

Variceal Upper GI bleed :
 more than meets the "endoscopic eye" ...

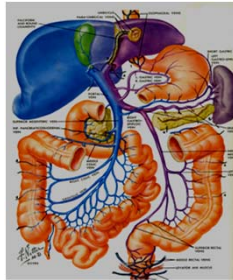
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 University Hospitals Leuven

BSGIE, 9-2016

1. The conundrum "Variceal hemorrhage"
2. Treating variceal hemorrhage : an integrated algorithm
3. Rescue therapies in acute situations
4. Conclusions

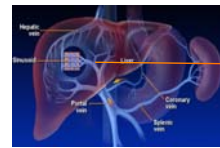
Variceal hemorrhage & portal hypertension:
 inevitably linked

"... a syndrome that naturally develops in diseases and conditions that impede the natural drainage of the portal vein..."

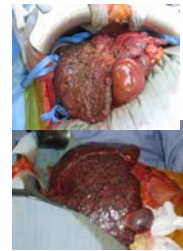


Herrick J Exp Med 1907

Cirrhotic portal hypertension



Hepatic

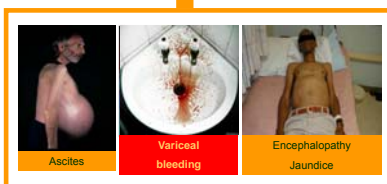


Cirrhotic portal hypertension = responsible for 90% of all causes of PHT

Natural history of chronic liver disease
 Variceal bleeding = a decompensating event and changes the setting !

Chronic liver disease → Compensated cirrhosis → Decompensated cirrhosis → Death

Development of complications

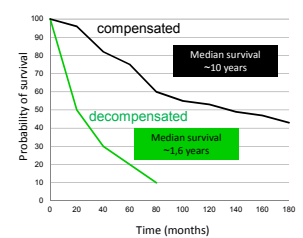
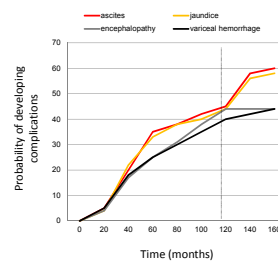


Ascites

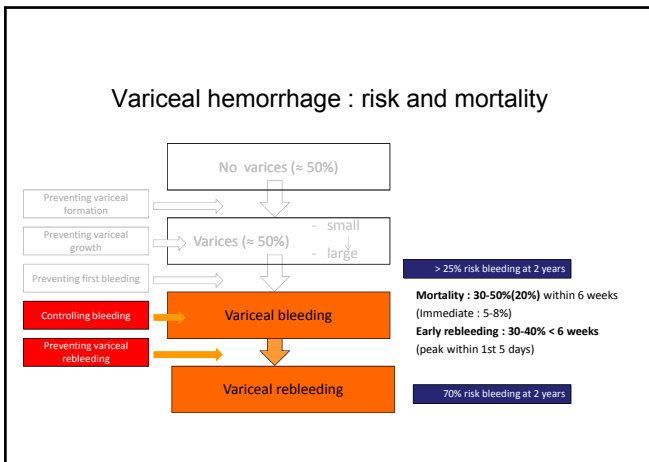
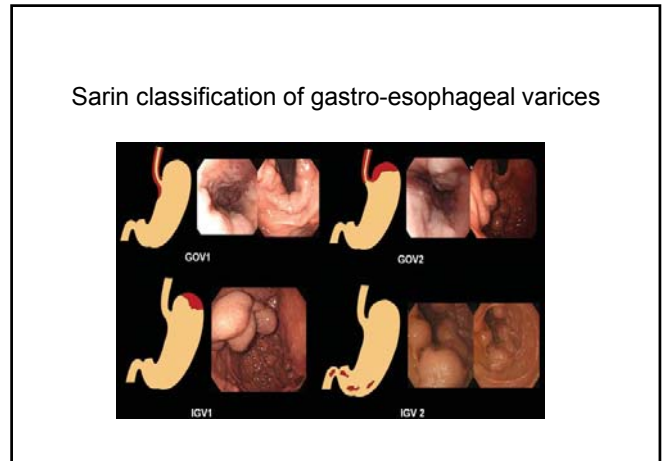
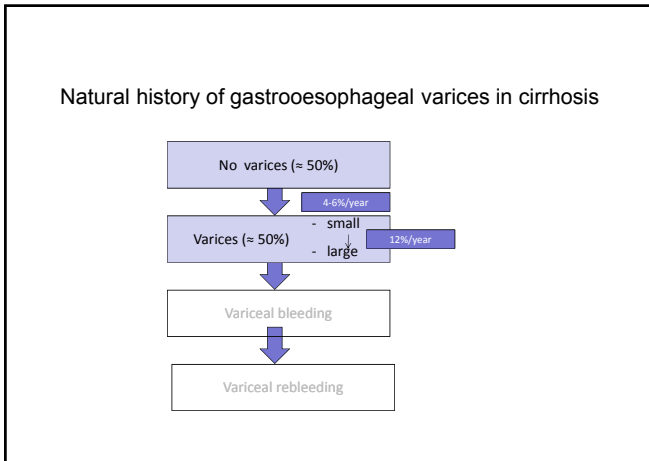
Variceal bleeding

Encephalopathy
Jaundice

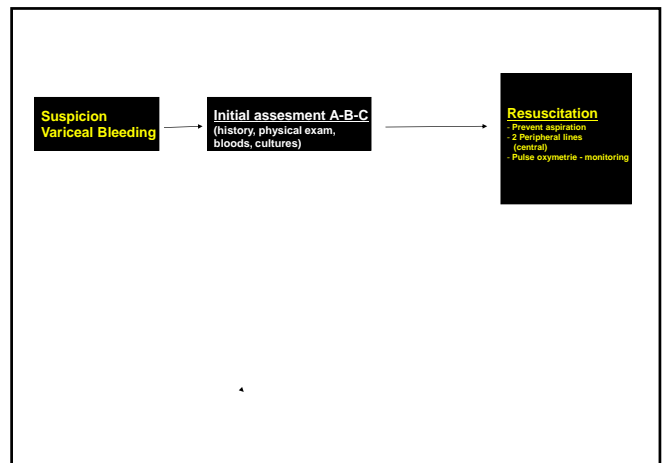
Natural history of chronic liver disease
 Decompensation warrants intensified FU & consideration for liver Tx



Gines et al. Hepatology 1987



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Suspicion Variceal Bleeding → **Initial assesment** (history, physical exam, bloods, cultures) → **Resuscitation**

Resuscitation
Prevent aspiration
- Peripheral lines (central)
- Pulse oxymetrie
- PC when Hb < 7g%

214 cirrhotics with AVB

Restrictive transfusion policy (Hb: 7 g/dl) N=109
Liberal transfusion policy (Hb: 9 g/dl) N=105

no recommendations for coagulation factors
platelet transfusion threshold ≤ 40,000

Colomo A et al. AASLD 2009 #232
Casasnovas et al. Hepatology 2001
Villanueva C et al NEJM 2013
BAVENO VI 2015

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Resuscitation
Prevent aspiration
- Peripheral lines (central)
- Pulse oxymetrie
- PC when < 7g%
- antibiotics (FQ)

Infection (20-50%) = frequent trigger of bleeding, failure to control bleeding and rebleeding

- 1st line : FQ 5-7 days
- 2nd line : 3rd generation cephalosporine
 - when on FQ prophylaxis
 - "advanced cirrhosis"

Thalheimer U et al. Gut 2004
Bernard B et al. Hepatology 1999
Fernandez J et al. Gastroenterology 2006
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Suspicion Variceal Bleeding → **Initial assesment** (history, physical exam, bloods, cultures) → **Resuscitation**

Resuscitation
Prevent aspiration
- Peripheral lines (central)
- Pulse oxymetrie
- PC till Hb 7g%
- antibiotics (FQ)
- vasoactive drugs ASAP
- Somatostatin
- Terlipressin

NEED TO BE STARTED BEFORE ENDOSCOPY !!!!

Somatostatin
250mcg bolus - 3mg/12hrs

Terlipressin
2mg loading dose - 1-2mg/4hrs

Duration : 2-5 days

Levacher S. et al. Lancet 1995
Burroughs et al. Gastroenterology 1990

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- HE prophylaxis

Control of bleeding in patients with active bleeding at endoscopy

Other outcomes	Low (250)	High (500)	P
Early rebleeding (%)	21	13	ns
Transfusions (UJ)	2.8 ± 2.3	1.7 ± 1.8	0.07
One-week mortality (%)	20	0	0.02
Six-week mortality (%)	32	8	0.02

Double dose of somatostatin (500mcg instead of 250mcg) upon ACTIVE bleeding at ENDOSCOPY

Motinho et al. J Hepatol 2001
Wells M et al. APT2012

Suspicion Variceal Bleeding → **Initial assesment** (history, physical exam, bloods, cultures) → **Resuscitation**

Resuscitation
Prevent aspiration
- Peripheral lines (central)
- Pulse oxymetrie
- PC < Hb 7g%
- antibiotics (FQ)
- vasoactive drugs
- Somatostatin
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Endoscopy

Failure to control bleeding (7 trials, 710 patients)
Early (5 day) mortality (8 trials, 1025 patients)
Late (< 42 day) mortality (6 trials, 1018 patients)

Drugs + endoscopic R/better | Drugs alone better

Banares R et al.

Suspicion Variceal Bleeding → **Initial assesment** (history, physical exam, bloods, cultures) → **Resuscitation**

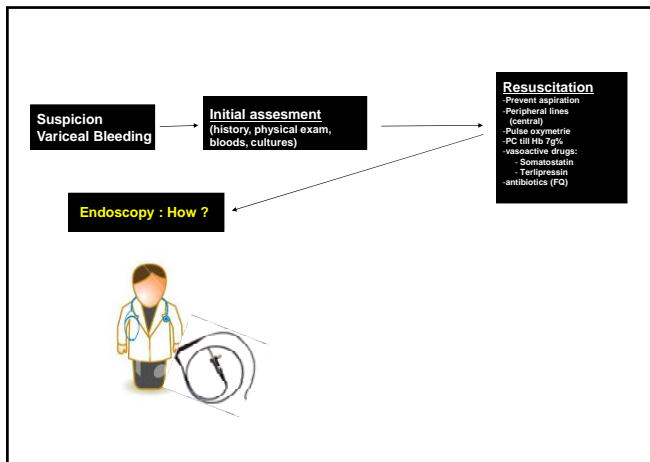
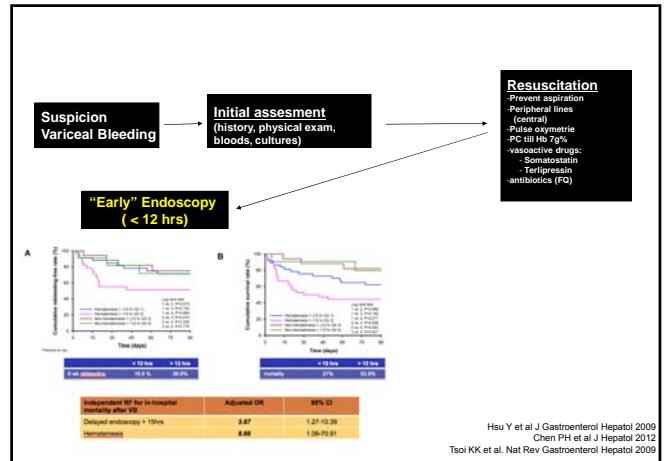
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Prevent aspiration
- Peripheral lines (central)
- Pulse oxymetrie
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- Somatostatin
- Terlipressin
- antibiotics (FQ)

Endoscopy : when ?

Tsai KK et al. Nat Rev Gastroenterol Hepatol 2009

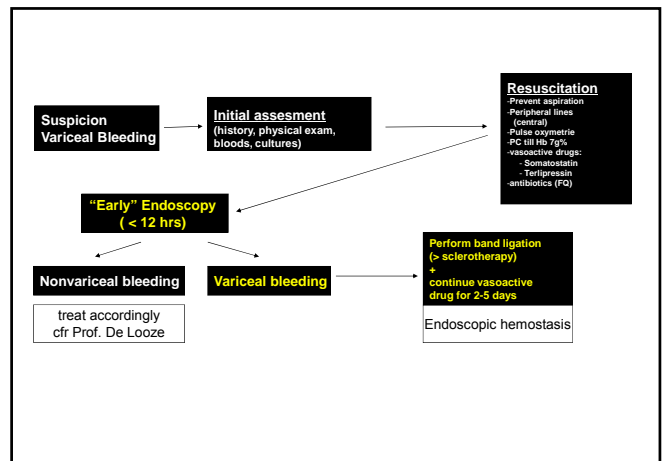
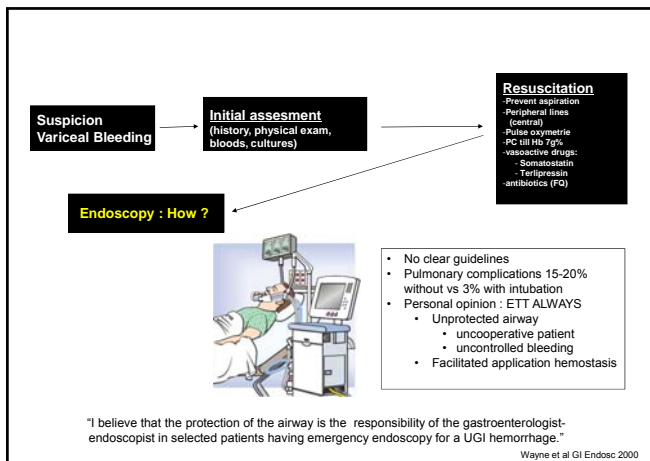
Q1 : Who scopes a patient stabilized and admitted with hematemesis and suspicion of variceal hemorrhage ...

1. within 3 hours after admission
2. at the latest always within 12 hours after admission
3. within 12 hours after admission, but only during the week
4. the next coming working day



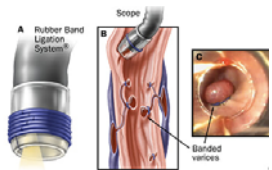
Q2 : Who has patients intubated for upper GI endoscopy and suspicion of variceal hemorrhage ?

1. Never
2. Always
3. Sometimes (in > 50% of cases)
4. Sometimes (in < 50% of cases)



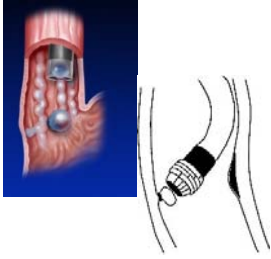
Endoscopic band ligation (EBL)

- Technique of choice for esophageal varices
- rubber band ligatures (4-6-10-shooter)
- Determine landmarks (Z-line, classification, grade, location, ...)



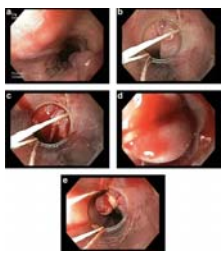
Endoscopic band ligation (EBL)

- Technique of choice for esophageal varices
- rubber band ligatures (4-6-10-shooter)
- Determine landmarks (Z-line, classification, grade, location, ...)
- Start at Z-line – spiral step up
- As vertical as possible on the varix and suck in the maximal possible bleb of varix




Endoscopic band ligation (EBL)

- Technique of choice for esophageal varices
- rubber band ligatures (4-6-10-shooter)
- Determine landmarks (Z-line, classification, grade, location, ...)
- Start at Z-line
- As vertical as possible on the varix and suck in the maximal possible bleb of varix
- DO NOT RELEASE WHEN SPURTING BLEEDING arises (bursting red spot)
- "Varix drop-off" after 3 days

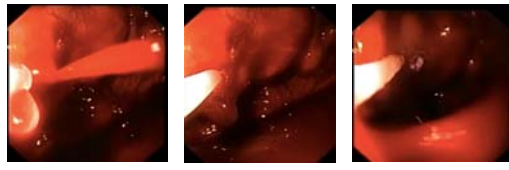


Tissue adhesives "Glue" - N-butyl-2-cyanoacrylate (Histoacryl)



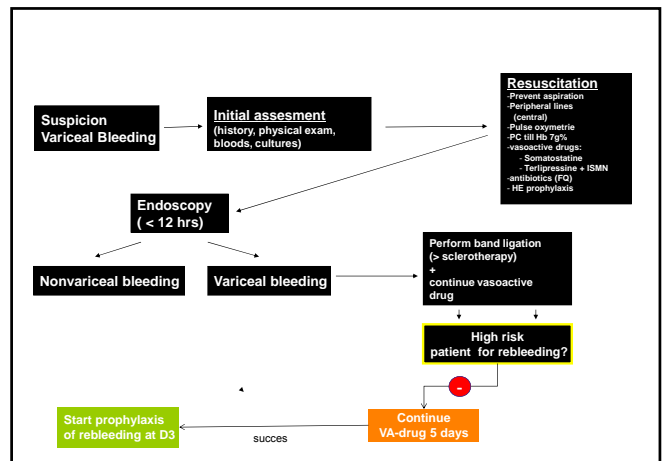
0.5ml lipiodol
 0.7ml lipiodol
 0.5ml lipiodol + 0.5ml glue
 0.7ml lipiodol + 0.3ml glue

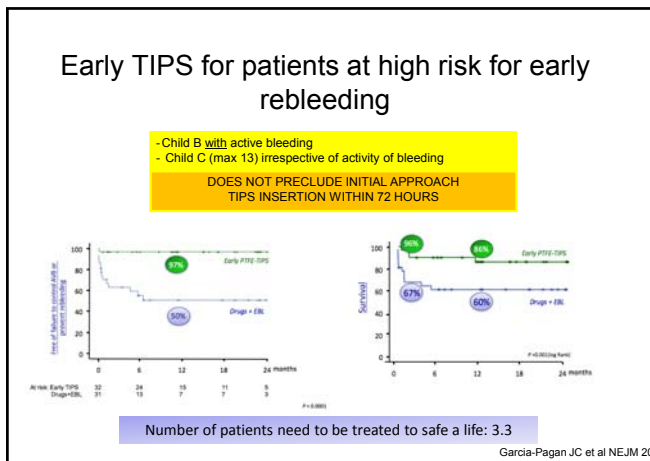
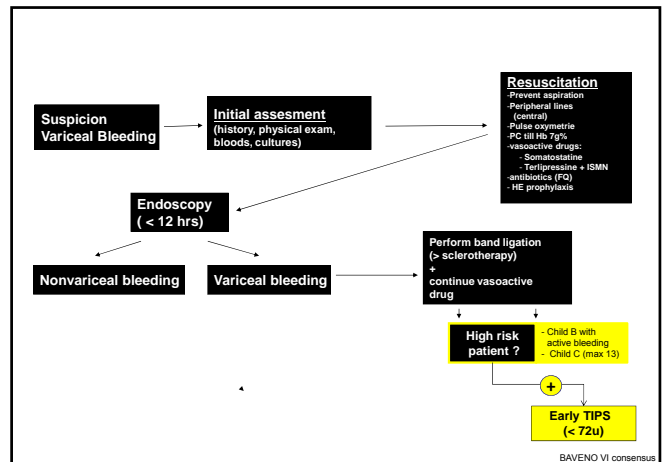
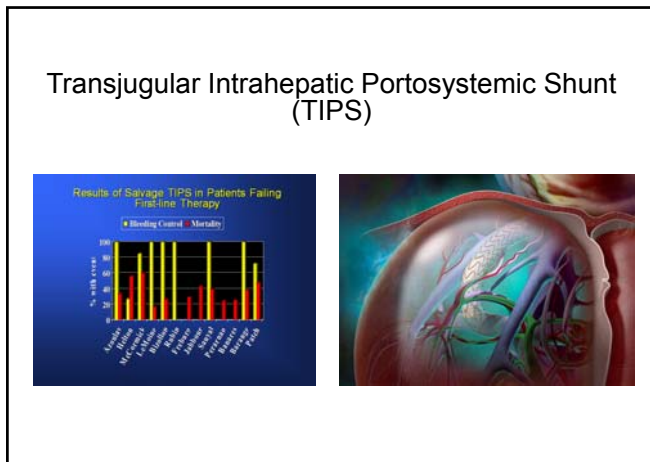
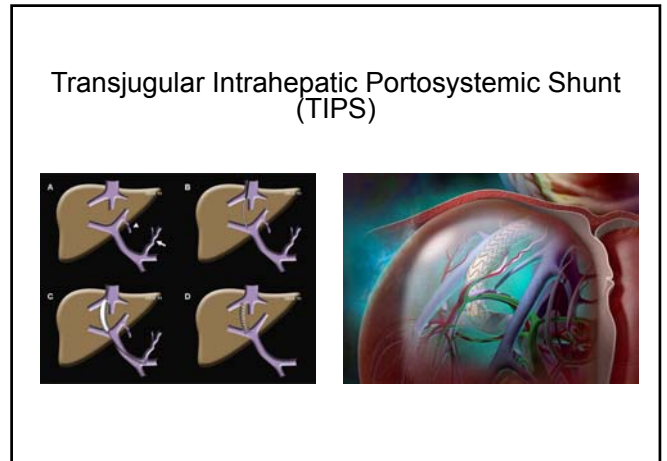
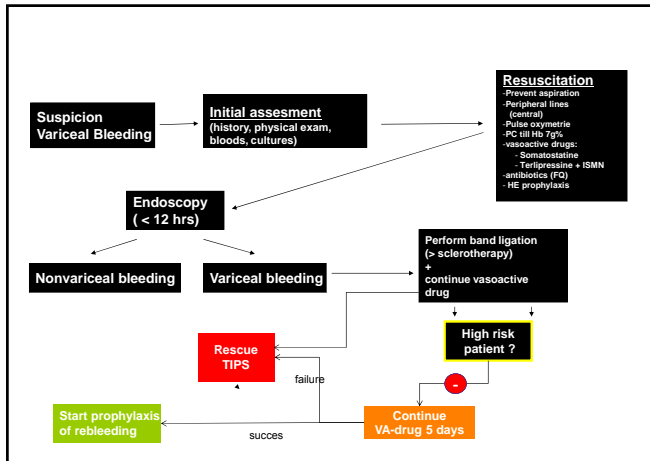
Tissue adhesives "Glue" - N-butyl-2-cyanoacrylate (Histoacryl)



- flush needle with 2cc glucose 5%
- Load syringe with glue
- choose puncture site
- Puncture varix
- Inject 1cc glue
- Immediately exchange for syringe with glu 5%
- Inject 2cc at least to clear the needle from remnant glue
- pull back out of varix at 1.5-2cc to have a "seal"
- Don't pull back needle-catheter in the scope

Repeat steps and inject until polymerization of the varix





- ### Q3 : Who applies the principle of early TIPS ?
1. Never
 2. Always
 3. Sometimes in > 50% of cases
 4. Sometimes in < 50% of cases


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Sengstaken-Blake more & Ella-stent
 Emergency tamponade for bridging



The image shows a Sengstaken-Blake tube, a long red tube with a yellow balloon at the top, and an Ella-stent, a self-expanding metal stent. It also includes a diagram of the tube's placement in the esophagus and stomach, and endoscopic views of the stent in place.

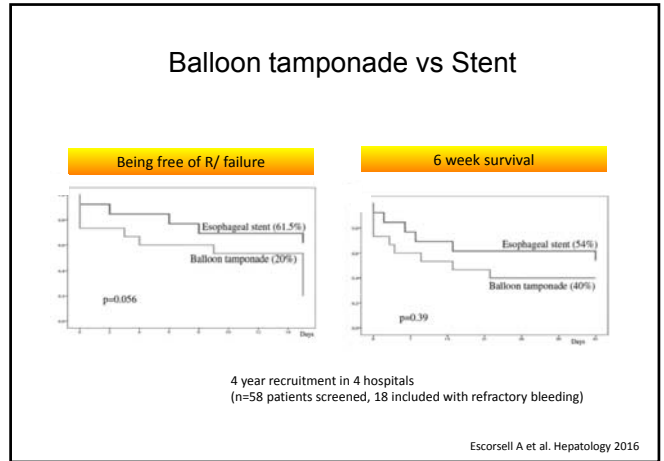
Self-expandable metal stent
 (SX-ELLA Danis stent, 12x35mm)



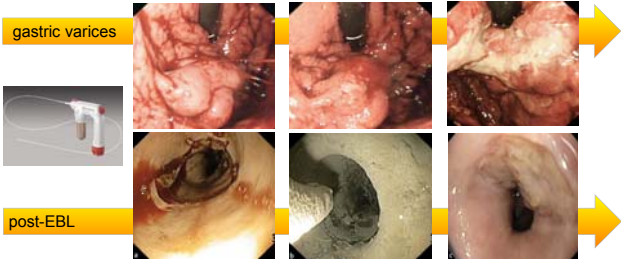
Year	n patients	ES-EBL successfully deployed (%)	Success rate (n/total) (%)	Rebleeding (%)	ES-EBL mortality (%)	Median survival time (n/total) (days)	Median follow-up period
Johnson et al. 2008	10	100	100	0	10	11.0 (10)	200-300 days
Johnson et al. 2008	14	100	100	0	14	17.0 (14)	200-300 days
Wright et al. 2010	10	90	70	10	100	1.8 (10)	300-42 days
Johnson et al. 2012	8	100	100	0	0	11.0 (7)	100-200 days
Johnson et al. 2013	7	100	100	0	0	11.0 (7)	400-1000 days
Johnson et al. 2013	10	100	100	0	0	100.0 (10)	200-300 days

ES, self-expandable esophageal stent; EBL, endoscopic balloon ligation.

El Sayed, O'Beirne J, et al. Frontline Gastroenterol 2015



Rescue therapy in refractory variceal bleeding or post-EBL ulcer bleeding : hemostatic powder (Hemospray)



The image shows the Hemospray device and endoscopic views of gastric varices and a post-EBL ulcer. Arrows indicate the application of the powder to the bleeding sites.

Stanley AJ et al Endoscopy 2013
 Holster et al. J Hepatol 2012.
 Ibrahim M et al. Endoscopy 2014

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Conclusions:

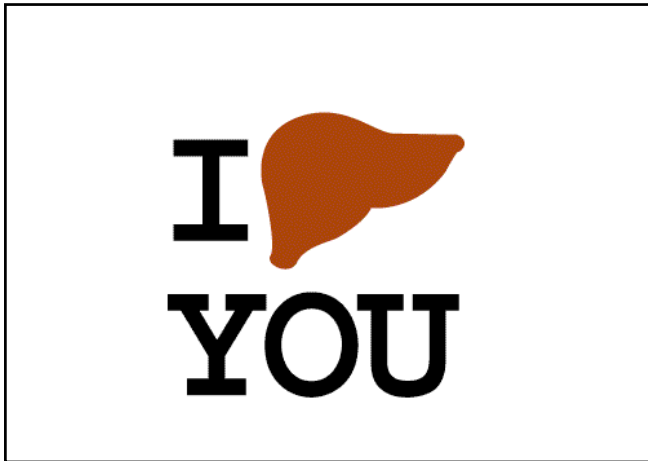
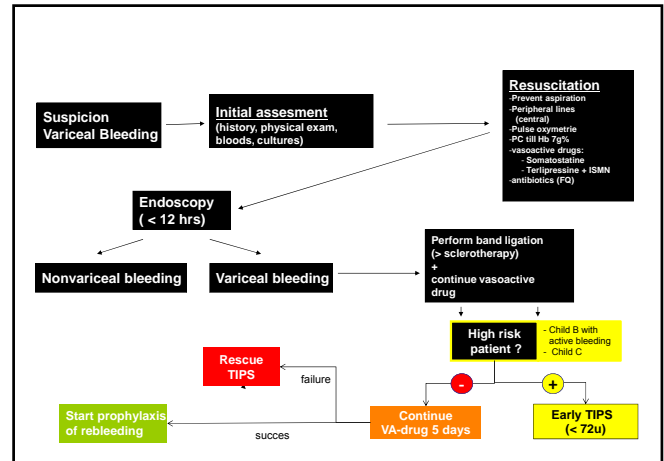
Variceal hemorrhage is a **serious bleeding complication in the UGI tract** and represents at the same time a **decisive transition in the natural history** of a patient with cirrhosis

Focus on **controlling bleeding** but also on **preventing rebleeding**

The management of VH is best served with **an algorithm** that is rigorously applied at each situation suspect for VH

This algorithm involves the **combined implementation of medical, endoscopic and potentially also interventional radiological measures**

Novel auxiliary bridging tools, such as hemostatic powder and selfexpandable stents, might prove beneficial



How to define failure ?

Failure to control bleeding

- within 6 hours after time zero
- transfusion ≥ 4 U PC
- impossibility to raise BP with 20mmHg or ≥ 70 mmHg

Bpsyst

- after 6 hours from time zero
- hematemesis
- drop in Bpsyst ≥ 20 mmHg
- transfusion of ≥ 2 U PC more to achieve Hb $\geq 7g\%$

Rebleeding

- new episode of melena or hematemesis after a period of 24 hours or more of stabilization
- considered **significant** if ≥ 2 U PC needed to attain Hb 7g%
- **early rebleeding**: if less than 1week after index bleeding

BAVENO V consensus

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BAVENO V consensus

Failure implies change of strategy !!!!!