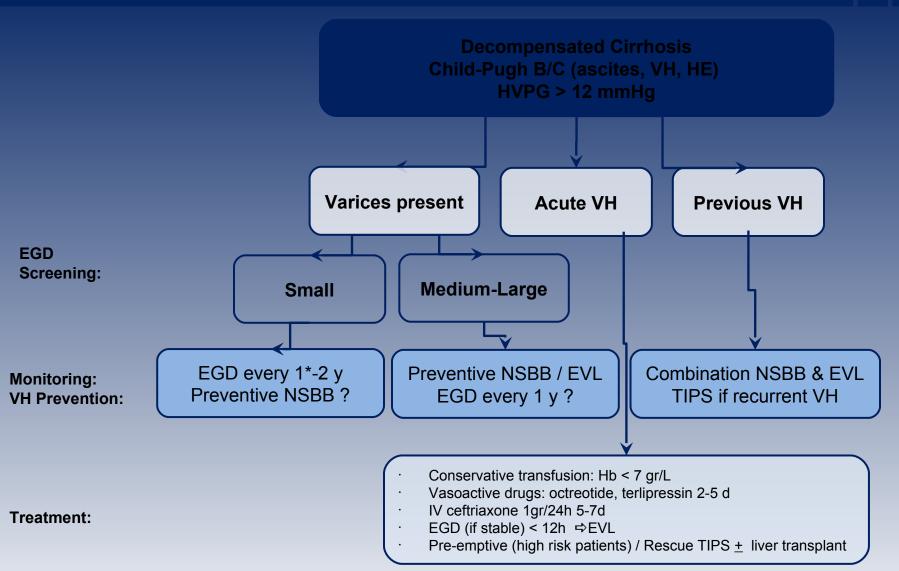
Patients who recovered from an episode of acute esophageal varices bleeding

#### **RISK**

NSBB + EVL (for eradication)

#### ► TIPS

- Recommended rescue therapy during acute phase and recurrent bleeding despite combination therapy.
- Consider referral for liver transplant.

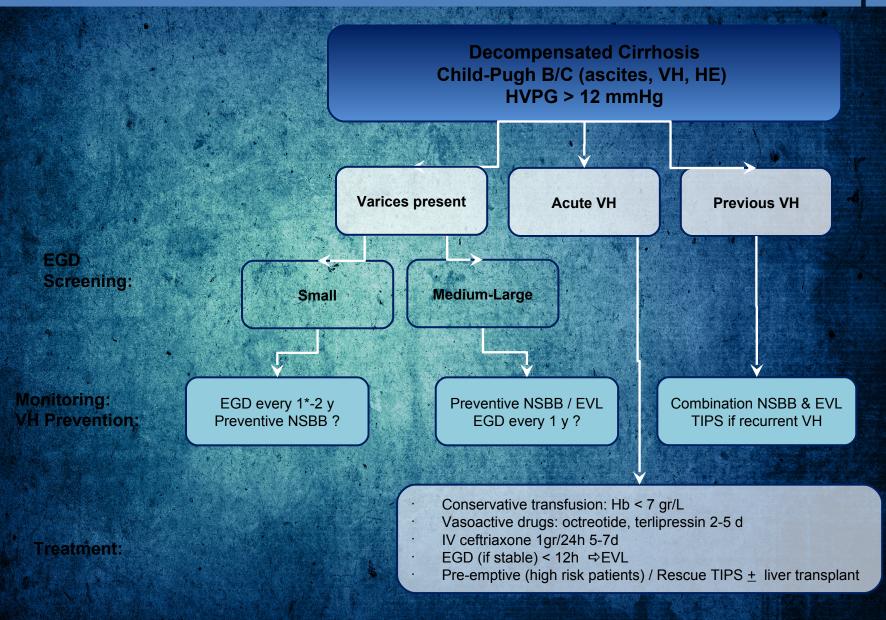


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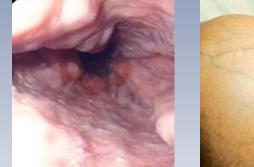


Active disease

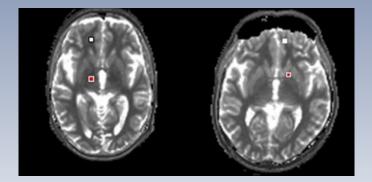
## DEFINITIONS

#### Decompensated cirrhosis

- Clinical evident of decompensated events; ascites, variceal hemorrhage and hepatic encephalopathy.
- Child-Pugh score B/C & median survival 1.8 years.







Esophageal Varices

**Ascites** 

Encephalopat hy