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All communications to:

EMH Swiss Medical Publishers Ltd.
 Swiss Medical Weekly
 Farnsburgerstrasse 8
 CH-4132 Muttenz, Switzerland
 Phone +41 61 467 85 55
 Fax +41 61 467 85 56
 office@smw.ch

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Percutaneous trans-hepatic cholangiography and drainage (PTCD) followed by a single balloon enteroscopy (SBE) assisted ERCP to treat a choledocholithiasis with cholangitis in a patient after Roux-en-Y surgery

Patrick Aepli¹, Martin Hoffmann²

¹ Division of Gastroenterology and Hepatology, Department of Medicine, Luzerner Kantonsspital, Lucerne

² Department of Radiology, Luzerner Kantonsspital, Lucerne

Background: Endoscopic access to the biliary system can be technically challenging in patients with surgically altered anatomy, such as a Roux-en-Y reconstruction. By using single (SBE) or double balloon enteroscopy (DBE) it is possible to perform an ERCP even in these patients thereby giving the potential to perform diagnostic or therapeutic intervention.

Methods: We present the case of an 80-year-old patient with cholangitis due to choledocholithiasis in a postoperative setting after Roux-en-Y surgery.

Results: SBE was used to gain access to the papilla under CO₂ insufflation but unfortunately a selective cannulation of the common bile duct (CBD) was not technically possible. As a result we performed a PTCD and pushed the stone into the duodenum. Recurrent cholangitis, inspite of PTCD, meant that even biliary stenting was necessary and 6 weeks later the patient presented for endoscopic stent removal. This proved difficult because the stent had migrated upwards into the CBD and couldn't be seen anymore endoscopically, but nevertheless stent removal was successful. In addition to the stent removal another stone was extracted from the CBD. The patient had a good clinical course without any adverse events.

Conclusion: SBE- or DBE-ERCP is a safe and often successful procedure for treating biliary obstruction in patients post Roux-en-Y reconstruction. However it is sometimes necessary to first complete a PTCD and the balloon assisted ERCP still has several disadvantages (absence of side-viewing perspective; lack of an elevator; need of extra-long ERCP accessories).

Evaluation of the Controlled Attenuation Parameter (CAP™) as a Non-Invasive Tool to Diagnose Liver Steatosis

02

Stephan Baumeler¹, Christine Alfter¹, Irina Bergamin¹, Jörg Neuweiler²,
Wolfram Jochum², David Semela¹

¹ Klinik für Gastroenterologie & Hepatologie und ²Institut für Pathologie, Kantonsspital St. Gallen, Rorschacherstrasse 95, CH-9007 St. Gallen

Background: Liver steatosis is a mostly benign and reversible condition. However, under certain conditions steatosis is associated with inflammation and liver fibrosis. Although liver biopsy with histology is the gold standard for assessment of hepatic steatosis, this technique is not readily available. Recognizing the increasing prevalence of steatosis (i.e. non-alcoholic fatty liver disease (NAFLD)) and the lack of specific laboratory findings, there is a need for non-invasive steatosis testing. In analogy to non-invasive fibrosis staging by vibration-controlled transient elastography (i.e. Fibroscan™) a novel ultrasonic controlled attenuation parameter (CAP™) has been developed to detect and quantify steatosis. Our aim is to validate the accuracy of non-invasive CAP™ measurements in comparison to histologic steatosis grading.

Methods: Interim analysis of a single-center, non-interventional study. Consecutive patients scheduled for liver biopsy for any reason (without focal liver lesion) underwent sonography, transient elastography (Fibroscan™) and CAP™ measurement immediately before biopsy to evaluate the diagnostic performance of CAP™ using the Fibroscan 502 Touch™ (3.5 MHz M probe, 10 measurements). Histologic grading of steatosis (S0 <10% steatosis, S1 11-33% steatosis, S2 34-66% steatosis, S3 >66% steatosis) was performed by an experienced pathologist (WJ) in a blinded manner. Spearman's rank correlation coefficient was used to measure the strength of association.

Results: 41 patients (16 female, 25 male) mean age 49 years were assessed by CAP™ and liver biopsy (10 with fatty liver disease (4 NAFLD, 3 NASH, 3 ASH), 18 with HCV, 5 HBV, 7 other). Mean CAP was 259 dB/m (range 173-394 dB/m, median 249 dB/m); mean success-rate 93%, accurate measurement (IQR/med <30%) in 38/41 (93%). The R value of the Spearman test was 0.58701 (two-tailed value of p = 0.00005).

Conclusions: This interim analysis of non-invasive steatosis assessment by CAP™ measurement showed significant correlation with histological grading of steatosis. However, the clinical value of CAP™ remains to be determined.

Toll-interacting protein Deficiency Protects Mice from Colitis-associated Cancer

Hristina Bega¹; Yan Pu², Dominique Velin¹ and Michel H. Maillard¹.

¹Service of Gastroenterology and Hepatology, CHUV, Lausanne;

²Institut Universitaire de Pathologie, CHUV, Lausanne.

Background: Genetic deletion of the Toll-interacting protein (Tollip) -an IL-1R and TLR2/4 regulator- leads to increased acute and chronic colitis in mice. We sought to investigate whether increased susceptibility to inflammation could also favor inflammation-driven colorectal carcinogenesis.

Methods: Colitis-associated cancer (CAC) was induced in 18-20-week old littermates C57BL/6 mice by azoxymethane (AOM) i.p. injection and 3 cycles of 2.5% oral dextran sodium sulfate (DSS) treatment. Tumor development was assessed by colonoscopy. Apoptotic and proliferative index in the colon were determined by Tunel assay and Ki67 immunohistochemistry and quantified using the Image J software.

Results: Tollip KO mice had significantly lower endoscopic tumor scores than WT littermates upon AOM-DSS exposure (8.38±7.78 vs 13.38±6.37, p>0.05). Likewise, tumor numbers (4.85±3.51 vs 7.06±2.99, p>0.05) and size (12.84% ±13.64 vs 37.34% ±20.53, p>0.05) were reduced. Immunohistological studies demonstrated reduced apoptotic index (79.29±75.07 vs 246.8±152.9, p>0.05) and a trend towards lower proliferation (20.96±8.53 vs 27.91±7.26, ns) in Tollip KO tumors when compared to wt controls.

Conclusions: Our data show that Tollip partially favors colonic oncogenesis despite being protective against colitis.

Maternal Microbiota-Derived Metabolites Shaping the Neonatal Immune System Are Transferred to the Offspring Ante- and Postnatally

04

Stephanie C Ganal-Vonarburg (1,3); Mercedes Gomez de Agüero (1,3); Anna Steinert (1); Tobias Fuhrer (2); Uwe Sauer (2); Kathy D McCoy (1) & Andrew J Macpherson (1)

(1)Division of Gastroenterology, Department of Clinical Research, University Clinic for Visceral Surgery and Medicine, University of Bern, 3010 Bern (2)ETH Zurich, Institute of Molecular Systems Biology, Auguste-Piccard-Hof 1, 8093 Zurich, (3)These authors contributed equally to this work

Introduction: Metabolic capacity and immune system development of the host are dependent on colonization with commensal microbiota. There is evidence that signals originating from commensals during early life or from maternal microbiota before birth are required to shape the neonatal immune system.

Methods: Our system of reversible colonization of germ-free (GF) mice with the auxotrophic *E. coli* strain, HA107, allows us to expose pregnant mice to microbiota without subsequently colonizing their offspring. We use this system to detect maternal microbiota-derived metabolites that are transferred to the offspring and to detect their influence on neonatal immunity.

Results: After colonization of pregnant GF mice with ¹⁴C-labeled HA107, microbiota-derived products were present in placenta, fetus and in maternal milk as detected by liquid scintillation. Using ¹³C-labeled HA107 and mass spectrometry, we identified ¹³C-labeled products in the maternal milk and the offspring, indicating that maternal microbiota-derived products can reach the offspring. Exposing pregnant mice to HA107 increased the number of intestinal NKp46⁺ type 3 innate lymphoid cells (ILC3s) and F4/80⁺ CD11c⁺ mononuclear cells (MNCs) in the offspring and altered its sensitivity to LPS challenge.

Conclusion: Cross-fostering between reversibly colonized and GF mothers revealed that both ante- and postnatal transfer of maternal microbiota-derived metabolites are required to fully shape the neonatal immune system.

Efficiency and safety in case of technical success of EUS- guided transhepatic antegrade biliary drainage. Report of a monocentric study.

Sébastien Godat (1), Erwan Bories (2), Fabrice Caillol (2), Christian Pesenti (2), Jean Philippe Ratone (2), Chiara De Cassan (2), Marc Giovannini (2).

(1)Division of Hepato and Gastroenterology, Centre Hospitalier Universitaire Vaudois CHUV, Lausanne, Switzerland.

(2)Division of Gastroenterology, Paoli-Calmettes Institute, Marseille, France.

Background

EUS-guided biliary drainage like choledochoduodenostomy, hepaticogastrostomy (HGS), antegrade stenting and rendezvous are alternative procedures in case of obstructive jaundice and altered anatomy or failed ERCP. Complications related to EUS-guided antegrade drainage (EUS-GAD) are still described as substantial in up to 10% and combination of procedures is sometimes suggested to avoid adverse events.

Aims & Methods

To evaluate the efficiency and safety of EUS-GAD with transhepatic access in case of technical success. We retrospectively reviewed patients who underwent EUS-GAD in a single, tertiary care center.

Results

20 patients were included (9F/11M, mean age 68). Malignant stenosis in 19/20 (95%). Reasons for EUS-GAD was failed ERCP in 13/20 (65%), duodenal stenosis in 4/20 (20%), altered anatomy after surgery in 3/20 (15%). Intrahepatic biliary duct puncture was done with a 19G EchoTip® Needle in 16/20 (80%), with an EchoTip® Access Needle in 7/20 (35%). A cystostoma 6fr was always used to create the hepaticogastric tract, without puncture site closure. Stenosis dilatation was done in 15% and calibration with cystostoma 6fr in 45%. SEMS was transpapillary in 95%. Drainage was completed in intraoperative stage by HGS in 1/20 and by percutaneous drainage of the right liver in 1/20. Overall clinical success was 17/20 (85%). 1/20 presented a persistent obstructive cholangitis treated by another SEMS via ERCP. 2/20 patients died of infectious complication with incomplete drainage in case of advanced cancerous disease. One of these 2 patients was treated by EUS-GAD and HGS in same time. None patients developed bilioma or bile leakage.

Conclusion

EUS-GAD by transhepatic way is clinical effective and safety. Closure of the gastric puncture site is not mandatory if drainage is efficient. Complementary method for biliary decompression should be combined in case of incomplete drainage and not to prevent potential adverse events.

05 Clinical impact of routine anti-Tumour Necrosis Factor (anti-TNF) therapeutic drug monitoring (TDM) in the management of Inflammatory Bowel Disease (IBD) patients

Guirgis M¹, Bianchetti D², Dorta G², Michetti P¹, Maillard M²

1 Crohn's and Colitis Centre, Lausanne,

2 Lausanne University Hospital, Gastroenterology Service

BACKGROUND. Few studies have focused on the utility of regular TDM in the management of IBD. Our primary objective was to evaluate routine use of TDM for 3 TNF α inhibitors and observe physician response.

METHOD. A retrospective study was conducted on all IBD patients, treated with TNF α -inhibitors, attending two IBD referral centres. TNF α -inhibitors trough levels and anti-drug antibody (ADA) were measured using Lisa Tracker Premium Kit. Patient demographics and clinical data were retrieved from medical records.

RESULTS. A total of 143 anti-TNF α and ADA assays were measured routinely in 79 IBD patients. 88% of patients were treated with infliximab, 8% adalimumab and 4% certolizumab pegol. The prevalence of ADA was 10%. A sub-therapeutic result was found in 40% of patients resulting in no change in dose (66%), an increase in dose or frequency (32%), cessation of treatment (3%), a decrease in dose (2%) or an unrecorded response (3%). In response to ADA detection, 30% of clinicians increased the dose, 40% switched or stopped the agent and 20% did not change management.

CONCLUSION. This is the first study in Switzerland to evaluate utility of anti-TNF TDM. Routine TDM revealed subtherapeutic levels in 40% of cases and led to a change in clinical management in one third of cases. The approaches to subtherapeutic levels and presence of ADA varied between physicians. Further studies are required to develop consensus.

06 Outcomes and predictive factors of endoscopic management in case of relapse of high grade dysplasia or intramucosal carcinoma on Barrett's esophagus after successful initial endoscopic resection.

Sébastien Godat (1), Fabrice Caillol (2), Aurélie Autret (3), Erwan Bories (2), Christian Pesenti (2), Jean Philippe Ratone (2), Chiara De Cassan (2), Flora Poizat (4), Marc Giovannini (2).

(1)Division of Hepato and Gastroenterology, Centre Hospitalier Universitaire Vaudois CHUV, Lausanne, Switzerland.

(2)Division of Gastroenterology, Paoli-Calmettes Institute, Marseille, France.

(3)Division of Biostatistics, Paoli-Calmettes Institute, Marseille, France.

(4)Division of Pathology, Paoli-Calmettes Institute, Marseille, France.

Background

The role of endotherapy in case of relapse of High-grade dysplasia (HGD) and intramucosal carcinoma (IMC) on Barret's Esophagus (BE) after successful initial endoscopic resection (ER) is unknown.

Aims & Methods

To evaluate the efficiency of endotherapy in case of relapse of HGD/IMC on BE after successful initial ER. We performed a retrospective study, in a single tertiary care center. Relapse was defined by histological presence of HGD/IMC, with at least 6 months after the end of initial ER.

Results

18 patients were included (1F/17M, mean age 70). Initial median Prague classification of BE was C3M4. Initial ER was done by EMR in 16/18 (89%), ESD in 2/18 (11%), with adjuvant radiofrequency (RF) in 5/18 (28%) or laser Argon in 1/18 (5%). Stenosis occurred in 17%, always treated endoscopically. Median time between initial endotherapy and relapse was 18 months (range 6-33). 2/18 (11%) underwent immediate surgery and 16/18 (89%) an endotherapy for relapse. EMR was done in 16/18 (89%), adjuvant RF in 3/16 (19%), with a technical and carcinological success in 11/16 (69%). In case of failed endotherapy, 2/5 underwent radiochemotherapy (RC) and 3/5 a salvage surgery. In case of successful endotherapy, 3/11 showed a secondary relapse after an average time of 20 months (range 13-34). 1/3 patients was treated by RC, 1/3 by salvage surgery and 1/3 by 6 sessions of ER without disease progression after 36 months. In total, endotherapy was efficient for treatment of relapse in 8/18 (44%) with a follow up of 28 months. 1/18 presented a metastasis progression and 3/18 died (1/3 of complications after salvage surgery, 1/3 of a pancreatic cancer, 1/3 of an accidental disease).

Conclusion

Endotherapy could be a treatment for management of relapse of HGD or IMC on BE, but should be carefully used with strict follow up.

08 The outer mucus layer hosts a distinct intestinal microbial niche

Hai Li¹, Julien P. Limenitakis¹, Markus B. Geuking¹, Tobias Fuhrer², Irene Keller¹, Melissa Lawson¹, Uwe Sawer², Kathy D. McCoy¹, Andrew J. Macpherson¹

¹ Maurice Müller Laboratories (DKF), Clinic for Visceral Surgery and Medicine, University of Bern, Bern, Switzerland

² Institute of Molecular Systems Biology, ETH Zurich, Zurich, Switzerland.

Background: The mammalian lower intestinal tract harbors great density and diversity of commensal microbes, and a physical gel-like barrier named mucus is adopted in terms of limiting direct contact of commensal bacteria to host during host-microbe coevolution. The inner mucus layer with its tight compiled structure is impenetrable to bacteria, whereas a comparable dose of bacteria to luminal contents are living in the colonic outer mucus layer. We asked whether the outer mucus layer was a separate microbial niche regarding bacterial kinetics and behaviors.

Methods: Longitudinal microbial community and distribution in cecal and colonic mucus and contents of mice containing specific pathogen free microbiota or 12 defined culturable organisms were analysed using 16s rRNA lontorrent sequencing. Transcriptional patterns of bacteria in colonic mucus and contents of *E. coli* and *B. thetaiotaomicron* monocolonised or bicolonised mice were compared by high-throughput RNAseq. Differentially expressed genes were subjected to STRING database using SEED mode for functional clustering gene sets. Mucus thickness measurement was performed following publication with modifications. Alternatively, Carney's fixed intestinal sections were administrated to immunohistological staining for detecting mucus. 2-photon microscopy was used to take time-lapse videos for detecting GFP positive bacterial tumbling in colonic mucus. The comparison of rates of bacterial replication in intestinal compartments on a per-bacterium basis was addressed by replication-caused-GFP-dilution taking advantage of plasmid pDIGI or chasing radioactivity decay of metabolically ³²P-radio-labeled bacteria. Metabolites in luminal contents and mucus or in *in vitro* bacterial cultures on germ-free mice mucus and contents extracts were compared using non-targeted mass spectrometry.

Results: Our data show that the outer mucus layer had different representations of microbes compare to adjacent luminal contents, and the exchange between the two compartments was very limited due to immobility of bacteria in mucus layer. Furthermore, by comparing the transcriptional and metabolic patterns of two model bacterial species, *Escherichia coli* and *Bacteroides thetaiotaomicron*, with exceedingly different mucolytic capability, we showed mucus and contents associated bacteria differentially shaped their metabolic pattern to adapt to the nutrients accessible in the two compartments. This was not limited to the carbon source, including host-derived phospholipids for *E. coli* and host or dietary glycans for *B. thetaiotaomicron*, but also included essential minerals.

Discussion: We proved the outer mucus layer is a unique niche for commensal microbes and intensive study of host-microbial mutualism in outer mucus layer is important to uncover the mechanism in clinic diseases as the mucus associated bacteria have a physical nearby impacts to the host.

Portal hypertension in a Crohn's patient with small intestinal bleeding treated by transjugular portosystemic shunt placement

Benjamin Heimgartner, Andrea De Gottardi, Andrew Macpherson, Jan Hendrik Niess

University Clinic for Visceral Surgery and Medicine, University of Bern

Introduction: Crohn's disease patients treated with azathioprine can develop nodular regenerative hyperplasia (NRH), portal hypertension (PHT) and portal vein thrombosis. PHT and inflammatory mucosal lesions increase the risk for gastrointestinal bleedings.

Case description: A 56 year old patient with Crohn's disease developed NRH under treatment with azathioprine. He was admitted with severe bleeding. Gastroscopy showed small esophageal varices without bleeding stigmata. Blood was detected in the terminal ileum. CT angiography revealed a partial portal vein thrombosis with extension to the superior mesenteric vein, thickening of the jejunal wall and splenomegaly. Because conservative treatment was not sufficient to control bleeding, a TIPS procedure and aspiration of the thrombus was performed. The immunosuppressive medication was switched to infliximab to control the activity Crohn's disease. The blood panel including thrombocytopenia improved and bleeding stopped.

Discussion: PHT in a Crohn's patient with NRH and partial obstruction of the mesenteric venous inflow to the liver can be efficiently treated by TIPS. In our case TIPS and systemic treatment with infliximab lead to the improvement of the blood panel and remission of the Crohn's disease.

O9

Identification of potent anti-proliferative agents against hepatic angiosarcoma

Christian Perez-Shibayama^{1,2}, Marianne Kraus³, Christoph Driessens³, David Semela¹

¹Klinik für Gastroenterologie und Hepatologie, Kantonsspital St. Gallen, 9007 St. Gallen. ²Institut für Immunbiologie, Kantonsspital St. Gallen, 9007 St. Gallen. ³Klinik für Onkologie und Hämatologie, Kantonsspital St. Gallen, 9007 St. Gallen

Background: Hepatic angiosarcoma (AS) is a rare and highly aggressive tumor. Conventional chemotherapy and first trials with anti-angiogenics showed only minor efficacy; new therapeutic approaches are therefore urgently needed. Our aim is to characterize potential new anti-tumor strategies against AS *in vitro* and *in vivo* by using our recently described murine angiosarcoma model (Rothweiler *et al*, 2015).

Methods: *In vitro* screening for effective pharmacological compounds was performed using MTS assays in isolated angiosarcoma cells. Identified anti-proliferative drugs were further assessed using *in vitro* capillary formation on Matrigel and by flow cytometry.

Results: Screening showed strong anti-proliferative effects by the proteasome inhibitor Carfilzomib, the protease inhibitor Lopinavir and the pro-apoptotic compound Navitoclax. These compounds were active in nanomolar to micromolar ranges when evaluated as mono-therapeutic agents. Importantly, combination strategies using these compounds showed synergistic effects. Consistently, the tube formation capacity of AS cells on matrigel was strongly affected after treatment with these compounds. Flow cytometric analysis of the hepatic AS cell line after treatment revealed that the possible mechanism of cell death induced by the anti-tumor drugs was apoptosis.

Conclusions: We have identified three clinically available compounds with potent anti-proliferative effects against difficult to treat angiosarcoma. Combinatory treatments showed synergistic effects and are therefore promising candidates in AS treatment. *In vivo* models will complement and further support possible new strategies to treat hepatic angiosarcoma.

O11

A severe complication after laparoscopic low anterior rectum resection: a compartment syndrome of the upper extremity

R. Kraus, G. Curti, Department of Surgery, Kantonsspital Aarau

O10

Survival after CRS/HIPEC for colorectal and appendicular peritoneal malignancy

O12

Marcel Schneider, Dilmurod Eshmuminov, René Vonlanthen, Philippe Gertsch, Kuno Lehmann

Background: Cytoreductive Surgery (CRS) and hyperthermic intraperitoneal chemotherapy (HIPEC) is a combined treatment for well-selected patients with peritoneal carcinomatosis (PC). Despite an increasing number of centers, survival data from Switzerland is scarce. **Methods:** Data of 130 patients with colorectal or appendicular carcinomatosis from two centers (Bellinzona n=17, 1998-2008) and Zurich (n=113, 2009-2014) were prospectively collected and analyzed. Patients with malignant disease received standard perioperative chemotherapy. Patients with low-grade appendix tumors were directly operated. HIPEC was performed after radical cytoreduction (CC-score 0, no visible tumor). Follow-up included clinical exams, tumor markers and CT scans every six months. **Results:** Patients had carcinomatosis from appendix tumors in 63% (82/130), including low-grade (40/82) and high-grade (44/82) tumors, and colorectal cancer in 37% (48/130). Curative surgery was possible in 73% of operated patients. Major morbidity and mortality were 8.1% and 2%, and follow-up was 25 months. For colorectal PC, median overall (mOS) and disease free (DFS) survival were 34 and 12 months, and 3-year survival was 48%. For low-grade appendix tumors, OS and DFS were 100% and 87% at 3y. For high-grade appendix tumors, mOS was not reached, DFS was 28 months, 56% were disease free and 69% alive at 3 years. Signet ring differentiation was a highly negative prognostic factor on survival for colorectal and appendix tumors ($p<0.001$). **Conclusion:** Disease free survival after curative CRS/HIPEC is excellent for appendix tumors, and well-selected patients with PC from colorectal cancer have a survival benefit. Patients with signet ring differentiation show worse outcomes.

Introduction

Laparoscopic surgical procedures decrease the tissue trauma compared to open surgery and result in less postoperative pain what have been shown in different studies. Despite proven advantages there are severe complications occurring during a laparoscopic procedure due to the positioning during surgery. We present a case of a compartment syndrome of the upper extremity after laparoscopic low anterior rectum resection.

Method

A 57y old man underwent an incomplete endoscopic resection of a pT1 adenocarcinoma 8cm above the anal margin line therefore the patient underwent a laparoscopic low anterior rectum resection with a defunctioning loop ileostomy.

Result

The patient has been placed on a bean bag in lithotomy position with the right arm close to the body. The right arm has been padded and then the vacuum has been applied to the bean bag forming the bag safely around the body. During the procedure the patient has been steep head-down and tilted to the right side for 6 hours. Shortly after the extubation the patient complained of heavy pain in his right forearm and showed clinically a compartment syndrome. Immediately the hand surgeons opened the carpal tunnel and all the muscle compartments of the hand and forearm.

Conclusion

This case illustrate the need of appropriate positioning and repositioning during surgery to prevent compartment syndrome not only of the lower extremity and to check the patient's limb after surgery.

Evaluation of minimal hepatic encephalopathy using standard psychometric tests and computer assisted inhibitory control test (ICT) in patients with cirrhosis

Laurent Spahr and Rachel Goldstein. Gastroenterology and Hepatology, Neurology, HUG

Minimal hepatic encephalopathy (MHE) is frequent (>50%) and presents as a spectrum of cognitive deficits in a cirrhotic patient with a normal neurological exam. Psychometric tests (PsyT) are used to detect MHE, but they are influenced by age and education and require psychological expertise and training. The ICT, a computerized test for assessment of attention and response inhibition, has been reported as sensitive to diagnose MHE (Bajaj, GE 2008). **Methods:** We explored the performance of healthy subjects (n=23 ; age 43 yrs) and patients with cirrhosis (n=30 ; age 55 yrs ; M/F : 20/10 ; alcoholic cirrhosis 77% ; Child A/B/C : 5/19/6 ; lactulose therapy: 11/30) on a battery of PsyT (Trail-A, Trail-B, Digit Symbol Test, DST) and the computerized ICT administered to detect MHE.

Results: values are given as Z-scores (raw score ; time ; correct lure response).

Test	Controls	Cirrhosis	P value
Trail-test A	0.21 ± 0.7	-3.14 ± 4.4	<0.0001
Trail-test B	-0.47 ± 1.1	-4.86 ± 4.3	<0.0001
DST	0.92 ± 0.6	-0.73 ± 0.6	<0.0001
ICT (correct lure response)	0.51 ± 0.5	-0.39 ± 1.1	<0.0001

The ICT could be completed by all patients. Patients with cirrhosis were significantly impaired with regard to PsyT and ICT performance compared to controls. Twenty one patients (70%) had abnormal values on at least 2 tests of the PsyT battery, meeting the accepted criteria for MHE diagnosis (EASL Guidelines). Using ICT, 15 patients (50%) were deficient and met MHE criteria. **Conclusion:** In this small group of patients with cirrhosis, both psychometric tests and the ICT allow to diagnose MHE. The use of ICT by non specialists could increase the testing rate for MHE. (Supported by FLAGS in Geneva)

O13

An unusual cause of an obscure gastrointestinal bleeding in the small bowel - sometimes capsule endoscopy only shows "the tip of the iceberg"

Andreas Stulz¹, Walter Arnold², Patrick Aepli¹

1 Division of Gastroenterology and Hepatology, Department of Medicine, Luzerner Kantonsspital, Lucerne

2 Department of Pathology, Luzerner Kantonsspital, Lucerne

Background: Obscure gastrointestinal bleeding (OGIB) is defined as recurrent or persistent bleeding or the presence of iron deficiency anaemia after evaluation with negative gastroscopy and colonoscopy. OGIB accounts for 5% of GI bleeding and presents a diagnostic challenge. Capsule endoscopy and balloon assisted enteroscopy remain the cornerstone of investigation in OGIB given their high diagnostic yield.

Methods: We present the case of an 86-year-old patient with a 1-year-history of severe iron deficiency anaemia in spite of repeated blood transfusions and iron therapy. Over the past 16 years the patient had been treated for several malignancies (colorectal, prostate, melanoma).

Results: Gastroscopy (incl. duodenal biopsies) and colonoscopy were normal, in particular no bleeding source was seen. We then performed a capsule endoscopy which showed a few small polypoid lesions and a minimal amount of blood in the proximal jejunum. In response to these findings, an anterograde (single balloon) enteroscopy was performed where several ulcers up to 3 cm in size with noticeable black colouration were found in the first 1.5 metres of the small bowel. Extensive biopsies were taken. The final histological results confirmed our suspicion that these lesions were metastasis from a melanoma.

Conclusion: Sometimes the findings in a capsule endoscopy are minimal because they show only the "tip of the iceberg". As a result every relevant lesion in a capsule endoscopy should be followed by an enteroscopy which allows detection, biopsy and treatment.

O15

Familial Eosinophilic Esophagitis (EoE) Uncovers a New EoE-like Syndrome without Tissue Eosinophilia O14

Straumann A,¹ Blanchard C,² Radonjic-Hoesli S,³ Bussmann C,⁴ Hruz P,⁵ Kozlowski E,³ Safroneeva E,⁶ Simon D,⁷ Schoepfer A,⁸ Simon HU,³; 1 Swiss EoE Center Olten, 2 Nestlé Research Center, 3 IKP University of Bern, 4 Violiier Pathology Basel, 5 University Hospital Basel, 6 ISPM University of Bern, 7 Dermatology, Inselspital/University of Bern, 8 CHUV, Lausanne

Background: Eosinophilic esophagitis (EoE) is a chronic-inflammatory, genetically-impacted disease with rapidly increasing incidence, defined clinically by symptoms of esophageal dysfunction and pathologically by an eosinophil-predominant infiltration. We identified in four EoE-families five members presenting with EoE-typical and steroid-responsive symptoms, but without tissue eosinophilia. We aimed to investigate this intriguing syndrome of "EoE without eosinophilia".

Methods: The five patients suffering from EoE-like syndrome were evaluated by endoscopic, histologic and quantitative immuno-histologic examinations, and by determination of mRNA expression.

Results: The frequency of first generation offspring of EoE-like syndrome patients affected by EoE or EoE-like syndrome was 40%. Immunohistochemistry confirmed definitely a lack of eosinophils in esophageal tissue of patients with EoE-like syndrome, but revealed a considerable T cell infiltration comparable with EoE. In contrast to EoE, eotaxin-3 mRNA was not, and eotaxin-3 protein markedly less elevated in EoE-like syndrome. The mRNA expression of selected EoE-genes was dysregulated in EoE-like syndrome, either at the level of EoE or at a level in-between EoE and normal epithelium.

Conclusions: These five members of EoE-families suffering from "EoE without eosinophilia" do formally not fulfill the diagnostic criteria of EoE. However, clinical manifestation, the bequeath of EoE to their offspring, immunohistochemistry and gene-expression pattern suggest a uniform underlying pathogenesis. Conventional EoE with the predominant eosinophilia is therefore likely only one phenotype of a broader "inflammatory dysphagia syndrome". The role of the eosinophils, the definition of EoE and its diagnostic criteria must be reconsidered.

Clinical Long-term Outcome of Patients Treated With Endoscopic Rendez-vous Dilatation of Complete Obliterations in the Proximal Oesophagus

Michael C. Sulz¹, Reto Bertolini^{1*}, Paul Martin Putora², Franziska Albrecht¹, Martina A. Broglie-Däppen³, Sandro J. Stöckli³, Christa Meyenberger¹. *The first two authors contributed equally to the abstract.

¹Gastroenterology and Hepatology, Kantonsspital St. Gallen, St. Gallen, ²Radiation Oncology, Kantonsspital St. Gallen, St. Gallen, ³Head and Neck Surgery, Kantonsspital St. Gallen, St. Gallen, Switzerland

Background: Complete obliteration in the proximal oesophagus can occur after radiotherapy in patients with head and neck cancers. If antegrade dilatation is unsuccessful, retrograde endoscopic rendez-vous dilatation is an option to restore oesophageal patency and to resolve patients from percutaneous endoscopic gastrostomy (PEG) tubes. Our aim was to focus on the technical and long-term clinical outcome.

Methods: This case series included all patients in one tertiary referral centre with complete obliteration in the proximal oesophagus over a 10-year period, who were unable to swallow and were treated by retrograde endoscopic rendez-vous dilatation. Technical success was defined as achievement of a retrograde puncture and passage with the endoscope to apply a nasogastric tube. The long-term clinical success was defined as either independency from PEG-tube and/or ability of oral food intake.

Results: 6 patients (5 males; median age 71 years) underwent retrograde endoscopic rendez-vous dilatation. Technical success was achieved in 5 of 6 patients. At follow-up (median 27 months) half of the patients stayed dependent on the PEG-tube, the others had oral diet.

Conclusions: This approach offers an alternative to high-risk blind dilatation or surgery in patients with complete proximal oesophageal obliteration. We found that technical success was fairly high. However in long-term follow up only half of the patients remained independent from PEG-tube.

Short-term Outcomes of Transanal Total Mesorectal Excision for Low Rectal Cancer

Rebekka Troller, Michel Adamina, Felix Grieder, Hans Gelpke, Stefan Breitenstein. Klinik für Viszeral- und Thoraxchirurgie, Kantonsspital Winterthur, Winterthur.

Background

Transanal total mesorectal excision (taTME) is an alternative to conventional mesorectal excision owing to its ability to achieve clear distal and circumferential resection margin in low cancers.

Methods

Consecutive patients were included in a prospective cohort study. Perioperative and functional outcomes were measured along regular clinic visits and with validated questionnaires. Results were reported as median and interquartile range (IQR).

Results

28 patients with a low rectal cancer (6.5cm to anal verge, IQR 6-8) underwent a taTME between Feb 2013 and Jan 2015. Age and body mass index were 66.5 years (IQR 62.5-76) and 26.5kg/m² (IQR 24.6-30.7). 17 patients had neoadjuvant radiochemotherapy. Median surgery time was 356.5 minutes (IQR 326.2-420.7), including an ileostomy. Median length of stay was 13.5 days (IQR 10.2-15.5). Mortality was 0 and 30-day morbidity totalized 40% minor and 18% major complications (Dindo-Clavien I-II, resp. III-IV). Dissection of the mesorectum was good (Quirke 3) and all distal and circumferential margins were clear. Median T stage was 3 (IQR 2-4). 3 patients had lymphnode metastases for a median number of retrieved nodes of 22 (IQR 14.5-34). Functional outcomes were assessed in 10 patients (10-month follow-up), of those 6 were stoma-free. Quality of life was good with a QLQ-C30 score of 70.8 (IQR 66.7-75), as was the urinary function (ICS voiding 7, IQR 6.3-10; incontinence 7.5, IQR 6-8). Sexual function in males (IIEF-5 score 2.5, IQR 2-17.5) and females alike (FSFI 6, IQR 3-6) was limited. Anal continence score was acceptable (FISI 17, IQR 2-28), as was the bowel emptying score (17, IQR 15.5, 17.8).

Conclusion

Transanal total mesorectal excision appears as an oncologic and functionally suitable procedure for distal rectum cancer.

O17

Successful stenting of a stenosis in the hepatic venous outflow tract in a patient with secondary Budd-Chiari-syndrome and refractory ascites

Daniel Venetz¹, Martin Hoffmann², Patrick Aepli¹

¹ Division of Gastroenterology & Hepatology, Department of Medicine, Luzerner Kantonsspital, Lucerne

² Department of Radiology, Luzerner Kantonsspital, Lucerne

Background: Transjugular intrahepatic portosystemic shunt (TIPS) is often the treatment of choice in patients with portal hypertension and refractory ascites due to chronic liver failure. Whether transjugular stent insertion in the hepatic venous outflow tract is effective in patients with portal hypertension and refractory ascites due to posthepatic obstruction is less clear.

Methods: We present the case of a 44 year-old patient with portal hypertension and refractory ascites due to a membranous stenosis at the level of the hepatic venous outflow tract, resulting in a secondary Budd-Chiari-syndrome with a singular remaining left hepatic vein, atrophic right liver lobe and hypertrophic left liver lobe. We don't know whether the right lobe was congenitally agenetic or underwent secondary atrophy due to visible liver vein obstruction.

Results: Transjugular percutaneous transluminal angioplasty (PTA) of the web-like, membranous stenosis between the singular remaining left hepatic vein and the inferior vena cava followed by stent insertion (18 x 40 mm) resulted in immediate reduction of the pressure gradient from initially 22 mmHg to 0 mmHg after the intervention. Stent insertion led to complete and rapid resolution of ascites.

Conclusion: Transjugular stent insertion between the hepatic outflow tract and the inferior vena cava can be a safe and effective treatment in patients with portal hypertension with refractory ascites due to posthepatic stenosis.

O19

Percutaneous Tibial Nerve Stimulation (PTNS): a successful treatment for patients with anal incontinence (urgency), leading to a sustained response and modulating anal sphincter pressure and rectal perception

Boudewijn Van der Weg B¹, Andreas Frenzer¹, Beat Muggli¹, Jan Borovicka², Ulrich Baumann³, Radu Tutuian⁴

¹Spital Thun, ²Kantonsspital St. Gallen, ³Spital Münsingen,

⁴Universitätsspital Bern

Background: To assess the long-term clinical response, anorectal motor and sensory function in incontinent patients treated by PTNS

Methods: Retrospective analysis of prospective gathered data of 100 patients with anal urge incontinence treated with PTNS. Patients had 21 treatment sessions within 8 months. St. Marks Incontinence-score and Visual Analogue Scale (VAS)-score were collected at baseline, after 3, 8, 10 and 20 months. All patients had a high resolution manometry at baseline and after 12 weeks.

Results: A successful clinical response ($\geq 50\%$ reduction of the St. Marks, resp. VAS-score) was reached at end of treatment (8 months) in 74% resp. 82.2% and at one year after end of treatment (20 months) in 71.1%, resp. 80.6% of the patients ($p<0.001$). The sphincter squeeze pressure significantly improved from 126.5mmHg at baseline to 137.6 mmHg ($p<0.001$) at 12 weeks. The constant perception for urge changed from 101.4ml to 120ml ($p=0.002$), the maximal tolerable volume from 155.5ml to 183.7ml ($p<0.001$) after 12 weeks PTNS treatment. A significant, although weak correlation between clinical response and improvement of rectal sensory function was observed ($p=0.04$, Spearman coefficient=-0.25)

Conclusions: PTNS is a minimal invasive treatment, with negligible side effects and a response rate of 80%. The therapeutic effect sustains at least one year after end of treatment. The treatment leads to a significant improvement of the sphincter squeeze pressure as well as the rectal perception.

O18

Two similar cases of elderly women with moderate abdominal pain and pneumoperitoneum of unknown origin: A surgeon's successful conservative management

Fabrizio Vinzens¹, Valentin Zumstein¹, Christoph Ackermann¹.

¹Department of Surgery, St. Claraspital, Basel.

O20

Background: Patients presenting with abdominal pain and pneumoperitoneum in radiological examination usually require emergency explorative laparoscopy or laparotomy. Mostly pneumoperitoneum associates with gastrointestinal perforation. There are very few cases where surgery can be prevented in patients with a combination of pneumoperitoneum and abdominal pain. People could benefit from a non-surgical treatment.

Methods: We present a successful conservative management of an 89 and 87-year-old female patient with moderate abdominal pain and free abdominal gas of unknown origin.

Results: Two elderly women presented with moderate abdominal pain in our emergency room in a short period of time. There was no medical intervention in the recent past history. Physical examination revealed mild tenderness in the upper respectively lower abdomen, but no clinical sign of peritonitis. Cardiopulmonary examination remained unremarkable. Blood studies showed no abnormalities, in particular inflammation parameter were within normal limits. Finally an obtained computed tomography (CT) showed free abdominal gas of unknown origin in both cases. We performed a conservative management with nil per os, total parenteral nutrition and prophylactic antibiotics. After 2 weeks both patients were discharged home.

Conclusion: Conservative management should be taken into account in patients with moderate abdominal pain and pneumoperitoneum of unknown origin in absence of clinical signs of peritonitis.

Non-CO2-insufflation and acute abdominal compartment during emergency endoscopy contributing to fatal outcome in advanced obstructive tumor of the gastro-esophageal junction with severe bleeding. ¹⁺Reiner Wiest, ¹⁺Jan Niess, ²Hans-Ulrich Rothen, ²Tobias Merz, ²Jukka Takkala; ¹Thomas Malinka, ¹Christian Seiler; ¹Department Visceral Surgery and Medicine (Inselspital); ²Department Intensive Care Medicine, (Inselspital), Freiburg-strasse, 3010 Bern; ¹⁺: equal contribution

Case: A 38 y old man suffering from advanced adenocarcinoma of the gastro-esophageal junction under palliative therapy was admitted with sever upper gastrointestinal bleeding. Emergency endoscopy utilizing room air was performed showing an highly obstructive tumor with active bleeding. The tumor could hardly but finally, successfully be passed and bleeding was stopped successfully by adrenalin injections and clipping. Immediately after endoscopy the patient showed clinical signs of an acute, severe abdominal compartment syndrome. CT imaging revealed a completely compressed abdominal aorta induced by massively distended and pressurized intestinum. Even before emergency laparotomy could be performed the patient presented with pulsless electrical activity and died. Conclusion: CO2 is recommended to be used in emergency endoscopies – at least in cases with obstructing tumors. Pressure sensing devices should be developed and utilized improving safety of endoscopic procedures.

O21

Systematic immunohistochemical screening for Lynch syndrome in colorectal cancer: a single-center experience of 488 patients

V. Zumstein¹, F. Vinzens¹, A. Zettl², K. Heinemann³, D. Köberle⁴, M. von Flüe¹, M. Bolli¹. ¹Department of Surgery, St. Claraspital, Basel; ² Department of Pathology, Biolier AG, Allschwil, ³Department of Medical Genetics, University Hospital Basel, ⁴Department of Oncology, St. Claraspital, Basel.

Objective: Mutations in DNA mismatch repair (MMR) genes cause autosomal dominantly Lynch syndrome (LS), characterized by early development of colorectal cancer (CRC) and other cancers. LS patients benefit from intensive cancer surveillance. We aimed to assess detection rate of LS at our institution after introduction a systematic immunehistochemical (IHC) screening of CRCs in 2011.

Methods: Following EGAPP recommendations, starting in June 11, all CRCs were prospectively tested by IHC for the presence of MMR proteins MLH1, PMS2, MSH2 and MSH6. In case of loss of MLH1, BRAF mutation V600E was assessed by molecular testing/IHC. Follow-up and genetic counseling was evaluated in patients suspected being LS carriers (i.e. tumors showing loss of MLH1 combined with absence of BRAF V600E, loss of PMS2, MSH2 or MSH6).

Results: Loss of MMR expression was found in 71 (14.5%) of 488 patients (76 ± 13 year) who underwent CRC surgery from June 11 to May 15. 27 (5.5%) patients were classified as potential LS carriers (68 ± 14 years). Sporadic MMR deficiency patients had a mean age of 78 ± 18 years. In 27 potential LS carriers, genetic counselling and germline testing was recommended, which revealed in 60% a positive result for LS, corresponding to a LS rate of 3.2% out of all our CRC patients.

Conclusions: All CRC should be tested by IHC. Tumorboard protocols should systematically evaluate IHC status; suspected LS patients need genetic counselling and testing.

O22

POSTERS GASTROENTEROLOGY

Measurement of functional blockade of TNF-alpha by anti-TNF agents is a stronger predictor than trough levels and anti-drug antibodies: 3-year prospective clinical data.

Pascal Juillerat^{1,2}, Peter Andrew¹, James Macpherson¹, Emma Slack¹, Nicola Putato^{1,2}, Julia Cahenzli¹, Flavia Schmid¹, Frank Seibold^{1,3}, Kathy D. McCoy¹, Andrew J. Macpherson^{1,2},
¹ University Bern, Maurice E. Müller Laboratories, Universitätsklinik für Viszeralre Chirurgie und Medizin, Inselspital, Bern.
² Clinic for Visceral Surgery and Medicine, Gastroenterology, Bern.
³ Gastroenterology, Tiefenauhospital, Spitalnetzbern, Bern.

Background: In case of loss of response (LOR) to anti-TNF agents in inflammatory bowel disease (IBD) patients, interventions, such as dose increase and shorten the interval, lead only to a transient improvement and a majority of patients will eventually loss response. A functional in vitro test (CD-62L shedding) measuring TNF functional blockade should help us identify those specific situations.

Method: An in vitro test was used to predict the response to the drug: the shedding of the L-selectin (CD62L) quantified by flow cytometry on the surface of granulocytes before and after the anti-TNF agent administration. In a subgroup of patients trough level of the drug (TL) and antibodies against the drug (ADA) have been performed in order to compare both tests. The treatment strategy during the 2 years of the study was blinded to the results of the CD62L shedding, TL and ADA and followed clinical-based interventions or switch by IBD specialists.

Results: From June 2012 to May 2015, 33 IBD treated with anti-TNF agents at Bern University Hospital were followed prospectively (clinicians blinded) to correlated clinical outcome with their response profile tested at baseline. The 22 responders (R) and 11 non responders (NR) had similar clinical characteristics. During a mean follow up of 25 months (range 7-41; 77 patient-years follow up), 25 medico- surgical events occurred (3 adverse events (AE), 1 CMV colitis, 16 flares treated with medication, 3 intestinal resections and 2 operations of fistula) 9 in R and 16 in NR, which means 14% vs. 60% ($p < .001$) of the patient-year follow up. The mean calprotectin during follow up (about 1 measurement per patient year available) was 119 (+/-139 SD) for R and 310 (+/- 226 SD) for NR. ADA and TL measurement could be performed in 15 patients (45%; 9 R and 6 NR). Only 2 patients developed ADA (one in each group). There was no significant difference in trough levels between R and NR (2.8 vs. 4.8; $p=0.4$) and 62% had a therapeutic level (>1.5). Patients stable without need for intervention were 16/21 (84%; 1 AE) in R vs. 1/9 (2 AE) in NR ($p<0.001$). In the NR group all the dose optimization failed, whereas in the responders group, interventions that would have been suggested on the basis of TL and ADA have not been performed (clinicians blinded), but were finally not required, based on the favorable clinical outcome.

Conclusions: Testing the in vitro functional blockade of TNF alpha (CD62L shedding) in anti-TNF treated IBD patients seem to be a better long term predictor than trough levels and antibodies against the drug measurements. This could minimize interventions and therefore reduce costs and risk of adverse events.

G1

The impact of the intestinal microbiota on eosinophils

Yasmin Kölle, Kathy McCoy

Department klinische Forschung, University of Bern

Background:

Gastrointestinal eosinophils are a major innate cell type in the lamina propria. In many inflammatory disorders eosinophils have been demonstrated to accumulate and are linked to fibrotic remodeling of the tissue. Since the intestinal microbiota has a strong impact on the immune status of the intestine (in health, active disease and remission), we asked how microbial colonisation influences the life cycle of eosinophils, their turnover and functional properties in the small intestine.

Methods:

We employed germ-free mice and deliberately colonized them with a complex microbiota. We used quantitative real-time PCR and RNA sequencing to assess transcriptome activity; electron microscopy to perform ultrastructural analysis in the tissue; cell cycle analyzing dyes (BrdU and DAPI) to analyse the life cycle turn over of eosinophils. Finally, we have re-derived the eosinophil-deficient mouse strain dblGATA1^{-/-} to investigate microbial responses in the absence of eosinophils.

Results:

We show that eosinophil proliferation is stimulated in the bone marrow following bacterial colonisation that brings along a transient eosinophilia in the blood. Under steady state conditions, eosinophils show higher intestinal turnover in the presence of microbes that reflects a more activated phenotype and alterations in gene transcription.

Conclusions:

Microbial colonisation feeds into the eosinophil life cycle and causes functional alterations that may turn out to be relevant in inflammatory conditions of the intestine.

G2

Acute Herpes simplex viral Esophagitis occurring in immuno-competent Individuals with Eosinophilic Esophagitis

Dorothee Zimmermann¹, Dominique Criblez¹, Evan Dellon², Christian Bussmann³, David Pfeiffer⁴, Matthias Froh⁵, Alex Straumann⁵

¹Division of Gastroenterology, Kantonsspital Lucerne, ²Division of Gastroenterology, University of North Carolina, Chapel Hill, USA, ³Division of Pathology, Vioillier, Basel, ⁴Division of Pathology, Kantonsspital Lucerne, ⁵Division of Gastroenterology, Kantonsspital Olten.

Background Herpes simplex esophagitis (HSE) is an acute, severe virus infection of the esophagus rarely occurring in immuno-competent individuals. Eosinophilic Esophagitis (EoE) is an immune-mediated esophageal disorder which is also rare.

Description of Cases We recently observed 5 severe HSE cases (m/f = 3/2, mean age 26 years), all affecting patients having a diagnosis of EoE. The coincidence of these two rare conditions raises the question of a causal relationship. Because 4 of the 5 patients had active, untreated EoE at the time of infection, a side effect of swallowed topical corticosteroids, the first line medical treatment of EoE, is likely not the explanation.

Conclusion The striking accumulation of HSE and EoE is suspicious that active, untreated EoE might predisposes to esophageal herpes simplex infection.

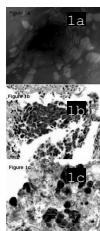


Figure 1a-c:
Endoscopy (1a),
histology (1b) and
immuno-
histochemical
staining (1c) in
one of the EoE
patients during
HSE.

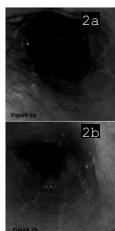


Figure 2a-b:
Endoscopy
during HSE (2a)
and follow up
EGD (2b) in one
patient who was
diagnosed with
EoE after an
episode of acute
HSE.

Parent-assessed EoE activity in children aged 3-9 years does not correlate with physician global assessment, endoscopic and histologic activity

Ekaterina Safroneeva¹, Micheal Coslovsky¹, Radek Panczak¹, Claudia Kuehni¹, Marcel Zwahlen¹, Sven Trelle¹, Andreas Limacher¹, Nadine Haas¹, Alex Straumann², Alain Schoepfer³; international pediatric EEsAI study group, ¹ISPM University of Bern, ²CHUV, Lausanne, ³Swiss EoE Center Olten.

Background: The international EEsAI study group is currently developing a symptom-based activity index for pediatric EoE patients aged 3-17 years. For children aged 3-9 years, parents report observed symptoms while for children aged ≥ 10 years there is parent and patient reporting of symptoms. We report on the findings in children aged 3 to 9 years.

Methods: Parents completed the symptom questionnaire with a recall period of 7 days. Children then underwent upper endoscopy with esophageal biopsy sampling. Parents (based on observed symptoms, ParGA) and physicians (based on patient history, endoscopic and histologic findings, PhysGA) provided in addition an overall assessment of EoE activity on a Likert scale from 0 (inactive EoE) to 10 (very active EoE).

Results: Ninety-three children were included (73% boys, median age 6.0 years, IQR 5.0-8.0 years, 42% with concomitant asthma, 65% with rhinoconjunctivitis). ParGA of EoE activity was 2.0 (IQR 0.0-4.0) while PhysGA was 2.0 (IQR 1.0-6.0). ParGA did not correlate with PhysGA ($r=0.01$, $p = 0.892$) and neither with peak eosinophil counts/hpf ($r = 0.18$, $p = 0.173$) and distinct endoscopic features (rings, exudates, edema, stricture, furrows; all p -value >0.1).

Conclusions: Parent assessment of children's EoE symptom severity does not correlate with endoscopic or histologic activity.

Short and long term mortality after percutaneous endoscopic gastrostomy (PEG) tube placement. Laurent Bochatay, Emiliano Giostra, Nicolas Goossens, Jean Louis Frossard.

Gastroenterology and Hepatology Service, Geneva University Hospital Geneva, Switzerland. **Background:** Patient with too short life expectancy should not receive PEG tube. Furthermore, it is well known some evolving disease do not benefit from PEG tube placement, e.g. patient with dementia. We aim to study the short and long term mortality after PEG tube placement at the gastroenterology unit of the university hospital of Geneva between 2011 and 2012. **Methods:** All patients that had a PEG procedure between January 2011 and December 2012 were included. Age, sex, indication for PEG tube placement, judgment capacity, type of sedation, ASA score, endoscopist experience, antiaggregant treatment, biological value and cause of death were analysed. Short term and long term mortality were defined as death occurring within or after 30 days following PEG procedure respectively. **Results:** 90 patients had a PEG tube placement and 10 died within 30 days (11%). On univariate analysis, age, indication, higher ASA score, use of antiaggregant treatment and higher creatinine level were significantly associated with 30 days mortality. On multivariate analysis, age ($p=0.039$), use of antiaggregant treatment ($p=0.08$) and higher creatinine level ($p=0.03$) were associated with 30 days mortality. Finally 36 death (40%) were observed. The mean survival time was 250 days (4-990) whereas the median of survival was 497 days. Progression of neurological condition and pulmonary sepsis were the main causes of death. **Conclusions:** As previously described in the literature, age, renal function and use of antiaggregant are associated with 30 days mortality. Even with a lack of power due to the small number of events, we can observe that patient with neurologic problems are most at risk of death within 30 days.

G4

G6

Escalating Incidence of Eosinophilic Esophagitis in Canton of Vaud, Switzerland, 1993 to 2013: a population-based study

Bénédicte Giriens¹, Ekaterina Safroneeva², Marcel Zwahlen², Alex Straumann³, Alain Schoepfer¹, ¹CHUV, Lausanne, ²ISPM University of Bern, ³Swiss EoE Center Olten.

Background: Eosinophilic esophagitis (EoE) is a chronic, inflammatory disease of the esophagus with a rapidly increasing incidence. However, population-based epidemiologic data on EoE are rare and limited to regions with less than 200,000 inhabitants. We evaluated the incidence and prevalence of EoE over time in Canton of Vaud, Switzerland.

Methods: Canton of Vaud lies in the French-speaking, Western part of Switzerland. As of 12/2013, it had a population of 743,317 inhabitants. We contacted all Pathology institutes ($n=6$) in this canton in order to identify patients that have been diagnosed with esophageal eosinophilia between 1993 and 2013. We then performed a chart review in all adult and pediatric gastroenterology practices in order to identify EoE patients and to assess their clinical, endoscopic, and histologic characteristics.

Results: Out of 263 patients with esophageal eosinophilia, a total of 179 fulfilled the diagnostic criteria for EoE. Median diagnostic delay was 4 (IQR 1-9) years. No patient was diagnosed with EoE prior to 2003. Incidence of EoE increased from 0.16/100,000 inhabitants in 2004 to 6.3/100,000 inhabitants in 2013 ($p<0.001$). The cumulative EoE prevalence in 2013 was 24.1/100,000. The incidence in males was 2.8 times higher (95%-CI 2.01-3.88, $p<0.001$) when compared to that in females. The annual EoE incidence was 10.6 times higher (95%-CI 7.61-14.87, $p<0.001$) in the period from 2010-2013 when compared to that in the period from 1993-2009.

Conclusions: The incidence and cumulative prevalence of EoE in Canton of Vaud, Switzerland, has rapidly increased in the past ten years.

Eosinophilic Esophagitis: Diagnostic Accuracy of Symptoms in predicting Endoscopic and Histologic Remission

Ekaterina Safronieva¹, Michael Coslovsky¹, Claudia Kuehni¹, Marcel Zwahlen¹, Sven Trelle¹, Nadine Haas¹, Alex Straumann², Alain Schoepfer³; International EEsAI study group.
¹ISPM University of Bern, ²Swiss EoE Center Olten, ³CHUV, Lausanne

Background: We aimed to evaluate the controversially discussed relationship between clinical severity, and endoscopic and histologic activity in adult patients with eosinophilic esophagitis (EoE).

Methods: Patients completed the validated EEsAI Patient Reported Outcomes (PRO) instrument (range 0-100 points) and then underwent endoscopy with esophageal biopsy sampling. Endoscopic remission was defined as follows: 1) absence of white exudates; 2) absence of moderate and severe rings; 3) absence of strictures; 4) furrows and edema could be present but not in combination. Histologic remission was defined as <20 eosinophils/mm². ROC curves were calculated to determine the best EEsAI PRO cutoff values to predict remission.

Results: A total of 269 EoE patients was recruited (67% males), of these 111 (41.3%) had a PRO score <20 points, 79 (29.7%) were in endoscopic and 75 (27.9%) were in histologic remission. The AUC was 0.6719, 0.6007, and 0.6797, to predict endoscopic, histologic, and combined endoscopic and histologic remission, respectively. The best overall accuracy to predict endoscopic, histologic, and combined endoscopic and histologic remission was found for an EEsAI PRO score of 21 points (accuracy 65.2%), 21 points (accuracy 62.1%), and 15 points (accuracy 67.7%), respectively.

Conclusions: The EEsAI PRO has a sufficient accuracy in predicting endoscopic and histologic remission. In addition to assessing PRO measures, assessment of endoscopic and histologic alterations remains to be an important element in the judgment of overall disease activity.

G7

30 days global mortality in patients undergoing endoscopy or liver biopsy in an academic center : A 2 years survey.

Laurent Bochatay, Nicolas Goossens, Emiliano Giostra, Jean-Louis Frossard. Gastroenterology and Hepatology Service, Geneva University Hospital, Geneva, Switzerland. **Background :** Global mortality associated with invasive procedures in a gastrointestinal unit is not clearly documented in the literature. Over 11'000

procedures have been performed between 2011 and 2012 in Geneva University Hospital. The aim of this study is to describe the 30 days mortality after an invasive procedure in our center.

Methods: All patients who had either a gastroscopy, colonoscopy, EUS, ERCP, PEG or liver biopsy were included. Gastroscopy was taken as the reference procedure for mortality comparison. Univariate and multivariate analysis for age, sex, type of procedure and physician experience were performed.

Results: 11'241 procedures were performed during the observation period. Among them, 291 were followed by death within 30 days (2.5%). Age was significantly higher in the 30 days mortality group (71.5 vs 60.9, p<0.001). In multivariate analysis, age over 60 (OR 3.4, p<0.01) and PEG procedure were associated with 30 days mortality. In contrast, physician experience (OR 0.71, p<0.001), echoendoscopy (OR 0.55, p=0.001), colonoscopy (OR 0.56, p<0.001) and percutaneous liver biopsy (OR 0.56, p=0.02) were inversely associated with 30 days mortality. **Conclusions :** Age over 60 years old and PEG procedure are associated with 30 days mortality. By contrast, colonoscopy, echoendoscopy and percutaneous liver biopsy are inversely associated with 30 days mortality. Although this study does not depict individual risk factor associated with 30 days mortality, our attention should be focused on older patient (>60 years old) and on patient undergoing PEG procedure.

G9

Eosinophilic Esophagitis: relationship of quality of life with clinical, endoscopic, and histologic activity

Ekaterina Safronieva¹, Michael Coslovsky¹, Claudia Kuehni¹, Marcel Zwahlen¹, Nadine Haas¹, Alex Straumann², Alain Schoepfer³; International EEsAI study group. ¹ISPM University of Bern, ²Swiss EoE Center Olten, ³CHUV, Lausanne

Background: Knowledge about determinants of quality of life (QoL) in eosinophilic esophagitis (EoE) patients helps to identify patients at risk of experiencing poor QoL and to tailor therapeutic interventions accordingly. We evaluated the impact of symptom severity, endoscopic and histologic activity on EoE-specific QoL in adult EoE patients.

Methods: Ninety-eight adult EoE patients were prospectively included (64% male, median age 39 years). Patients completed two validated instruments to assess EoE-specific QoL (EoO-QoL-A) and symptom severity (adult EoE activity index patient-reported outcome) and then underwent EGD with biopsy sampling. Physicians reported standardized information on EoE-associated endoscopic and histologic alterations. The Spearman's rank correlation coefficient was calculated to determine the relationship between QoL and symptom severity. Linear regression and analysis of variance was used to quantify the extent to which variations in severity of EoE symptoms, endoscopic and histologic findings explain variations in QoL.

Results: QoL strongly correlated with symptom severity ($r=0.610$, $P<0.001$). While the variation in severity of symptoms, endoscopic and histologic findings alone explained 38%, 35%, and 22% of the variability in EoE-related QoL, respectively, these together explained 60% of variation. Symptom severity explained 18–35% of the variation in each of the five QoL subscale scores.

Conclusions: EoE symptom severity and biologic disease activity determine QoL in adult EoE patients. Therefore, reduction of both EoE symptoms as well as biologic disease activity is essential for improvement of QoL in adult EoE patients.

G8

Health questionnaire effectiveness for planning the type of sedation for colonoscopy: a pilot study

Rocco Grignoli¹, Philippe Bichard¹, Bernard Walder², Jean-Louis Frossard¹. ¹ Service de Gastroentérologie HUG ² Service d'anesthésie HUG

Introduction Since 2005, propofol is the most frequently used drug for sedation in patients undergoing endoscopy. In accordance with the guidelines of the SGGSSG, patients who may benefit from an endoscopy with Propofol sedation by the operator, are those with low anesthetic risk (ASA1 or 2). In our practice, the medical history of near half of patients are unknown at the time of programming endoscopy. In order to assess the pre-endoscopy risk, we developed a simple questionnaire completed by the patient that allows to determine the type of sedation. **Patients and method** Outpatients scheduled for colonoscopy between 1 September and 20 December 2013 fulfilled a standardized questionnaire evaluated by a gastroenterologist for the use of Propofol: if all responses were negative, endoscopy was scheduled with sedation by the operator. In case of > 1 positive answers, the patient was referred to the anesthesia consultation. Retrospectively the ASA score was calculated and the incidence of complications was recorded. **Results** 100 ambulatory patients underwent colonoscopy. 94 correctly answered the questionnaire. 37 patients were referred to the anesthesia consultation due to one or more positive answers. In the absence of risk factor based on their responses to the questionnaire, 57 patients had directly access to colonoscopy with Propofol administered by the operator. Characteristics of these 57 patients: mean age 54.5 years (23-86), 61.2% men. Retrospectively, we could calculate the ASA score of 49 patients. 16 (32.7%) were ASA 1, 33 (67.3%) were ASA 2. No patient with sedation by the gastroenterologist had an ASA score > 2. 9 patients (15.8%) had a fair or poor tolerance (8 for unmanageable pain, 1 for vomiting). **Conclusion** The results of this preliminary study demonstrates the ability of our questionnaire to identify patients at very or low anesthetic risk, allowing access directly to the endoscopy without anesthetic consultation and therefore reduce the waiting times.

G10

Multi-Country, Cross-Sectional Study to Determine Patient-Specific and General Beliefs Toward Medication and Their Treatment Adherence to Selected Systemic Therapies in 6 Chronic Immune-Mediated Inflammatory Diseases (ALIGN)

Pierre Michetti¹, John Weinman², Ulrich Mrowietz³, Josef S. Smolen⁴, Dieter Schremmer⁵, Namita Tundis⁶, Nicole Selenka-Gebauer⁷

¹La Source-Beaulieu, Lausanne, Switzerland; ²King's College, London, UK; ³University Schleswig-Holstein, Kiel, Germany; ⁴Medical University of Vienna, Vienna, Austria; ⁵GKM, Munich, Germany; ⁶AbbVie Inc., North Chicago, IL, USA; ⁷AbbVie Inc., Zürich, Switzerland

INTRODUCTION: Adherence to therapy is critical to achieve and sustain optimal outcomes in patients (pts) with immune-mediated inflammatory disease (IMID). Pts' beliefs about the necessity of treatment and potential adverse effects could strongly influence adherence.

AIMS & METHODS: ALIGN was a multi-country cross-sectional study exploring pts' beliefs, concerns, attitudes and adherence toward TNF inhibitors (TNFi) and conventional therapies used alone or in combination across multiple IMIDs. Adults age ≥ 18 y with rheumatoid arthritis (RA), ankylosing spondylitis (AS), psoriatic arthritis (PsA), Crohn's disease (CD), ulcerative colitis (UC) or psoriasis (PsO), receiving conventional therapy and/or disease-modifying antirheumatic drugs (including TNFi), were recruited. Pts completed validated questionnaires, e.g., the Beliefs about Medicines Questionnaire (BMQ) and short Morisky Medication Adherence Scale (MMAS-4). BMQ scores, MMAS-4 scores and pts' attitudes toward their medications are presented.

RESULTS: 7197 pts in 33 countries met eligibility criteria. Pts had RA (27.5%), AS (11.3%), PsA (8.9%), CD (17.3%), UC (8.8%) or PsO (26.2%). Mean age was 47.5 y (range, CD=38.0; RA=54.8). Mean disease duration was 11.7 y (range, UC=8.1; PsO=18.7). The largest proportion of pts received conventional therapies (40.3%), followed by TNFi mono- (32.0%) and combination therapy (27.7%). An attitudinal analysis of BMQ necessity and concern scores revealed that most pts were either "accepting" (high necessity/low concern) or "ambivalent" (high necessity/high concern) toward their medication irrespective of disease or treatment. Adherence across diseases was generally higher in pts receiving TNFi with or without conventional therapy (range of mean MMAS-4 scores, 3.4-3.7; 0=low, 2=medium, 4=high adherence), vs pts receiving conventional mono- (2.6-3.3) or combination therapy (2.8-3.4). Across treatments, high adherence according to MMAS-4 was consistently lower for "ambivalent" pts (46.1%-69.0%) vs "accepting" pts (55.8%-77.6%) according to combined BMQ scores (Table).

Table. N (%) of Patients Accepting and Ambivalent Toward Their Medications Who Were Adherent.

Monotherapy, n (%)				TNFi + Conventional Combination Therapy, n (%)			
Conventional		TNFi		Conventional		TNFi	
Accepting (n=1,347) (n=1,151)	Ambivalent (n=1,251)	Accepting (n=1,251) (n=834)	Ambivalent (n=834)	Accepting (n=900) (n=739)	Ambivalent (n=739) (n=1031)	Accepting (n=786) (n=786)	Ambivalent (n=786)
751 (55.8)	531 (46.1)	919 (73.5)	504 (60.4)	550 (61.1)	374 (50.6)	800 (77.6)	542 (69.0)
p<0.0001		p<0.0001		p<0.0001		p<0.0001	

CONCLUSION: Compared with "accepting" pts, "ambivalent" pts appeared to be less often highly adherent (MMAS-4 score=4), which could negatively affect treatment efficacy. The high percentage of "ambivalent" pts reveals the need to better explore concerns about medication and address erroneous beliefs regarding benefit-risk of treatments.

The Cytokine IL-19 is Expressed by Macrophages

Anna Steinert, Andrew Macpherson and Jan Niess

Department of Clinical Research, University Clinic for Visceral Surgery and Medicine, University of Bern

Background: Macrophages are rather an undefined cell population in the gut. We hypothesized that myeloid cells in the gut can be identified by the expression of IL-19, a member of the IL-20 cytokine family.

Methods: To identify the source of IL-19, a reporter mouse line was constructed in which the IL-19 promoter drives the expression of the red fluorescent protein tdTomato. Expression levels of IL-19 and of the type I IL 20 receptor (formed by the IL-20 receptor alpha (IL-20RA) and beta (IL 20RB) subunits) were determined in mice with different hygiene statuses.

Results: Bone marrow-derived macrophages expressed significantly more IL-19 than dendritic cells. IL-19 expression was upregulated after stimulation with Toll-like receptor ligands. The addition of IL-19 to macrophages prestimulated with LPS decreased secretion of the proinflammatory cytokine IL-6. IL-19 expression was only found in mesenteric lymph nodes, but not in other organs tested. IL-19 is produced by phagocytes isolated from the large intestine of mice with colitis. IL-20RB was expressed in all investigated organs with the highest expression in the skin. IL-20RA was expressed in the epidermis, the stomach and the proximal colon. IL-20RA and IL-20RB expression depend on the hygiene status because highest expression was observed in germ-free animals.

Conclusions: Macrophage derived IL-19 might act on mucosal sites where the type I IL-20 receptor is expressed.

G11

Title: ¹³C-labeled bacteria metabolomics reveals the identity of bacterial metabolites in the host and the host metabolic responses.

Authors: Yasuhiro Uchimura [1], Tobias Fuhrer [2], Uwe Sauer [2], Kathy D. McCoy [1] and Andrew J. Macpherson [1]

Affiliations:

[1] Maurice Müller Laboratories, Department of Clinical Research (DKF), Clinic for Visceral Surgery and Medicine (UVCMD), University of Bern, Bern

[2] Department of Biology, Institute of Molecular Systems Biology, ETH, Zurich

Background: A combination of physical barriers and innate and adaptive immunity prevents penetration of live bacteria from the intestines to systemic sites within our bodies. Nevertheless, we have found that bacterial products can propagate throughout the host organism.

Methods: In order to identify bacteria-derived metabolites *in vivo*, we colonized germ-free mice with bacteria that had been fully labeled with carbon-13 and then traced ¹³C-metabolites in various host sites by mass-spectrometry. In addition, the host metabolic response to bacterial colonization was also analyzed through mass-spectrometry.

Results: We identified bacterial metabolites in more than 25 different host tissues. The pattern of propagation of individual bacterial metabolites depended on their molecular features, such as hydrophobicity. We also found prominent host metabolic responses in the peritoneal cavity and portal organs such as liver and spleen as compared to other sites.

Conclusion: Our study demonstrates that ¹³C-bacteria metabolomics is a powerful technique to address metabolic relationship between gut microbes and the host.

G12

Repertoire analysis of hygiene-induced IgE in germ-free mice

Sandra C. Rupp; Julien P. Limenitakis; Markus B. Geuking; Andrew J. Macpherson; Kathy D. McCoy

Department for Clinical Research, University of Bern, Bern

Background: Microbial colonization of the gastrointestinal tract educates and shapes the development of the immune system. Accordingly germ-free (GF) mice show an immature immune system with reduced serum immunoglobulin titers of all isotypes except for IgE. This hygiene-induced IgE is T cell and IL-4 dependent and can only be prevented by microbial colonization early in life. Since alterations of the microbiota have been linked to autoimmune and allergic diseases and IgE is important in both, we investigated the repertoire of IgE in GF mice using a high throughput sequencing approach.

Methods: RNA was isolated from the mesenteric lymph nodes of GF mice aged 10, 20 and 40 weeks. First strand cDNA synthesis was achieved using the SMARTer 5'RACE kit (clontech) followed by a 5'RACE and amplicon PCR of the VDJ region. The PCR product was gel purified for 650-800bp length. After the addition of barcodes, samples were sequenced on the Illumina MiSeq platform with 300bp paired end reads. Data analysis was executed with IMGT Hi V-Quest or IgBlast. Graphical output was created with IgGalaxy, Excel and R.

Results: The CDR3 length in GF mice suggests an oligoclonal IgE repertoire that manifests as the mice mature. Further analysis will reveal if there is selective usage of V, D, or J gene segments or evidence of somatic hypermutation, indicative of a germinal centre reaction.

Conclusion: The oligoclonal IgE repertoire in GF mice suggests that this response is antigen-driven, which may indicate self-reactivity. The early exposure to a diverse microbiota is consequently required for a basal regulation of the immune system.

G13

Maternal microbiota educates the immune system of the offspring through natural antibodies

Gomez de Agüero, Mercedes¹; Ganal, Stephanie C.¹; Rupp, Sandra; Steinert, Anna; McCoy Kathy. D.¹ & Macpherson, Andrew. J.¹

Division of Gastroenterology, Department of Clinical Research, University of Bern, 3010 Bern, Switzerland;
*These authors contributed equally to this work

Background: Mammalian embryonic immune system development occurs in the absence of live bacterial exposure, although it is possible that the developing fetus may be exposed to microbial products from the maternal microbiota. However, the potential impact of the maternal microbiota on immune development and health of the offspring is not clearly understood.

Methods: Germ-free (GF) mice were transiently colonised with bacteria during the pregnancy by oral gavage with an auxotrophic *Escherichia coli* strain HA107. Innate immune cells of the offspring were analysed by FACS at different time points.

Results: We have found that the offspring from mothers exposed to intestinal microbes only during pregnancy showed an increase in intestinal NKp46⁺Roryt⁺ innate lymphoid cells and F4/80⁺CD11c⁺ mononuclear cells. Moreover, the expression of genes involved in microbial adaptation was imprinted by maternal microbiota. The offspring from microbial treated mothers were protected against microbiota challenge preventing bacterial translocation to mesenteric lymph nodes and modulating gene expression. In the absence of natural antibodies and B cells, maternal microbiota failed to shape the innate immune system in the offspring resulting in increased bacterial translocation following challenge. Furthermore, maternal antibodies required loading with maternal bacterial products to control the development and the efficiency of the innate immune system of the offspring.

Conclusions: Our results reveal the tremendous role played by maternal microbiota and natural antibodies in setting the baseline of the innate immune system in the offspring.

G15

Buried Bumper Syndrome: a case-series of a new technique for the endoscopic management of an old complication. Paulo J. Castro Soares¹, Jean-Louis Frossard¹, Philippe Bichard^{1,2}. Gastroenterology and Hepatology Department, Geneva University Hospital, Geneva.

Background: Buried bumper syndrome (BBS) is rare complication of percutaneous endoscopic gastrostomy (PEG) and percutaneous endoscopic jejunostomy (PEJ) due to excessive external traction that leads to erosion of the internal bumper with its inclusion into the visceral and parietal walls. Since it was first described in 1988 by Shallman et al. several different techniques for endoscopic treatment have been proposed. Among the most commonly used we can include the needle-knife incision of the gastric wall, the push-and-pull "T" technique, the extraction using the tapered tip of a new PEG tube and the dilation of the opening using a external placed balloon catheter. **Methods:** From July 2013 to May 2015 four cases of BBS were diagnosed in our tertiary center. Three of the cases were complications of a PEG and one a complication of a PEJ. All these cases were treated with the following technique: we cut the external end of the tube at 10 cm from the abdominal wall and insert a 0.025-inch VisiGlide® guide wire through the feeding tube lumen into the stomach. The guide wire is recuperated with a 13mm snare. We then advance a Hurricane™RX Biliary Ballon Dilation Catheter with 6mm of outside diameter and 4cm in length and inflate it to maximum manufacturer recommended pressure with an Encore™26 Inflator. Traction of the guide wire and balloon catheter allows the extraction of the bumper and sectioned tube into the stomach. A new balloon-type feeding tube is then inserted. **Results:** The average time to diagnosis of BBS after the PEG or PEJ procedure was 21.5 months. All of the cases were successfully treated and immediate placement of a new balloon-type feeding tube through the same PEG/PEJ was performed with no complications. All the PEG/PEJ are still functional and in use. **Conclusions:** Our technique as shown to be effective and safe and can be added to the arsenal of endoscopic procedures available to treat BBS of PEG and PEJ.

G17

Infliximab treatment outcome in steroiddependent/refractory Crohns Disease patients: a single center report

G16

T. Kaymak, F. Moriconi, C. Beglinger, P. Hruz

Background:

The anti-TNF α antibody Infliximab (IFX) is an efficient therapy in patients with Crohns Disease (CD) who have a steroid-dependent, steroid-refractory or a complicated (e.g. fistula) disease course. Many patients receive a long-term treatment. Recommendations with respect to discontinuation of the IFX therapy are not well established. As IFX is an expensive therapy with potential side effects the outcome of stopping an IFX is an important question. The aim of the current study was 1) to analyze long-term treatment outcome in CD patients and 2) to evaluate withdrawal of IFX in CD patients in remission.

Methods:

In a single center retrospective analysis we report the outcome of CD patients treated with IFX between 2008 and 2015 (n=109) at the University Clinic of Basel. The indication for IFX treatment was steroid-dependent, steroid-refractory or a complicated (e.g. fistula) disease course. Every patient was initially treated with a regular induction scheme (IFX 5mg/kg body weight at 0, 2, 6 weeks). Clinical remission was defined as Calprotectin <150 μ g/g stool and absence of clinical symptoms. "Discontinuation of treatment" was defined as a permanent stop of IFX therapy after 2 years of clinical remission with a bland endoscopic and histological examination before stopping the treatment.

Results:

After the induction treatment with IFX 41/109 (37.6%) patients achieved remission. Of these patients, 13/41 (31.7%) were on long-term remission and discontinued from IFX during disease course, 18/41 (43.9%) remained on long-term therapy with IFX, 5/41 (12.2%) needed a change to a different TNF α inhibitor and 5/41 (12.2%) were lost to follow-up.

In 68/109 (62.3%) patients a complete clinical response could not be achieved after induction treatment with IFX. In 3/68 (4.4%) patients an immediate change to another biological treatment was necessary. A permanent IFX therapy without any changes was observed in 16/68 (23.5%) patients. However, in 49/68 (72%) an IFX dose increase and/or interval reduction was necessary. An interval reduction of IFX therapy led to a remission in 13/33 (39.4%) whereas 20/33 (60.6%) remained in a chronic active state with minimal or absent clinical symptoms. In the group "dose increase and interval reduction", only 1/13 (7.7%) achieved a remission.

In total, IFX therapy could be discontinued in 15/109 (13.8%) patients, including 2 patients that did not initially achieve remission after the induction therapy. Relapse was observed in 11/15 (73.3%) patients; 10/11 (90.9%) patients responded to re-induction therapy with IFX.

Conclusion:

Our single center analysis demonstrates that only a minority of patients started on IFX therapy due to a steroid dependent, steroid-refractory or a complicated (e.g. fistula) disease course can be discontinued from the TNF α inhibitor treatment. Withdrawal of IFX therapy led to a relapse in 73.3% of our patients in remission with good response to re-induction with IFX.

Mucosal consequences of systemic antimicrobial CD4 $^{+}$ T cell reactivity

G18

Cheong KC Kwong Chung, Kathy D McCoy, Andrew J Macpherson, and Markus B Geuking

Mucosal Immunology Lab, Department of Clinical Research, Universitätsklinik für Viszerale Chirurgie und Medizin Inselspital, University of Bern, Murtenstrasse 35, CH-3010 Bern, Switzerland.

Background: Healthy host-microbial mutualism relies on compartmentalization and proper regulation of systemic and mucosal immune responses. Despite this, the systemic immune system is frequently exposed to some levels bacteraemias, which can trigger systemic antimicrobial immune reactivity including CD4 $^{+}$ T cells. This can occur for example when immune compartmentalization is compromised, in the presence of deficient innate immunity, or use of nonsteroidal anti-inflammatory drugs (NSAID). Importantly, systemic hyper reactive CD4 $^{+}$ T cells might be involved in the pathogenicity of inflammatory bowel diseases (IBD).

Methods: We generated a genetically modified *Escherichia coli* strain expressing a defined T helper epitope to study antigen-specific antimicrobial CD4 $^{+}$ T cells.

Results: We found that the bacterial load threshold required for the induction of systemic antimicrobial CD4 $^{+}$ T cell proliferation was very high and not easily reached under physiological conditions. However, when intestinal barrier function was compromised by induced damage to the intestinal epithelium, the presence of systemic anti-microbial CD4 $^{+}$ T cells resulted in dramatically increased levels of bacterial translocation.

Conclusions: Systemic antimicrobial CD4 $^{+}$ T cell reactivity might impact adversely on the mucosa under conditions of reduced barrier function.

IL-22-induced antimicrobial peptides are key determinants of vaccine-induced protection against *Helicobacter* in mice

Mati Moyat,¹ Hanifa Bouzourene,² Wenjun Ouyang,³ Jean-Christophe Renaud,⁴ and Dominique Velin¹

¹Service of Gastroenterology and Hepatology, CHUV, Lausanne.

²UNISciences, UNIL, UniLabs, Lausanne. ³Department of Immunology, Genentech, South San Francisco, USA. ⁴Ludwig Institute for Cancer Research, Brussels Branch, Belgium.

BACKGROUND: Despite the proven ability of immunization to reduce *Helicobacter* infection in mice, the precise mechanism of protection has remained elusive. In this study, we evaluated the role of IL-22 in vaccine-induced reduction of *Helicobacter* infection.

METHODS: Vaccine-induced reduction of *Helicobacter* infection was compared in vaccinated mice with or without functional IL-22.

RESULTS: Gastric IL-22 levels were increased in mice immunized and challenged with *H. felis*, as compared to controls (18±11 and 8±8 ng/mg for IL-22 protein, p<0.05, respectively). FACS showed that a peak of CD4⁺IL-22⁺ T cells infiltrating the gastric mucosa occurred in immunized mice in contrast to control mice (2.06±0.60% and 0.79±0.26%, p<0.01, respectively). Injection of anti-IL-22 to immunized mice significantly prevented the reduction of *Helicobacter* infection (2 out of 9 mice (anti-IL-22) versus 8 out of 9 (control) mice reduced *Helicobacter* infection p<0.001). Finally, AMPs extracted from stomachs of vaccinated mice or mice injected with IL-22Fc, but not from the stomachs of non-immunized or immunized mice injected with anti-IL-22 efficiently killed *Helicobacter* in vitro.

CONCLUSION: These results demonstrate that IL-22 plays a critical role in vaccine-induced reduction of *Helicobacter* infection, by promoting the expression of AMPs capable of killing *Helicobacter*.

G19

Microbiota-mediated fine-tuning of the threshold of intestinal inflammasome activation in host-microbial mutualism.

Francesca Ronchi, Andrew Macpherson* and Kathy McCoy*

Maurice Müller Laboratories (DKF), Universitätsklinik für Viszeral Chirurgie und Medizin Inselspital, University of Bern, Murtenstrasse 35, CH-3010 Bern Switzerland

these authors equally contributed to the work

Background: The inflammasome is a complex of proteins that controls the activity of caspase-1, pro-IL-1beta and pro-IL-18. It acts in inflammatory processes and in pyroptosis. The lower intestine is densely populated by a community of commensal bacteria that, under healthy conditions, are beneficial to the host. Some evidence suggests that the gut microbiota influences regulation of the inflammasome. Components of inflammasomes have been shown to have a protective function against development on experimental colitis, dependent on IL-18 production. However the precise mechanisms and the role of the inflammasome in maintaining a healthy host-microbial mutualism in the absence of pathogens remains unknown.

Methods: To address this question, we have performed *in vivo* experiments colonising germ-free (GF) mice with limited and defined or diverse but undefined commensal bacteria to investigate how the inflammasome components, particular within intestinal epithelial cells (IECs), are regulated under different hygiene conditions.

Results: We have established that gene expression of the inflammasome components NLRC4, NLRP3, NLRP6, NLRP12, caspase-1, ASC and IL-18 do not differ between GF and colonised conditions under steady-state, even when colonised with a highly diverse intestinal microbiota. Under steady state, we observed high expression of NLRs and IL-18 in the gut, even in absence of microbiota. As expected, induction of IL-18 was observed following infection with the pathogen *Citrobacter rodentium*. In contrast, IL-18 was induced transiently upon colonisation of germ-free mice. We are now investigating the dynamics of microbial colonisation at early time points to determine whether this early induction of IL-18 is required for homeostasis.

Conclusion: Our data suggest that in contrast to pathogenic bacteria commensal bacteria do not dramatically induce expression or activation of the inflammasome in IEC. Nevertheless, modulation of IL-18 may occur during the first phase of colonisation. We are now investigating the impact of steady-state expression of the inflammasome under different hygiene conditions using mice that are deficient for protein members or products of the inflammasome complex, such as ASC and IL-18 gene-deficient mice.

G21

Success and complications of an intraductal fully covered metal stent to treat anastomotic biliary strictures (ABS) after orthotopic liver transplantation (OLT)

G20

A rare case of a pigmented neuroendocrine tumour of the stomach in the setting of autoimmune gastritis

G22

Henrik Csaba Horvath¹, Eva Diamantis-Karamitopoulou²

¹University Clinic of Visceral Surgery and Medicine Inselspital, Bern

²Division of Clinical Pathology, Institute of Pathology, University of Bern

Background: Neuroendocrine tumours of the gastrointestinal tract (GI-NETs) account for 2% of all GI-Tract neoplasms. Individual GI-NETs may exhibit widely different clinical courses, reflecting their underlying heterogeneity at the genetic, molecular and cellular level that influences their biological behaviour.

Results: We report an extremely rare case of a "black" pigmented gastric tumour of 3mm in size found in a 68 year-old male patient with newly diagnosed chronic atrophic gastritis and known chronic alcohol abuse. Histologic examination revealed a NET with trabecular growth and monomorphic cells which showed immunopositivity for pancytokeratin, chromogranin and synaptophysin and were negative for S100, Melan A and HMB45. The MIB1 Index was <2%, compatible with a NET G1. Perls' stain was negative, while electron microscopy revealed melanosomes in the pigmented tumour cells compatible with presence of melanin pigment.

Conclusion: To our knowledge, this is the first reported case of pigmented gastric NET. Pigmented NETs may represent a diagnostic challenge due to their endoscopic similarity to melanoma. Since epidemiological data show an increasing incidence of gastrointestinal neuroendocrine tumours, endoscopists may be faced with pigmented NETs more frequently in the future.

Patrick Aepli¹, Rhys Vaughan², Luke Hourigan³, Arthur Kaffes¹
 1 Gastroenterology and Hepatology, Royal Prince Alfred Hospital, Sydney, New South Wales, Australia
 2 Gastroenterology and Liver Transplantation, Austin Health, Melbourne, Victoria, Australia
 3 Gastroenterology, Princess Alexandra Hospital, Brisbane, Queensland, Australia

Background: Metal stents for benign biliary strictures are gaining acceptance with many published series. Traditional metal stent designs seem to have poor durability in ABS after OLT. Novel intra-ductal stents show promise in these strictures. **Methods:** A retrospective multi-centre Australian study of ABS after OLT treated with a novel intraductal stent. Records from 3 liver transplant centre databases were reviewed.

Results: Totally 36 removable fully covered self expanding metal stents (RCSEMS) were inserted in 31 cases to treat an ABS after OLT. The mean age of the patients was 56 years. Treatment with the RCSEMS was performed with an average treatment time of 3.8 months. Stricture resolution was achieved in 93.5 %. All attempted stents removals were successful without any difficulty. Follow-up showed 7 cases of ABS recurrence (24.1 %) and all were treated successfully with repeat ERCP and stenting (some metal and some plastic). The mean time of ABS presentation after OLT was 20.3 months. Complications were reported in 6.5 % (one patient had mild cholangitis on stent removal, one patient required re-stenting because of cholangitis due to a secondary stricture). Finally it was pleasing, that no cases of stent migration were seen.

Conclusions: This novel intra-ductal fully covered metal stent has a high clinical success and low complication rate, in particular there were no cases of stent migration. As a result this stent type is preferred to traditional metal stents for treating ABS after OLT.

Alcohol Consumption and Impact on Gastrointestinal Symptoms among Swiss IBD Patients

Authors:

Felix Brunner¹, Nicolas Fournier², Roland von Känel³, Stefan Begré⁴, Carole Clair⁵, Andrew Macpherson¹, Pascal Juillerat¹
¹Gastroenterology, Inselspital, Bern; ²Social and Preventive Medicine, CHUV, Lausanne; ³Klinik Barmelweid, Barmelweid; ⁴Klinik Hohenegg, Meilen; ⁵PMU, CHUV, Lausanne.

Introduction

Little is known about alcohol drinking behavior and its influence on gastrointestinal symptoms among patients with inflammatory bowel disease (IBD).

Aims & Methods

A questionnaire was sent to patients randomly selected from 3 different groups according to reported frequency of alcohol consumption at enrolment to the Swiss IBD cohort. It was designed to assess the average alcohol intake per day, type of drinking, number of days drinking per week, influence of alcohol on gastrointestinal symptoms and smoking. Other data on demographics and disease characteristics were gathered from the most recent follow up visit.

Results

383 of 537 (72%) questionnaires were answered. The rate of abstainers was similar to the Swiss population (12%). Among drinkers, 52% drank up to 1x weekly, 29% on 2-3 and 19% on 4-7 days per week. The amount of alcohol per occasion was 0-20g in 70%, 20-40g in 24% and 40-60g in 5% and similar for both diseases. After being diagnosed with IBD, 61% did not change, 29% reduced, 6% quit or 4% increased alcohol consumption. In both, CD and UC, 30% of patients reported worsening of diarrhea, 15% worsening of abdominal pain and bloating after drinking alcohol. The type of preferred beverage did not significantly influence gastrointestinal symptoms.

Conclusion

Drinking behavior in Swiss IBD patients is similar to the general population. A third of patients reduced or quit drinking after being diagnosed with IBD. Worsening of diarrhea, abdominal pain and bloating after drinking alcohol was reported in 15-30% by both UC and CD patients, independently of the preferred drinks.

Predictors of success for double balloon assisted endoscopic retrograde cholangiopancreatography (DBE-ERCP) in patients with Roux-en-Y anastomosis

Patrick Aepli¹, Ken Liu¹, Arthur Kaffes¹

¹ Gastroenterology and Hepatology, Royal Prince Alfred Hospital, Sydney, New South Wales, Australia

Background: Performing ERCP in patients with Roux-en-Y anastomosis (REYA) is a major challenge to endoscopists since the biliary anastomosis is not reachable with a standard duodenoscope. The use of DBE-ERCP has been shown to permit endoscopic intervention in this setting.

Methods: We retrospectively studied all patients with REYA who had a DBE-ERCP between 2009 and 2015 at our liver transplant centre.

Results: In totally 47 patients we performed 76 DBE-ERCPs. Patients had REYA for different indications (mostly orthotopic liver transplantation, OLT). DBE-ERCP was mainly performed for biliary obstruction (often with concomitant cholangitis). Overall, DBE-ERCP was technically successful in 65.8 %. The success rate was 84.0 % in patients with REYA from non-transplant surgery performed in adulthood, 57.8 % in patients post OLT and only 20.0 % in biliary atresia patients. Therapeutic interventions (e.g. biliary stenting, stent removal, dilatation) were successful in 72.1 %. Complications occurred in only 2 cases, both developed cholangitis which resolved with conservative management.

Conclusion: The initial indication for REYA appears to be predictive for successful DBE-ERCP. The procedure is most successful in patients after non-transplant adult surgery (previous bile duct injury, gastrectomy, pancreaticoduodenectomy), intermediate successful in patients post OLT and least successful in patients with biliary atresia. Patients predicted to have low success rates should be initially referred for alternative procedures (PTCD or EUS-guided biliary access).

G23

Rare Double -„Fish Eye“- Papilla Vateri in Main-Duct-Intraductal Papillary Mucinous Neoplasia (IPMN): Curative Surgical Resection by Total Duodeno-Pankreatectomy.

¹Reiner Wiest, ¹Andrew Macpherson, ²Andreas Frenzer,

²Christian Ruis, ¹Thomas Malinka, ¹Beat Gloor; ¹University Hospital Visceral Surgery and Medicine, Inselspital Bern, Freiburgstrasse, 3010 Bern; ²Department Gastroenterology, Spital Thun, Krankenhaus-strasse 12, 3600 Thun

85 y old male patient presented himself with painless jaundice and loss of body weight (20 kg/6 months). Abdominal imaging revealed an excessively dilated biliary tree and pancreatic duct. ERCP was performed showing a “double-fish-eye” papilla – as a rare finding in the literature - with extrusion of massive viscous mucus from biliary as well as pancreatic orificium. Repeated endoscopic interventions via ERC including cholangioscopy failed to relieve biliary obstruction due to inability to wash-out the mucus. After percutaneous transhepatic cholangiographic drainage the patient finally was operated despite age, comorbidities and high nutritional risk score. A total duodeno-pancreatectomy was performed showing an invasive colloid pancreatic cancer derived from a “main-duct-type” in intraductal papillary mucinous neoplasia = IPMN of intestinal typ: TNM pT2(m) pN0 (0/32) L0 V0 Pn0 R0. The patient is doing fine underlying the safety of this procedure even under these high-risk conditions.

G24

Swiss Snapshot Diverticulitis Trial: Current Practice of in-hospital Management of Acute Colonic Diverticulitis - a Nation's Experience

Seraina K Faes, Alexandre P Cuerel and Dieter Hahnloser for the Swiss Diverticulitis Study Group.

Department of Visceral Surgery, University Hospital Lausanne

Background

To evaluate the current practice of in-hospital management of acute colonic diverticulitis in Switzerland.

Methods

Prospective snapshot research during three months (and one month follow-up) of all patients hospitalized for acute colonic diverticulitis.

Results

Over a time period of three months, 74 participating hospitals treated 783 patients with acute colonic diverticulitis. 510/783 (65.1%) patients were diagnosed with uncomplicated diverticulitis using CT scans in 94.5%. Management was conservative in 99.8% with intravenous antibiotics in 98% and a failure rate of 1.4%. Patients with complicated diverticulitis (34.9%) were managed conservatively (67%), by percutaneous drainage (7%) or immediate surgery (26%). Failure of non-operative therapy was 15.3%. Laparoscopic surgery was performed in 46.2% with a conversion rate of 34.7%. Primary anastomosis was achieved in 49.1% (27.8% with protective loop ileostomy). Surgeons in training performed 31.5% of operations.

Conclusions

Conservative management of uncomplicated diverticulitis is very successful, questioning the need for hospitalisation. Hartmann's procedure in emergency surgery for complicated diverticulitis is still high.

G25

Swiss Snapshot Diverticulitis Trial: Influence of Hospital Category on the Management of Acute Colonic Diverticulitis

Seraina K Faes, Alexandre P Cuerel and Dieter Hahnloser for the Swiss Diverticulitis Study Group.

Department of Visceral Surgery, University Hospital Lausanne

Background

To evaluate the influence of hospital category on the management of acute colonic diverticulitis in Switzerland.

Methods

Swiss Snapshot Diverticulitis Trial: Prospective research during 3 months of all patients hospitalized for acute colonic diverticulitis or elective diverticular resection. Subanalysis for hospital category in acute diverticulitis.

Results

Of 151 Swiss hospitals, 74 (49%) contributed 783 acute diverticulitis cases. Hospitals of category A included most patients (34%). Category distribution varied significantly between uncomplicated and complicated diverticulitis ($p=0.001$; 23% complicated in B1 vs 54% in U). Diagnostic use of CT scan differed significantly (100% in U/Private (P), 83% in B1, $p=0.001$). Prescription of iv antibiotics ranged from 80% (A) to 90% (B3) ($p=0.03$). Conservative treatment was applied in 99.8% of uncomplicated and 74% of complicated cases and its failure rates varied for hospital category (uncomplicated: min 0% B3, max 8% P; complicated: min 8% B3 max 43% B1). Length of hospital stay was not differing upon category (uncomplicated: median 4, range 1-34, $p=0.57$; complicated: median 7, range 2-74, $p=0.088$). Execution of primary anastomosis was diverging (20% of U, 100% of P). Laparoscopy rate ranged from 32% B2 to 100% P. Conversions were performed in 27% A to 75% P of laparoscopies.

Conclusions

Management of acute colonic diverticulitis in Switzerland highly varied between different hospital categories. A nationwide guideline might uniform our treatment approaches.

G27

An acute jejunal variceal bleeding at choledochojejunostomy treated with surgically-assisted endoscopical rubber band ligation

Bernhard Friedli¹, Annette Wochner¹, Henrik Horvath¹, Stefan Tschopp², Franz Eigenmann², Daniel Candinas¹, Pascal Juillerat¹.

¹ University Clinic for Visceral Surgery and Medicine, Inselspital Bern.

² Department of Gastroenterology, Baden Hospital, Baden.

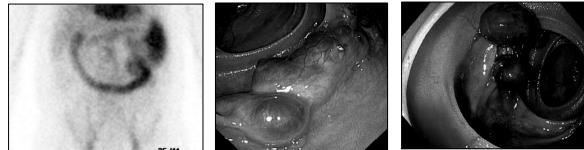
Introduction: Acute bleeding from jejunal varices located at the Roux-en-Y choledochojejunostomy is rare and its treatment is often complex due to the postoperative gastrointestinal anatomy. No guidelines have so far been established. In our literature review, we found 15 case reports which describes mostly endoscopical sclerotherapy, but also surgical ligation, revision of the anastomosis or radiological embolization and/or decompression of the portal venous hypertension (by TIPS or surgical shunt).

Case description: We report the case of a 74 year-old man admitted for acute gastrointestinal bleeding. He got liver transplantation due to HCV cirrhosis 8 years prior to this episode and had current portal hypertension secondary to chronic portal vein thrombosis. The bleeding source couldn't be initially identified despite repeated colonoscopy and gastroscopy, push enteroscopy, abdominal angiography and capsule endoscopy. A technetium 99m-labeled red blood cell scintigraphy could eventually locate it at the Roux-en-Y choledocho-jejunostomy. It was then decided to perform an surgically-assisted enteroscopy. After a laparotomy and enterotomy the scope was introduced at 30 cm from the anastomosis. Jejunal varices with "nipple sign" were found and treated using rubber band ligation.

Picture 1: Technetium 99m-labeled red blood cell scintigraphy

Picture 2: jejunal varices with "nipple sign"

Picture 3: after rubber band ligation



Results: Following the surgically-assisted endoscopical band ligation of the jejunal varices, no rebleeding occurred during a 6-month period. Due to the development of a severe portal hypertensive gastropathy with diffuse gastral bleeding a surgical mesocaval shunt was then performed. **Conclusions:** Bleeding from jejunal varices located at the site of choledochojejunostomy after OLTX is rare and challenging to detect and to access for treatment. Radio-labelled scintigraphy could help confirming the source of bleeding at the inaccessible anastomosis sign and a surgically-assisted push enteroscopy with band ligation therapy lead in our case to impressive results.

G29

Esophageal work-up prior to bariatric surgery: who and how to investigate

G28

Benjamin Heimgartner, Marcus Herzig, Radu Tutuian

University Clinic for Visceral Surgery and Medicine, University of Bern

Background: Gastroesophageal reflux disease (GERD) is common in obese patients and bariatric operations have different influence on GERD. We assessed the prevalence of symptoms and objective findings prior to surgery.

Methods: Work-up included quantification of symptoms, upper GI endoscopy and 24-h impedance-pH (imp-pH) monitoring off PPI. Imp-pH monitoring was classified as abnormal if either %time pH<4 was abnormal, total number of reflux episodes was elevated or symptom index (SI) was positive.

Results: Among 100 consecutive patients 54% reported heartburn and/or regurgitation, 71% had objective evidence of GERD (38% endoscopic lesions and 33% abnormal imp-pH results). Twenty-nine percent of patients with endoscopic lesion had normal imp-pH monitoring, 45% of patients with abnormal imp-pH recordings had erosions. Endoscopy was inferior to imp-pH in identifying GERD (sensitivity 54% vs. 85%, $p < 0.01$). Symptomatic and asymptomatic patients had similar prevalence of esophageal lesions (37% vs. 39%) and abnormal imp-pH findings (68% vs. 50%). Only 31% of patients with abnormal %time pH<4 had elevated number of reflux episodes suggesting that poor acid clearance is the main mechanism leading to GERD in obese patients.

Conclusion: Half of obese patients report typical GERD symptoms and >70% had evidence of GERD. Since symptoms don't predict objective findings, endoscopy and reflux monitoring should be part of the pre-bariatric surgery work-up.

Endoscopic recanalization of a post-radiogenic complete esophageal occlusion by combined anterograde / retrograde esophagoscopy

G30

Thomas Kratt^{1,2}, Markus Küper², Martin Goetz^{2,3}, Karl Ernst Grund²

Affiliations:

¹Department of Gastroenterology, Limmatalspital Schlieren, Switzerland

²Department of Interdisciplinary Endoscopy, University Hospital Tübingen, Germany

³Department of Gastroenterology, University Hospital Tübingen, Germany

Background: Esophageal stenosis is a potential side-effect after radiochemotherapy of tumors of the head and neck region. In rare cases complete luminal obstruction can result. Most of the patients tolerate this situation for some time due to PEG placement prior to this.

Methods: Retrospective analysis of long-term results after direct recanalization via fluoroscopy and combined antegrade (*transoral route*) and retrograde (*via PEG-stoma*) esophagoscopy by use of a stiff guide-wire (Eder-Puestow) or direct needle knife cutting followed by bougienage or balloon dilation after successful recanalization. Luminal patency was maintained by repeated balloon-dilation or self bougienage.

Results: 14 patients (pt.) with time of esophageal closure 1–24 months, length of occlusion 3mm-6cm. Duration of endoscopic interventions 55 - 90 min. Success rates: primary success: 13/14 pt. (93%, 1x failure in case with 6cm occlusion); long-term success: 10/13 pt. (77%, 2x tumor-recurrence with discontinuation of therapy). PEG-removal possible: 6/10 pt., long-term self bougienage necessary: 5/10 pt.

Conclusions:

Recanalization of a complete postradiogenic esophageal long segment occlusion can be safely performed by combined anterograde/retrograde esophagoscopy, even in cases with long-lasting (max. 24months) and long-distant (max.4cm) esophageal occlusion.

Early colonoscopy in patients with complicated sigmoid diverticulitis even with covered perforation and abscesses: Preliminary results of a prospective study

Thomas Kratt^{1,2}, Markus Küper², Dietmar Stüker², Martin Goetz^{2,3}

Affiliations:

¹Department of Gastroenterology, Limmatalspital Schlieren, Switzerland
²Department of Interdisciplinary Endoscopy, University Hospital Tübingen, Germany
³Department of Gastroenterology, University Hospital Tübingen, Germany

Background: Early elective colonoscopy within the first days after clinical manifestation of an acute diverticulitis is discussed controversially and often seen as a contraindication for performing a total colonoscopy because of higher perforation risk.

Methods: Single center prospective study with preliminary results of 50 colonoscopies in 45 patients with acute sigma diverticulitis.

Inclusion criteria: Patients with acute sigma diverticulitis radiologically confirmed with CT including Hinchey stage II; intended high colonoscopy within 2 weeks after initial manifestation of an acute episode of sigma diverticulitis.

Results: Hinchey classification stage II in 19 cases. 3 cases with chronic-acute diverticulitis and complications (stenosis, fistula). Colonoscopy results: 76% performed 7 days, 94% within 10 days after initial manifestation of sigma diverticulitis. In 93.5% a total colonoscopy was possible. 11 cases (22%) without visible intraluminal inflammatory transformation of the sigmoid colon. In 22 patients (49%) tumorous lesions (21 colonic adenomas, 1 asymptomatic rectal cancer) were revealed. No endoscopy-related complications occurred.

Conclusion: Early elective total colonoscopy in acute and complicated sigmoid diverticulitis does not seem to be associated with an increased perforation risk. Surprisingly one out of five patients showed no signs of intraluminal inflammation, whereas we found a prevalence of colorectal tumors (adenomas, rectal carcinoma) in 49%, independently of the sigmoid diverticulitis.

G31

Endoscopic Treatment of Severe Bleeding in the Upper and Lower GI-Tract with the OTSC®-System: Analysis of 103 Cases

Thomas Kratt^{1,2}, Marcos Oberacher¹, Branislav Risti¹, Markus Küper², Dietmar Stüker², Maximilian v. Feilitzsch², Martin Goetz^{2,3}, Andreas Kirschniak²

Affiliations:

¹Department of Gastroenterology, Limmatalspital Schlieren, Switzerland
²Department of Interdisciplinary Endoscopy, University Hospital Tübingen, Germany
³Department of Gastroenterology, University Hospital Tübingen, Germany

Background:

The OTSC® (Over-The-Scope-Clip, Ovesco Endoscopy AG, Tübingen, Germany) has been developed for closure of iatrogenic, postsurgical or spontaneous perforations of the GI tract. However, only limited data have been published concerning the clinical outcome after endoscopic therapy of severe GI bleedings with the OTSC clip.

Methods:

Retrospective analysis of a prospective registry: From June 2006 through May 2015, 103 consecutive patients with severe acute upper or lower gastrointestinal bleeding, having been referred to our endoscopy units (Tübingen, Schlieren), received OTSC® treatment either as initial therapy or after failure of other methods.

Results:

103 cases intended to treat; 3 cases with technical failure. Age ≥ 64.7 yrs. (range 24-89). 63% upper GI tract bleed; 62% spontaneous bleed (37% ulcer bleed); 27% postendoscopic bleed; 11% postsurgical bleed. 73% Forrest I; 24% hemorrhagic shock; 29% with relevant anticoagulation therapy (Marcoumar, Pradaxa). 21% additional endoscopic hemostatic therapy necessary (fibrin glue, APC). Success rates: early success (≥ 72 h): 95.7%; long-term success 96.6%; overall success: 92.4%. Complications: 4% recurrent bleeding ≤ 72 h; 3.4% recurrent bleeding ≥ 72 h; 7.7% overall recurrent bleeding. 2.2% esophageal mucosal tear (OTSC system dep., no therapy necessary).

Conclusions:

The OTSC® system is feasible, safe and effective for treatment of patients with severe GI bleeding (persistent and recurrent) even with relevant anticoagulation therapy. Currently, a prospective randomized controlled multicenter study is ongoing to investigate the possibilities of the new device.

Identification of rejection after small bowel transplant changes: from histopathology to endoscopy. Preliminary results of a prospective study

G32

Subepithelial lesions of the stomach : monitoring or resection.

G34

Sébastien Godat (1), Maxime Robert (2), Fabrice Caillol (2), Erwan Bories (2), Christian Pesenti (2), Jean Philippe Ratone (2), Flora Poizat (3), Marc Giovannini (2).

(1) Division of Hepato and Gastroenterology, Centre Hospitalier Universitaire Vaudois CHUV, Lausanne, Switzerland.

(2) Division of Gastroenterology, Paoli-Calmettes Institute, Marseille, France.

(3) Division of Pathology, Paoli-Calmettes Institute, Marseille, France.

Background

Gastric subepithelial tumors represent a diagnostic and therapeutic challenge, given their histologic heterogeneity and potential malignant behavior.

Aims and Methods

To evaluate the interest, efficiency and safety of endoscopic resection for subepithelial gastric lesions of size < 20 mm. Single-center retrospective study in a tertiary care center.

Results

A total of 33 lesions (10 malignant/23 benign lesions) Mean histological size was 14.5 mm. 9 EMR, 18 ESD and 6 Hybrid Resection were performed. 93.9% lesions were resected in one piece. Complete and definitive resection, with a follow up of 6 months, was obtained in 96.7% of cases. A vertical resection was insufficient in 4 cases. One GIST needed a complementary surgical resection, 1 neuroendocrine tumor was successfully treated by a new ESD session and 2 pancreatic rests were not additionally treated given the benign character and the absence of residual tissue in endoscopic control after 6 months. There was only one severe adverse event (2.9%): 1 pneumoperitoneum with ESD, 3 bleeding with 1 ESD and 2 EMR, always treated conservatively or endoscopically.

Conclusion

Endoscopic resection is safe and should be the procedure of choice for both diagnosis and definitive resection for subepithelial gastric lesions of size under 20 mm.

Identification of rejection after small bowel transplant

changes: from histopathology to endoscopy. Preliminary results of a prospective study

Thomas Kratt^{1,2}, Dietmar Stüker², Alfred Königsrainer¹, Falko Fend⁴, Martin Goetz^{2,3}

Affiliations:

¹Department of Gastroenterology, Limmatalspital Schlieren, Switzerland
²Department of Interdisciplinary Endoscopy, University Hospital Tübingen, Germany
³Department of Gastroenterology, University Hospital Tübingen, Germany
⁴Department of Pathology, University of Tuebingen, Germany

Background: After small bowel transplantation acute tissue rejection and infection are possible complications with divergent emergency therapies. Questions: 1. Is zoom-endoscopic evaluation sufficient for the identification of tissue rejection? 2. Is the "optical biopsy" able to replace time-consuming histological findings in emergency situations? 3. Is it sufficient to inspect only the transplant stoma?

Methods: Prospective-consecutive observational study on 5 pat. Endoscopic survey (173 endoscopies with 160 biopsies) from the 2nd postop. day up to 36 months. SASAKI score for evaluation.

Results: Ad 1+2: 144 examinations without and 12 examinations with rejection, confirmed by histology with excellent correlation. 2 cases with mild rejection were not observable with endoscopy; 2 cases showed an inflammation in endoscopy, initially histologically described as rejection.

Ad 3: Remarkable differences in the findings of rejection due to "skipping rejection lesions"

Conclusion: Endoscopic staging according to the SASAKI score provides a sensitivity of 98% and a specificity of 93% and is suitable for the follow-up care after small intestine transplantation. The utilization of zoom-endoscopes is helpful but not obligatory necessary. Examining only the transplant stoma is not sufficient; a tissue rejection can occur as a "skipping lesion" in distant areas. Apoptosis-rate only partially correlates with the grade of tissue rejection; early histological signs of rejection are rather an increase of tissue eosinophilia and lymphocytic infiltrates.

Association between non alcoholic fatty liver disease severity and body composition in obese patients

Laurent Spahr, Laurence Genton, Sylvain Terraz, Claude Picard, Alain Golay, Zoltan Pataky, Gastroenterology/Hepatology ; Clinical Nutrition, Therapeutic Education for Chronic Diseases, Radiology, University Hospitals of Geneva.

Background: Non alcoholic fatty liver disease, or NAFLD, is the hepatic manifestation of the metabolic syndrome. The prevalence of NAFLD in obese patients increases in an epidemic proportion. The link between overweight or obesity is well accepted, but little evidence exists on the relationship between body composition and NAFLD severity. **Methods:** 15 overweight or obese patients (age 51 ± 8 yrs ; alcohol intake < 20 gr/week; BMI 35.1 ± 5 kg/m 2) with NAFLD were investigated at the HUG using a liver biopsy (with determination of the NAFLD activity score (NAS, semi-quantitative): steatosis, ballooning hepatocytes, lobular inflammation), bioelectrical impedance analysis (BIA) for body fat assessment and liver fat content using MRI. Linear relationship among variables were computed by Spearman's correlations. Multivariables analysis was performed with the linear regression model. **Results:** A positive association was observed between percent fat mass and the NAS score ($r = 0.72$, $p = 0.002$), which persisted after adding BMI to the model ($\beta = 0.22$, $p = 0.002$). A positive association was also found between serum C-reactive protein levels and liver steatosis as measured by MRI ($r = 0.63$, $p = 0.01$). We found however no association between body fat by BIA and liver fat content measured by MRI. **Conclusion:** In this small group of obese and overweight patients, the severity of NAFLD is associated with body composition. Increased fat mass is linked to a higher NAS score on liver histology. Body composition as assessed by BIA may contribute to a better patient selection for liver biopsy. However, these findings needs to be confirmed in a larger group of patients (Funded by FLAGS in Geneva)

H1

T and B Cell Responses and Previous Exposure to Hepatitis B Virus in "Anti-HBc alone" Patients

Qixia Wang^{1,4}, Philip Sachse¹, Mariam Semmo², Megha Lokhande^{1,7}, Matteo Montani³, Jean-François Dufour^{1,7}, Fabien Zoulim⁵, Paul Klennerman⁶, Nasser Semmo^{1,7}

Affiliations

1. Hepatology, Department of Clinical Research, Inselspital, University of Bern, CH-3010 Bern
2. Department of Nephrology, Inselspital, University of Bern, CH-3010 Bern
3. Institute of Pathology, Inselspital, University of Bern, CH-3010 Bern
4. State Key Laboratory for Oncogenes and Related Genes, Key Laboratory of Gastroenterology & Hepatology, Ministry of Health, Division of Gastroenterology and Hepatology, Ren Ji Hospital, School of Medicine, Shanghai Jia Tong University, Shanghai Institute of Digestive Disease, 145 Middle Shandong Road, Shanghai 200001, China
5. Inserm, U1052, UMR CNRS 5268, Centre de Rech. en Cancérologie de Lyon, F-69003 Lyon
6. Peter Medawar Building for Pathogen Research, University of Oxford, UK-Oxford, OX1 3SY
7. University Clinic of Visc. Surgery and Medicine, Dep. of Hepatology, Inselspital, CH-3010 Bern

Background & Aims: A serologic response to hepatitis B virus (HBV), defined as "anti-HBc alone" is commonly observed, but its significance remains unclear. This study aimed to define the relationship between "anti-HBc alone" serostatus and HBV infection, including HBV-specific T and B cell memory responses.

Methods: 31 "anti-HBc alone" patients were enrolled. Total HBV DNA and cccDNA were tested by nested polymerase chain reaction (PCR) analysis in liver samples from 22 "anti-HBc alone" patients vs controls (chronic or resolved HBV infection), followed by HBsAg/HBcAg immunohistochemical (IHC) staining. IFN- γ secretion by HBV-specific T cells was compared in individuals who were "anti-HBc alone" (n=27), resolved HBV (n=21), chronic HBV (n=24) and 12 healthy controls using enzyme-linked immunospot (ELISpot) assays. An HBsAg-IgG B-cell ELISpot assay was performed in "anti-HBc alone" patients before and after one dose of recombinant HBsAg vaccine.

Results: The majority (23/31, 74.2%) of the "anti-HBc alone" individuals were co-infected with HCV. Infrequent intrahepatic total HBV DNA (2/22, 9.1%) and cccDNA (1/22, 4.5%) were detected in biopsies (Fig. 1); HBsAg and HBcAg IHC staining was negative in Anti-HBc alone patients in contrast to those with chronic HBV infection (Fig. 2). HBV-specific T cell responses were similar between "anti-HBc alone" individuals and HBV resolvers (Fig. 3). Circulating HBV-memory B cell responses were detected in all "anti-HBc alone" individuals, consistent with an HBsAg-specific memory pool (Fig. 4). After one HBV vaccine dose, increased anti-HBs antibody levels were observed, accompanied by an expansion of HBsAg-specific memory B cells (Fig. 4) ($P = 0.0226$).

Conclusions: "Anti-HBc alone" individuals showed HBV-specific T cell and memory B cell responses typical of previous viral exposure and protective memory, suggesting a resolved infection. B and T cell ELISpot seem to be more sensitive in tracking HBsAg-specific responses than routine Anti-HBs EIA test kits, suggesting that the ELISpot might be the better tool in daily clinical routine in this kind of patient population.

H2

Liver transplantation for hepatocellular carcinoma: the Geneva experience.

L Elkrief, I Morard, L Spahr, PA Elton Rouge, P Morel, T Berney, P Majno, C Toso, E Giostra. Service de Gastroentérologie et Hépatologie, Service de Transplantation, Service de Chirurgie viscérale, Hôpitaux Universitaires, Genève

Background We compared survival and prognostic factors in patients transplanted for hepatocellular carcinoma (HCC) to patients with other liver diseases. Patients were included in the waiting list if they met the "Milan criteria" and since 2009 the "TTV/AFP criteria".

Results: from 1987 to 2014, 594 adults patients were transplanted: 159 (27%) for hepatocellular carcinoma, and 434 (73%) for other causes. The other causes of were: HCV (20%), alcoholic cirrhosis (12%) and HBV (7%). There was a significant difference in survival rate: 77% at 5 years for CHC patients versus 85% for non-CHC, and 63% at 10 years for CHC versus 75% (log rank= 0.027). Among the 159 adults patients transplanted for HCC 43 died, including 11 (6.9%) from recurrence, which represents the main cause of death in this group of patients. Of the 435 patients transplanted for other causes 111 died, the main reasons of death being: cancers (20%), HCV recurrence (13%) and cardiovascular diseases (13%). The causes of death were not significantly different between the 2 groups, except for CHC recurrence.

In the 102 patients on the waiting list between January 2010 and January 2015, 23(22.5%) were withdrawn. 16 out 102 (15.6%) patients had tumor progression as cause of dropout, and this represents 70% of the total dropout.

Conclusion: survival is significantly lower in our patients transplanted for HCC because of recurrence. New treatments are eagerly needed because death from recurrence is the reason for the difference in survival.

Evolution of portal haemodynamics in patients with decompensated alcoholic liver disease (ALD) and relationship with alcohol relapse

Laurent Spahr, Flavie Furrer, Cassandra Oropesa. Gastroenterology and Hepatology, HUG

Decompensated ALD presenting as acute on chronic liver failure is characterized by an increased portal pressure due to intrahepatic vascular resistance in relationship with inflammation and endothelial dysfunction. The evolution of portal haemodynamics over time, however, is ill defined. We explored the evolution of portal pressure, inflammation and liver failure following decompensation. **Methods:** We selected 21 patients with cirrhosis from a cohort study (age 51 yrs; recent alcohol intake: 135 gr/day; alcoholic hepatitis: 81%; 47% steroid treated; MELD score 17.2 ± 1.4 ; ascites: 71%; betablockers: 9%) who were reevaluated after 3 months. We measured the hepatic venous pressure gradient (HVPG, wedge (WHVP) minus free FHVP, mmHg) both at baseline and at 3 months, and inflammation using plasma C-reactive protein (CRP: N: < 10 mg/L). Alcohol relapse was monitored over time. **Results:**

	Baseline	3 months	p value
MELD	17.2 + 6.5	12.8 + 4.4	< 0.05
HVPG (mmHg)	17 + 4.2	15.7 + 4.2	0.28
WHVP (mmHg)	30.2 + 7.5	25.2 + 5.9	< 0.05
CRP (mg/L)	22.1 + 17.6	9.7 + 12	< 0.001
Ascites (%)	71	33	< 0.05

Nine (n=9) patients return to alcohol during FU (heavy regular consumption in 7, mean 620 gr/week, isolated relapse in 2). Abstinent patients compared to alcohol relapsers presented a trend towards a better liver improvement (delta MELD -16 vs -3, $p=0.06$) but a parallel evolution of portal haemodynamics (% changes in HVPG: -2 vs -12, $p=0.15$; % changes in WHVP: -12 vs -15, $p=0.56$). **Conclusion:** in this selected population of patients with chronic ALD, portal haemodynamic parameters and liver insufficiency improve at 3 months after decompensation. Alcohol relapse doesn't seem to influence the evolution of these parameters. (partially funded by FLAGS)

H4

Real World Effectiveness, Safety and Costs of Direct-Acting Antiviral Agents in Chronic Hepatitis C Treatments

 Patrizia Künzler¹, Irina Bergamin¹, David Semela¹
¹Clinic for Gastroenterology and Hepatology, Cantonal Hospital of St. Gallen, CH-9007 St. Gallen

Background/Aims: Second generation direct-acting antiviral agents (DAA) have revolutionized hepatitis C virus (HCV) treatment since their introduction in Switzerland in 2014. We evaluated efficacy, safety and direct costs of current DAA regimens in comparison to therapies with first generation protease inhibitors (PI) in 2011-2013.

Methods: Single center retrospective analysis of outpatients treated for chronic hepatitis C between 2011 and 2015.

Results: Overall 112 (28 PI vs. 84 DAA) patients received treatment. *PI Group (2011-2013):* 28 GT1 received first generation PI (Telaprevir or Boceprevir) with Interferon (IFN)/Ribavirin (RBV), mean age 49 (34-62), fibrosis F3 or F4 in 20 (71%), 18 (64%) treatment experienced. *DAA Group (2014-ongoing):* 84 with predominantly GT1 (56 (66%)), mean age 55 (33-83), fibrosis stage F3 or F4 in 75 (89%), 28 (33%) treatment experienced. Patients were treated according to Swiss label or within early access programs: 25 (30%) received IFN/RBV/Sofosbuvir (SOF), 16 (19%) 3D-combination, 15 (18%) SOF/Ledipasvir, 11 (13%) SOF/RBV, 11 (13%) SOF/RBV with Simeprevir, 6 (7%) SOF/RBV with Daclatasvir.

Outcomes PI Group: 16 (57%) with SVR12, 9 (32%) stopped treatment due to futility rules, 2 (7%) due to serious adverse events (SAE), 1(4%) due to breakthrough. SAEs occurred in 6 (21%); in addition 7 (25%) needed blood transfusion or EPO. Average treatment duration was 26 weeks (range 4-48 weeks) with mean drug costs per SVR12 of CHF 63'997.

Outcomes DAA Group: End of treatment response in 61/61 (100%) while ongoing therapy in 23 patients, SVR4 in 51/52 (98%) and SVR12 in 29/31 (94%) with relapse in 2/31 (6%). SAEs occurred in 7 (8%), all due to comorbidities and unrelated to treatment. Two (2%) RBV-associated blood transfusion occurred. One (1%) patient stopped DAA therapy three weeks earlier due to an episode of depression. Average treatment duration was 15 weeks (9-24 weeks) with mean drug costs per SVR12 of CHF 73'386.

Conclusions: Current DAA regimens are safe, well tolerated and show excellent cure rates even in treatment-experienced patients with advanced liver disease and advanced age. Overall SVR12 rates increased significantly with second generation DAAAs while costs remain high.

H5
Intestinal microbiota significantly alters hepatic gene expression and plasma bile acid profile in mice with acute cholestasis.

 Sheida.Moghadamrad¹, Irene.Keller³, Kathy.D.McCoy², Johan.Mattsson⁵, Cédric.Bovet⁵, Rolf.Jaggi⁴, Andrea. De Gottardi¹.

 Department of Clinical Research, Hepatology¹, Gastroenterology², Interfaculty Bioinformatics³, Genomics⁴, Clinical Metabolomics Facility⁵, University-hospital of Berne, Switzerland.

Background: Intestinal microbiota is recognized as a new "organ" which plays an important role in the regulation of metabolism of bile acids (BA). We were therefore interested in assessing the effects of intestinal microbiota on hepatic gene expression profile and plasma BA composition in basal conditions and in cholestasis.

Methods: We induced acute cholestasis in germ free (GF) and altered Schaedler flora colonized (ASF) mice by performing bile duct ligation (BDL) for 5 days. We evaluated the gene expression profile in the liver of these mice compared to non-cholestatic control groups using next generation sequencing and pathway analysis, and we measured the plasma concentration of 20 BA by UPLC-MS analysis.

Results: We found that acute cholestasis was associated with a significantly different expression of groups of genes involved in the regulation of the immune system, the accumulation of extracellular matrix and of oxidative processes in both ASF and GF mice. Moreover, 448 genes were significantly over expressed in the absence of intestinal microbiota involving in pathways such as organic acid catabolic and fatty acid metabolic processes, leukocyte migration, external side of plasma membrane. In non-cholestatic mice the absence of microbiota significantly induced or suppressed the expression level of 80 genes. Under basal conditions, chenodeoxycholic acid and muricholic acid were detected only in ASF-mice, whereas tauromuricholic acid was detected only in GF-mice. The concentration of the majority of BA increased after BDL, but more markedly in ASF mice.

Conclusion: Intestinal microbiota significantly alters the hepatic gene expression profile and the plasma bile acid composition in basal conditions in mice. Changes observed after BDL suggest that microbial-induced differences may impact the course of cholestasis.

H6
POSTERS SURGERY
One year experience in enhanced recovery pathway for colorectal surgery

Cornelia Künzli, Andrea Goldmann, Michel Adamina, Stefan Breitenstein

Department of Visceral Surgery, Kantonsspital Winterthur

Background: Implementation of enhanced recovery after surgery (ERAS) pathways is associated with a reduction in morbidity and length of hospital stay (LOS). The complexity of current ERAS pathways has hampered their widespread use. This paper reports our first-year experience in implementing ERAS in elective colorectal surgery.

Methods: ERAS pathways were implemented along a structured process including all stakeholders. Outcome and pathway compliance were prospectively followed. Data were compared to a retrospectively analysed consecutive pre ERAS cohort.

Results: Within the first year 114 patients were operated on. They were compared to a 153-patient consecutive pre ERAS cohort. Patients' age, morbidity and procedures performed did not differ significantly. In both groups most patients underwent colorectal resections for diverticular disease or cancer. Morbidity was halved from 45.7% in the pre ERAS group to 24.5% in the ERAS group ($p = 0.007$). Severe complications (Dindo-Clavien III-V) showed a decrease from 12.4% to 9.6%. Median LOS decreased from 9 (IQR 7, 13) to 7 (IQR 3, 11) days ($p = 0.004$). The readmission rate remained stable.

Conclusion: Successful implementation of ERAS pathway in colorectal surgery is feasible and worth the efforts. It requires significant work, close monitoring and a dedicated team.

S1
Major liver resection in non-cirrhotic liver without inflow occlusion is feasible and safe

Anja Schärli, Stefan Gutknecht, Markus Weber

Department of Visceral, Thoracic and Vascular Surgery, Triemli Hospital Zürich

BACKGROUND: Several studies showed conflicting results concerning the benefit of the Pringle maneuver caused by a hepatic ischemia-reperfusion injury. Our aim is to investigate the clinical outcome of patients who underwent liver resection without inflow occlusion.

METHODS: Over the last 48 months all liver resections at our clinic were consecutively performed without inflow occlusion. We retrospectively analyzed 63 patients who underwent major liver resection (3 \geq segments) for various reasons. The analyzed parameters were estimated blood loss (EBL), transfusion requirement, mortality, and morbidity during the first 30 postoperative days as well as liver function during the first 7 days.

RESULTS: Mean EBL was 523 ml ($SE \pm 52ml$). 12 patients (19%) were in need of a blood transfusion (1.8 units per patient in need). The average ICU stay was 1.62 days ($SE \pm 0.38$), the average hospital stay was 12.32 days ($SE \pm 0.76$). 4 patients (6%) had to undergo an abdominal reoperation. There was no postoperative death and one case in need of hemodialysis. In comparison to other study populations our case series has no higher EBL or complication rate. Peak liver enzymes and prothrombin time were equal or better to other studies.

CONCLUSION: Major liver resection without inflow occlusion did not result in any discernible disadvantage and had a short ICU and hospital stay and a minimal reoperation rate. Based on our experience we suggest restricting inflow occlusion in major liver resection to exceptional patients.

S2

Incidence of early postoperative complications comparing linear versus circular stapling technique in laparoscopic Roux-en-Y gastric bypass

Larissa C. Vines¹, Nabil Kalak¹, Stefan Aczel², Marc Schiesser¹
¹Klinik für Allgemein-, viszeral- u. Transplantationschirurgie, Kantonsspital St. Gallen
²Klinik für Endokrinologie, Diabetologie, Osteologie u. Stoffwechselkrankungen, Kantonsspital St. Gallen

Background

Different surgical techniques to perform a Laparoscopic Roux-en-Y gastric bypass (RYGB) do exist. Currently there are two techniques, which have emerged as standard for the gastrojejunostomy: The linear stapling and circular stapling technique. The aim of this study was to compare the two techniques regarding postoperative morbidity and weight loss.

Methods

We compared two consecutive cohorts at a single institution between November 2012 and June 2014 assessing the incidence of postoperative complications (30d) and weight loss at one year after the operation. 109 patients underwent a RYGB with a 21-mm circular stapler. 134 patients underwent a RYGB performed by linear stapling technique.

Results

Preoperative BMI and incidence of comorbidities were similar in both groups. The incidence of postoperative complications was significantly higher in the group using the circular stapling technique with 23.9% vs 4.5% ($p=0.000009$). Stenosis at the gastrojejunostomy occurred significantly more often in the circular stapling group with 14(13%) cases versus 0 in the linear group. The rate of leakages (2 vs 1) and length of stay (6.5 vs 6.2 days) was similar in both groups. There was no significant difference regarding weight loss at one year (29.4 vs 29kg/m²).

Conclusion

Linear stapled gastric bypass displays less postoperative complications, with similar weight loss at one year and should therefore be the procedure of choice.

S3

Long-term outcome of laparoscopic adjustable gastric banding (LAGB): results of a Swiss single-centre study of 405 patients with up to 18 years follow-up

Fabrizio Vinzens¹, Ashley Kilchenmann¹, Valentin Zumstein¹, Ralph Peterli¹.¹Department of Surgery, St. Claraspital, Basel.

Background: LAGB seemed to be a very promising bariatric procedure but many studies showed high rates of re-operation due to complications or insufficient weight loss. There is lack of long-term studies with a follow-up beyond 15 years.

Methods: Retrospective analysis of prospectively collected clinical data on weight loss, co-morbidities, re-operations, complications, and quality of life including BAROS score (Bariatric Analysis and Reporting Outcome System) in a cohort of 405 patients having undergone LAGB was performed. Follow up (FU) was conducted in our outpatient clinic or via telephone interview.

Results: 405 patients (age 41 ± 10 years, BMI 44.3 ± 6 kg/m²) were treated with a LAGB between 1996 and 2010. Mean FU was 13 ± 3 years, with a FU rate of 85% (range 8–18 years) corresponding to 343 patients. 100 patients exceeded a FU period of 15 years. In 216 patients (63%) the LAGB was removed and another bariatric procedure performed: 32 (9%) patients underwent lap. sleeve gastrectomy, 102 (30%) lap. gastric bypass and 82 (24%) biliopancreatic diversion with duodenal switch due to either band intolerance, slippage, insufficient weight loss or secondary weight regain; 27 (8%) patients refused further bariatric surgery after band removal. Total failure rate was 63%. Finally, 100 (30%) patients still have the band in place with a mean BMI of 35 ± 7 kg/m², corresponding to an excessive BMI-loss of $48 \pm 27\%$. Of these, the failure rate was 25%, according to BAROS; 50% had a good to excellent outcome.

Conclusion: More than 10 years after LAGB 70% of patients lost their bands and only 12% have the band in place and a good to excellent result according to BAROS.

S5

Six years results of revisional laparoscopic Roux-en-Y gastric bypass (rLRYGB) after failed gastric banding (LAGB)

Fabrizio Vinzens¹, Ashley Kilchenmann¹, Valentin Zumstein¹, Ralph Peterli¹.¹Department of Surgery, St. Claraspital, Basel.

Background: Long-term failure of LAGB is high. At our institution we initially performed bilio-pancreatic diversion with duodenal switch (BPD-DS) in cases of LAGB failure as part of a staged concept, possibly over treating a number of patients. Lately, rLRYGB has become popular after failed LAGB. Most published series are based on a mean follow-up of 2-3 years.

Methods: Retrospective analysis of prospectively collected data on weight loss, co-morbidities, re-operations, complications, and quality of life including BAROS score (Bariatric Analysis and Reporting Outcome System) of rLRYGB was performed, with a mean follow up of 6 years (range 2-9 years).

Results: Overall 74 patients met our inclusion criteria. Baseline characteristics at the time of LAGB: age 40 ± 10 years, 55 women and 19 men, weight 120 ± 15 kg, BMI 42 ± 4 kg/m². On average it took 6.3 ± 3.1 years from initial LAGB to rLRYGB. The most common indications for rLRYGB were band intolerance (59%), band slippage (12%), pouch dilatation (6%), insufficient weight loss (3%) and/or secondary weight regain (9%). Baseline BMI decreased to 35.2 ± 5.9 kg/m² at the time of rLRYGB and further to 30 ± 4.8 kg/m² after a mean follow-up of 6 ± 2.2 years, resulting in an excessive BMI loss of $72.6 \pm 27.9\%$. According to BAROS score, 60% had a good to excellent result. Remission/improvement rate for diabetes and arterial hypertension was over 50%.

Conclusion: Long-term results of rLRYGB as a revisional procedure after failed LAGB proved satisfactory concerning weight loss and quality of life/co-morbidities using BAROS score.

S4

The value of extended lymphadenectomy in distal oesophageal cancer

S. Wolf, R. Warschkow, B. Schmied, M. Schiesser
 Kantonsspital St. Gallen, Department of Surgery, Switzerland

Background: Curative esophagectomy provides the best prognosis in patients with esophageal cancer. The extent of lymphadenectomy and its potential benefit has been controversially discussed over the last 2 decades. The aim of this study was to assess the benefit of an extended lymphadenectomy regarding survival.

Methods: 3087 patients with stage I-III distal esophageal cancer were identified from the Surveillance, Epidemiology, and End Results (SEER) database between 2004 and 2012. The impact of extended lymphadenectomy with 20+ retrieved regional lymph nodes (RLN) on survival was assessed using both multivariate Cox proportional hazards model and propensity score matching (PSM).

Results: In 727 patients (23.6%) 20+ RLN were retrieved. This rate increased from 14.1% in 2004 to 30.1% in 2012 ($P<0.001$). The 5-year overall and cancer-specific survival in patients with 20+ retrieved RLN were 48.4% (95%CI: 44.0-53.2%) and 55.2% (95%CI: 50.8-60.0%) compared to 43.3% (95%CI: 40.9-45.7%) and 49.7 (95%CI: 47.3-52.3%) in patients with less retrieved RLN. Extended lymphadenectomy of 20+ RLN was associated with an increased overall survival in unadjusted (hazard ratio (HR) of death=0.85, 95%CI: 0.75-0.97, $P=0.013$) and PSM-adjusted Cox regression (HR=0.78, 95%CI: 0.66-0.91, $P=0.002$). Similar results were observed for cancer-specific survival in unadjusted (HR=0.86, 95%CI: 0.75-0.99, $P=0.037$) and PSM-adjusted Cox regression (HR=0.81, 95%CI: 0.69-0.97, $P=0.016$).

Conclusions: Extended lymphadenectomy resulted in a significant survival benefit in unadjusted and PSM-adjusted population-based analysis. Therefore, extended lymphadenectomy should be advocated in all patients undergoing esophagectomy.

S6

Laparoscopic Sleeve Gastrectomy and Roux-Y-Gastric Bypass are equally effective up to three years. Results of the prospective randomized Swiss Multicentre Bypass Or Sleeve Study (SM-BOSS)

Ralph Peterli¹, Bettina Wölnerhanssen², Diana Vetter³, Philipp Nett⁴, Yves Borbely⁴, Beatrice Kern¹, Markus Gass¹, Thomas Peters¹, Marc Schiesser⁵, Bernd Schultes⁵, Marco Bueter³

¹ Departments of Surgery and Internal Medicine, St.Claraspital, Basel

² Department of Biomedicine, University of Basel

³ Department of Visceral and Transplantation Surgery, University Hospital Zürich

⁴ Department of Surgery and Medicine, University Hospital, Bern

⁵ Department of Surgery, Kantonsspital St.Gallen

⁶ eSwiss Medical and Surgical Center, St.Gallen

Background: Laparoscopic Sleeve Gastrectomy (LSG) is performed almost as often in Europe as laparoscopic Roux-Y- Gastric Bypass (LRYGB). We present the 3-year results of this randomized clinical trial comparing the two procedures.

Methods: Initially 217 patients (LSG, n=107; LRYGB, n=110) were randomized to receive either LSG or LRYGB at four bariatric centres in Switzerland. Mean BMI of all patients was $44 \pm 11 \text{ kg/m}^2$, mean age was 43 ± 5.3 years, and 72% of patients were female. Minimal follow-up was three years with a rate of 97.7% at three years after surgery. Both groups were compared for weight loss, co-morbidities, quality of life according to GIQLI and BAROS score, and complications.

Results: Excessive BMI loss was similar between LSG and LRYGB at each time point (one year: $72 \pm 22\%$ vs $77 \pm 21\%$, p=0.2; two years: $74 \pm 30\%$ vs $78 \pm 30\%$, p=0.4; three years: $70 \pm 24\%$ vs $74 \pm 23\%$, p=0.2). Prevalence of comorbidities was significantly reduced after both procedures except for GERD, which showed a higher remission rate after LRYGB. Quality of life increased significantly in both groups after one and three years post surgery. Within three years of follow-up there was no difference in number of complications treated by reoperation (LSG, n=9; LRYGB, n=16, p=0.15) and number of complications treated conservatively: peptic ulcer (LSG, n=0; LRYGB, n=1), stricture (LSG, n=0; LRYGB, n=1), kidney stones (LSG, n=2; LRYGB, n=1), micronutrient deficiencies (LSG, n=86; LRYGB, n=92).

Conclusions: LSG and LRYGB are equally efficient regarding weight loss, improvement of comorbidities, quality of life, and complications up to 3 years after surgery.

Postoperative inflammatory syndrome after CRS/HIPEC

S8

Dilmurodjon Eshmuminov, Marcel Schneider, Ksenija Slankamenac, Kuno Lehmann

Background: Cytoreductive surgery (CRS) and hyperthermic intraperitoneal chemotherapy (HIPEC) is a treatment option in well-selected patients with peritoneal tumors. There is a risk of severe morbidity and even mortality; however knowledge about the complex pathophysiology of CRS/HIPEC is scarce. Here we describe the postoperative dynamics of inflammatory parameters in presence or absence of infectious complications after CRS/HIPEC. **Methods:** Ninety patients after completed CRS/HIPEC between 2009-2013 were taken from the prospective Zurich database, including perioperative complications, clinical parameters and laboratory values. Patients with infectious complications (CDC Definition 1999) were compared to patients without infection. Routine diagnostics included clinical examination, chest X-ray, urine and catheter cultures, and CT scan in selected cases. To exclude the influence of postoperative management, the patients were additionally compared to patients (n=77) after esophagectomy. **Results:** The majority of patients had appendix tumors (53%) and colorectal carcinomatosis (30%), with a female preponderance (61%), and a median age of 51 years. The two groups (+/- infection after CRS/HIPEC) were not different regarding the extent of the disease (median PCI 10), and surgery (splenectomy, number of colon anastomoses). Surprisingly, also the group without infection, showed a second postoperative peak in CRP levels, observed only after CRS/HIPEC but not after oesophagectomy. If this CRP value was $>80 \text{ mg/l}$, it was predictive for major complications (OR 3.3, p=0.042) in patients after CRS/HIPEC. Patients after CRS/HIPEC developed less leucocytosis in case of infection compared to patients after esophagectomy. **Conclusion:** There is a second rise of inflammatory parameters (CRP) after uncomplicated CRS/HIPEC, and a reduced capacity for leukocytosis in case of infectious complications. This observation is novel and highlights the complex pathophysiology of CRS/HIPEC.

Double-barrelled wet colostomy following exenteration for locally advanced and recurrent pelvic cancer: early experience

S9

Rebecca Kraus, Hannah Palacci, Catherine Bowes, Roel Hompes, Bruce George, Richard Guy
Department of Colorectal Surgery, Oxford University Hospitals, UK

Introduction

In patients undergoing pelvic exenteration for advanced cancer, two stomas are usually required – a colostomy and ileal conduit. Whilst inconvenient for the patient, this approach limits options for perineal flap reconstruction. We describe the technique, and our early experience of, the double-barrelled wet colostomy (DBWC) as an alternative option.

Methods

Exenterations were undertaken via a midline incision, with proximal vascular ligation, lymphadenectomy, en bloc organ resection and removal of specimens via the perineum. Each DBWC was constructed by anastomosing the ureters separately, over infant feeding tubes, into a 15cm blind length of colon distal to the colostomy site. Following Vertical Rectus Abdominis Muscle (VRAM) flap mobilisation and abdominal closure, the DBWC was exteriorised and matured over a bridge in the left iliac fossa, recovering the infant feeding tubes to drain into a colostomy bag.

Results

Eight patients (6 males; median age 61) underwent exenteration and DBWC over a 9 month period. Pathologies were: locally advanced primary rectal cancer with prostatic invasion (5 patients); recurrent rectal cancer or anal cancer with vaginal and urethral involvement (2 patients); prostate cancer invading rectum (1 patient). Seven patients received preoperative chemoradiotherapy and 7 had perineal reconstruction with a VRAM flap. Median hospital stay was 21 days. Procedure-related complications included: urinary sepsis which resolved after ureteric stent removal (2 patients), and one collection around a VRAM flap needing drainage.

Conclusion

Early experience with DBWC following exenteration is encouraging. No major complications were seen, urinary sepsis does not appear to be a significant risk, as faecal and urinary streams do not mix, and patient satisfaction is high. Avoidance of a second stoma eliminates the risk of ileal conduit complications and allows the harvesting of a VRAM flap for perineal reconstruction

Tutorial assistance for board certification in surgery: Frequency, associated time and costs

S10

Mechera Robert*, Department of Surgery, University Hospital Basel

Dell-Kuster Salome*, Institute for Clinical Epidemiology and Biostatistics, University Hospital Basel

Marco von Strauss und Torney; Department of Surgery, Cantonal Hospital Aarau

Igor Langer; Department of Surgery, Lindenhof Hospital Bern, Rachel Rosenthal; Department of Surgery, University Hospital Basel

*These authors contributed equally to this work.

Background: Tutorial assistance is related to extra time and cost and the hospitals' financial compensation for this activity is under debate. We aimed at quantifying the time and resulting cost required to train one surgical resident in the operating theatre for board certification in Switzerland.

Methods: We analysed data of 212'948 surgeries carried out between 2008 and 2012. The percentage of procedures carried out by residents as compared to three higher levels of surgical seniority was assessed. Average duration of procedure categories by different seniority levels was calculated and extra time and resulting cost were analysed.

Results: On average, residents performed about a third of all surgeries even in typical teaching procedures like appendectomies. An increase in duration and cost of well-defined procedure categories was demonstrated if a resident performed the procedure. In less well-defined ones, residents seemed to perform less difficult procedures than senior consultants resulting in shorter durations of surgery. Total cost for duration of surgeries necessary for board certification reaches a maximum of CHF 2,7 Millions.

Conclusion: The low percentage of procedures performed by trainees may make it difficult to fulfil requirements for board certification in a broad training system within a reasonable period of time, especially in times when reimbursement of training remains an unsolved issue and a relevant cost driver.

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