

ΗΠΑΤΕΚΤΟΜΗ ΚΑΙ ΜΕΤΕΓΧΕΙΡΗΤΙΚΕΣ ΕΠΙΠΛΟΚΕΣ

Μετεγχειρητική Χολόρροια:

Συντηρητική και Χειρουργική Αντιμετώπιση

Δημήτρης Π. Κορκολής

Χειρουργός

Διευθυντής Χειρουργικής Ογκολογικής Κλινικής

ΓΑΟΝΑ «Άγιος Σάββας»

ΕΙΣΑΓΩΓΗ

- Ηπατεκτομή σήμερα:
 Νοσηρότητας < 30%
 Θνητότητας < 2%
- Η χολόρροια μετά ηπατεκτομή παραμένει συχνή
- 3.1% - 15.6% των ηπατεκτομών χωρίς HJ - 30% με HJ
- ↑↑↑ αριθμού ηπατεκτομών
- ↑↑↑ ογκολογικών ενδείξεων
- MIS - ERAS
- Staged Hepatectomy
- ALPPS
- Πολύπλοκες ηπατεκτομές με αποκαταστάσεις αγγείων/χολαγγείων

ΕΙΣΑΓΩΓΗ

- Χολή – Νεκρωτικό υλικό – Αίμα στον κενό χώρο της εκτομής
- Ανάπτυξη μικροβιακών πληθυσμών
- Έκπτωση host defense μηχανισμών = ΣΗΨΗ
- Απόστημα - Περιτονίτις - Ηπατική ανεπάρκεια - Θνητότης 30%
- Παράταση νοσηλείας σε ΜΕΘ
- Διατήρηση ενδοκοιλιακών παροχετεύσεων
- Μακρά νοσοκομειακή νοσηλεία
- Διαγνωστικές - ενδοσκοπικές παρεμβάσεις (20-70%)
- Χειρουργική επανεπέμβαση
- QoL
- Κόστος
- RFS – OS???

Bile leakage after hepatobiliary and pancreatic surgery: A definition and grading of severity by the International Study Group of Liver Surgery

Consensus proposal of the ISGLS for a definition and grading of bile leakage after hepatobiliary and pancreatic surgery

Definition	Bile leakage is defined as fluid with an increased bilirubin concentration in the abdominal drain or in the intra-abdominal fluid on or after postoperative day 3, or as the need for radiologic intervention (ie, interventional drainage) because of biliary collections or relaparotomy resulting from bile peritonitis. Increased bilirubin concentration in the drain or intra-abdominal fluid is defined as a bilirubin concentration at least 3 times greater than the serum bilirubin concentration measured at the same time.
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Grade	
A	Bile leakage requiring no or little change in patients' clinical management
B	Bile leakage requiring a change in patients clinical management (eg, additional diagnostic or interventional procedures) but manageable without relaparotomy, <i>or</i> a Grade A bile leakage lasting for >1 week
C	Bile leakage requiring relaparotomy

ΠΡΟΔΙΑΘΕΣΙΚΟΙ ΠΑΡΑΓΟΝΤΕΣ

- Ασθενής
 - Ηλικία
 - Φύλο
 - Νοσηρότητες (ΑΥ, ΣΔ, ↓Hg, ↑WBCs, albumin<3.5, Ασκίτης, ΧΜΘ)
 - PVE - TACE
- Ηπατική Παθολογία

Tanaka et al, ¹⁰³ 2002	HCC (316)	23 (7.3%)
	CCC (9)	3 (33.3%)
	Metastatic (33)	0 (0%)
	Other (5)	0 (0%)
Nagano et al, ⁸² 2003	HCC (126)	9 (7.1%)
	Metastatic (187)	17 (5.4%)
Sadamori et al, ¹⁰ 2013	HCC (359)	46 (12.8%)

ΠΡΟΔΙΑΘΕΣΙΚΟΙ ΠΑΡΑΓΟΝΤΕΣ

Εκτομή S1, S4, S5, S8

R Anterior

L Medial

Central bisectionectomy

L + L extended hepatectomy

Extended R

OR time >300

EBL >500

Pringle>45min

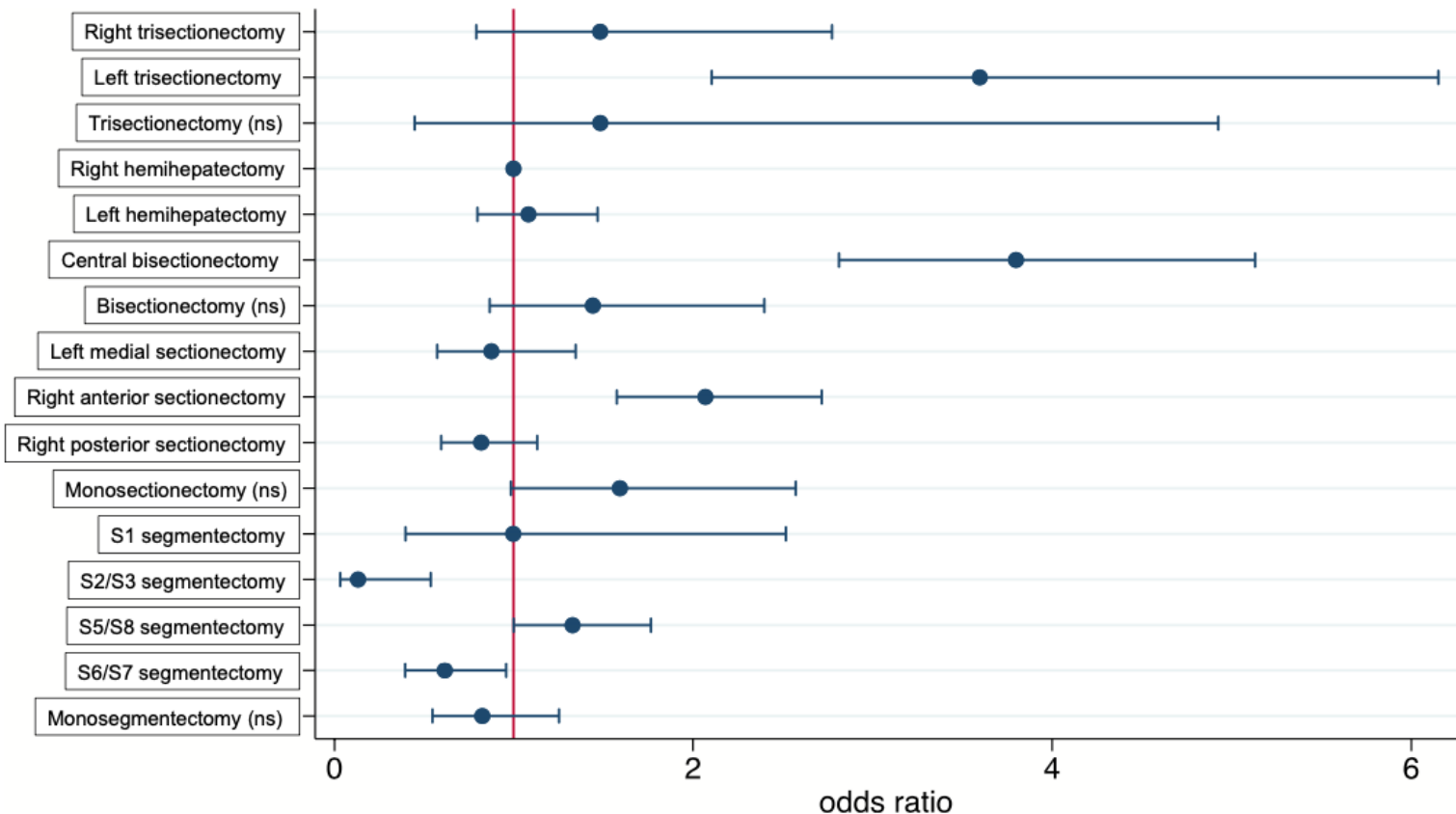
↑ Cut surface area

Glisson's exposure

HJ – Oddi's dysfunction

Study Author	% Leak	Significant Factors	
Lo et al ⁷⁶	8.1	Age Non-HCC Mean hemoglobin Estimated blood loss Platelets Noncirrhotic livers Logistic Regression: Advanced age Preoperative leukocytosis	Childs B and C Major hepatectomy Caudate resection T-tube drainage Concomitant bilioenteric or bowel anastomosis Mean operative time Left hepatectomy Prolonged operative time
Sadamori et al ¹⁰ (all HCC)	12.9	Trisegmentectomy Repeat hepatectomy MVA OR time >300 min	OR time >300 min Blood transfusion
Zimmitti et al ¹²	4.8	Preoperative jaundice Portal vein embolization Biliary tumors Repeat hepatectomy Two-stage resection Extended resection Caudate resection En bloc diaphragm resection MVA Repeat hepatectomy Bile duct resection Intraoperative transfusion	Bile duct resection/reconstruction Liver-associated procedures OR time >180 min EBL >1000 mL Tumor diameter >30 mm Portal node dissection Intraoperative transfusion En bloc diaphragm resection Extended hepatectomy
Benzoni et al ⁹	6	Major hepatectomy Left hepatectomy Trisegmentectomy Bisegmentectomy/left lobectomy	Segmentectomy/wedge Pringle >20 min Transfusion >600 mL Abscess Liver dysfunction Pleural effusion
Sadamori et al ⁸³	12.8	Repeat hepatectomy OR time >300 min EBL >2000 mL MVA Repeat hepatectomy OR time >300 min	Blood transfusion Duration of parenchymal transection
Nagano et al ⁸²	5.4	Age Cut surface area High-risk operation	
Okumura et al ⁸⁵	6.5	Fibrosis or cirrhosis OR time >5 h Major hepatic resection MVA OR time Resection of segment 4	Hepatectomy including Couinaud segment 4 or segment 5

Risk factors for bile leakage: Latest analysis of 10 102 hepatectomies for hepatocellular carcinoma from the Japanese national clinical database



Bile Leak Reduction with Laparoscopic Versus Open Liver Resection: A Multi-institutional Propensity Score-Adjusted Multivariable Regression Analysis

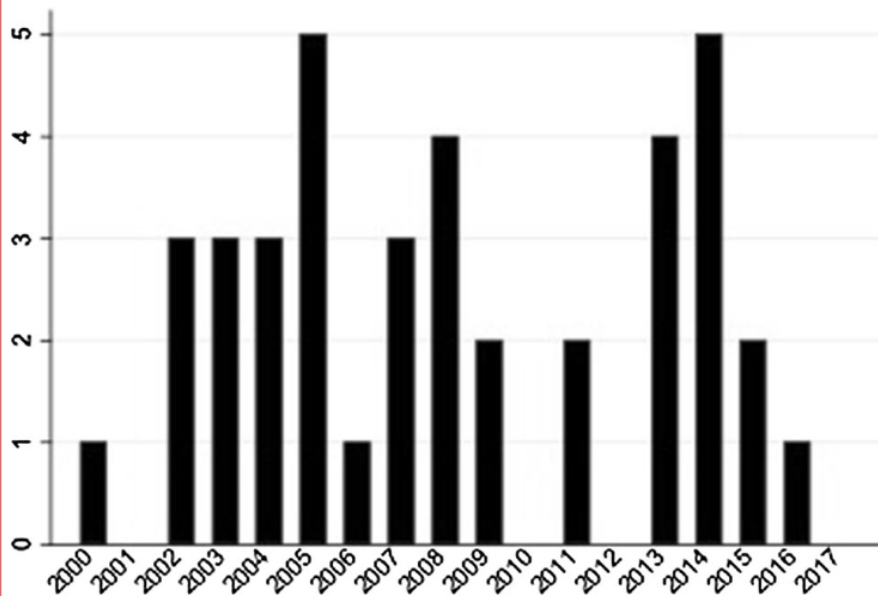


Fig. 1 Bile duct leak incidence per year, 2000–2017 ($N = 1388$)

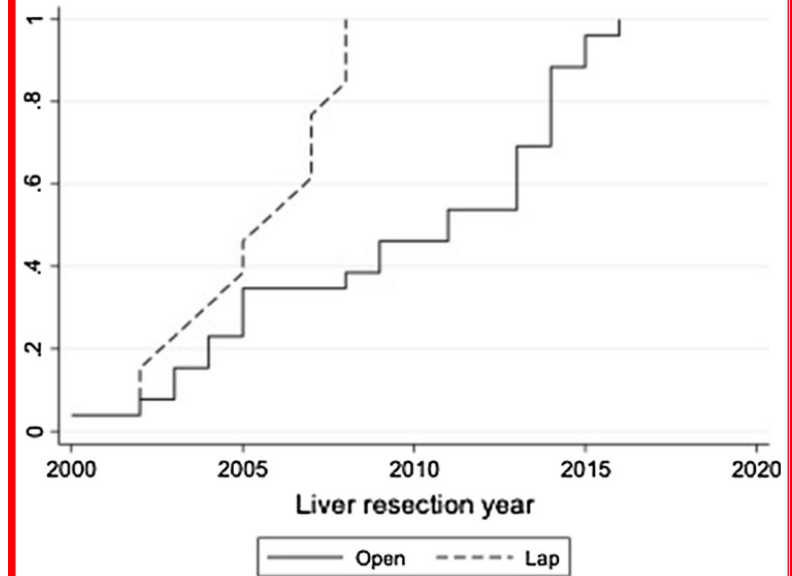


Fig. 3 Cumulative distribution of bile duct leaks by laparoscopy, 2000–2017 ($N = 1388$)

Μεγέθυνση Ιστών

Διαρκής Διεγχειρητική χρήση IOUS

Σύγχρονη τεχνολογία (*Ligaclip, Harmonic, Cavitron*)

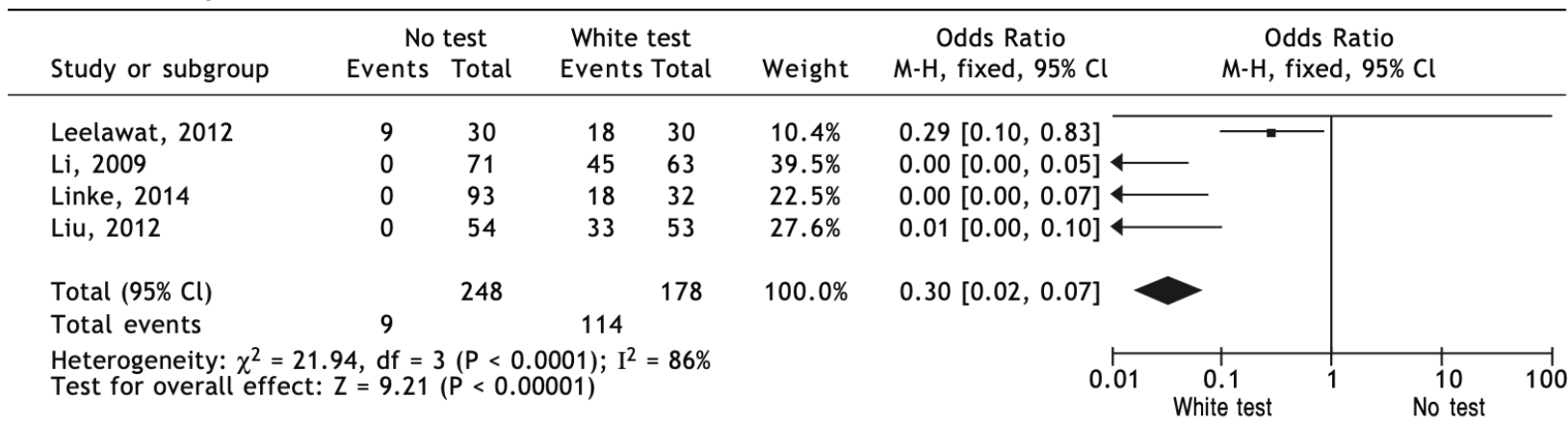
Smith AA, et al. World J Surg 2020

ΠΡΟΛΗΨΗ

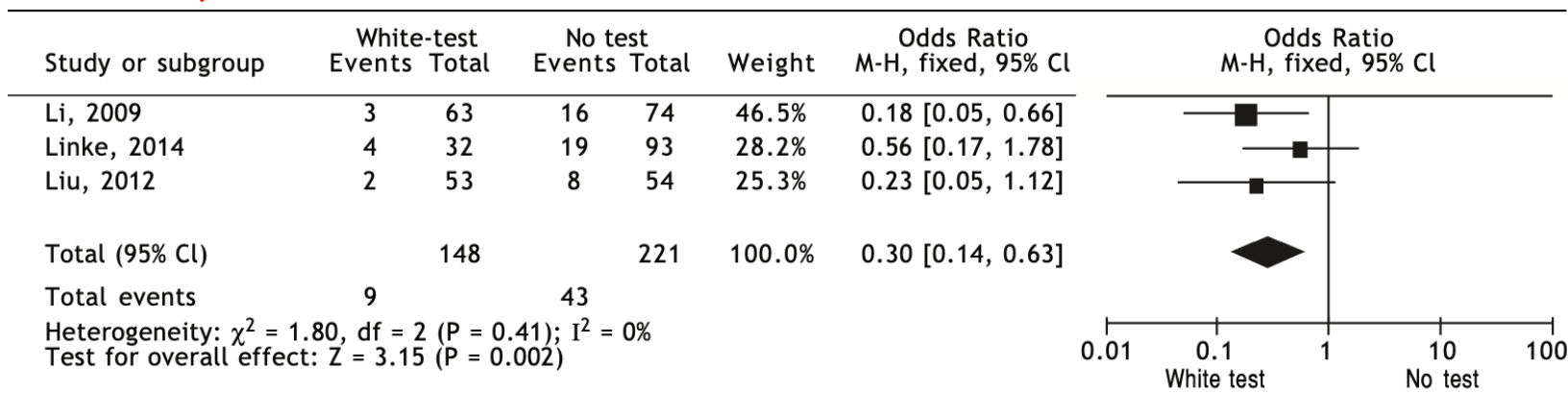
- Εγχειρητική Τεχνική
- White - test
- Bile - leakage test - ICG fluorescence
- C-tube / T-tube decompression
- Fibrin sealers
- Συστηματική Παροχέτευση

The White-test helps to reduce biliary leakage in liver resection: a systematic review and meta-analysis

Intra- Op Bile Leak



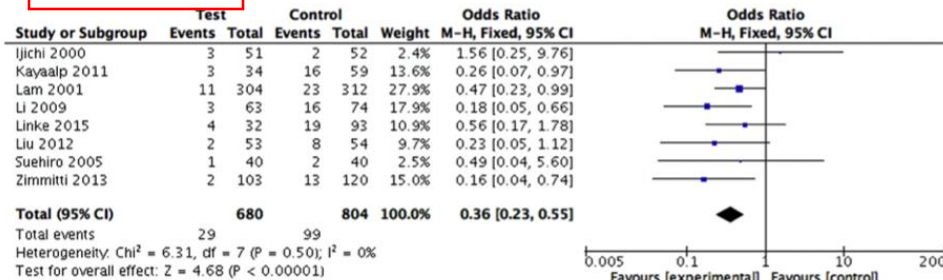
Post-Op Bile Leak



The role of bile leak testing in liver resection: a systematic review and meta-analysis

Ashish I. Vaska¹ & Saleh Abbas^{1,2}

Postoperative bile leak rate



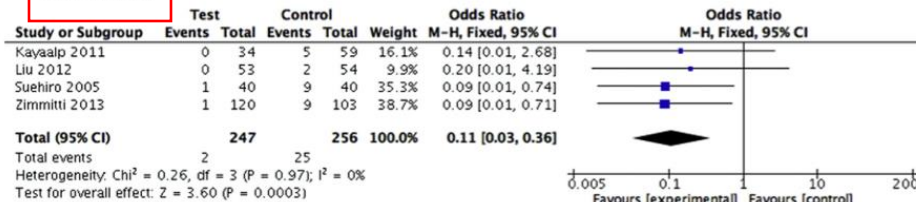
Normal saline

Betadine

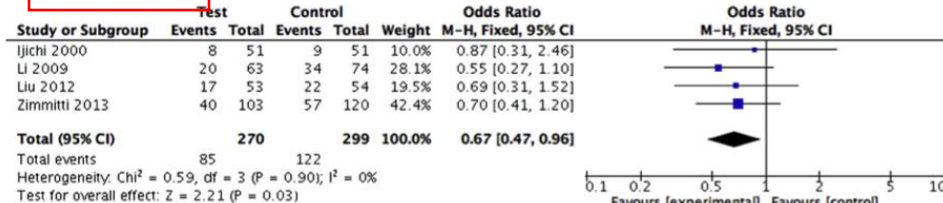
Methylene blue dye

Fat emulsion (white test)

Overall morbidity



Need for reintervention

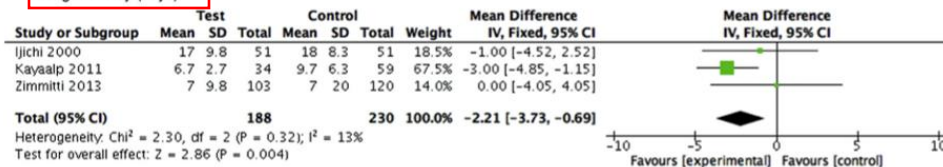


Indocyanine green ICG

Propofol

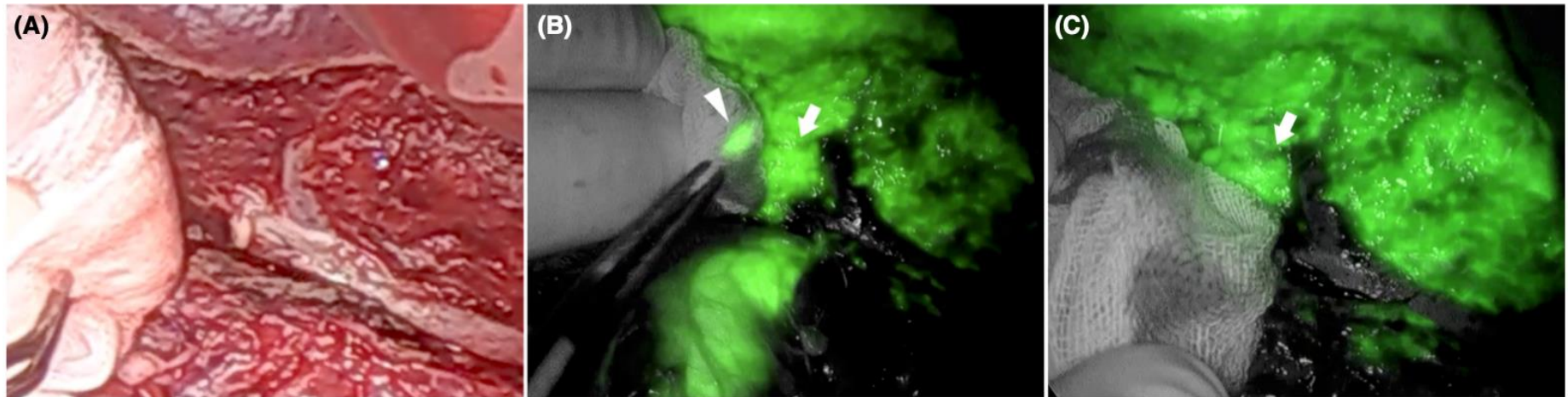
Air

Length of stay (days)



Hepatectomy guided by indocyanine green fluorescent imaging for visualizing bile leakage (with video)

Takehiko Hanaki  | Naruo Tokuyasu | Teruhisa Sakamoto | Yoshiyuki Fujiwara



IV ICG infusion (10mg/body)

ICG is taken up by the hepatocytes and gradually excreted in bile

Identification of bile duct system with ICG camera

Bile leakage as an ICG fluorescence contamination of the gauge

Prevents BLs

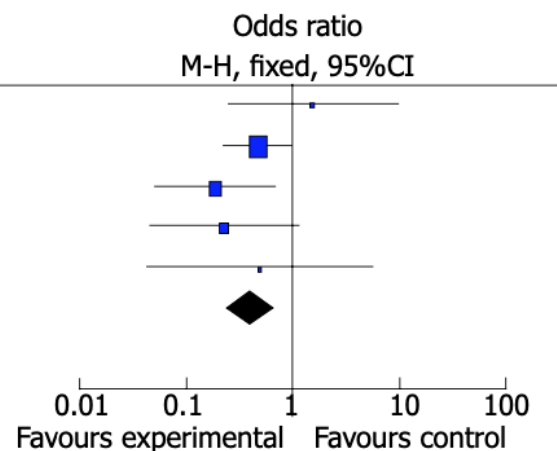
Beneficial for Peripheral type bile leakage!

Bile leakage test in liver resection: A systematic review and meta-analysis

Study or subject	BL test group		N-BL test group		Weight	Odds ratio M-H, fixed, 95%CI
Events	Total	Events	Total			
Ijichi 2000	3	51	2	51	4.0%	1.53 [0.24, 9.57]
Lam 2001	11	304	23	312	46.6%	0.47 [0.23, 0.99]
Li 2008	3	57	16	70	29.0%	0.19 [0.05, 0.68]
Liu 2012	2	53	8	54	16.2%	0.23 [0.05, 1.12]
Suehiro 2005	1	40	2	40	4.2%	0.49 [0.04, 5.60]
Total (95%CI)		505		527	100.0%	0.39 [0.23, 0.67]
Total events	20		51			

Heterogeneity: $\chi^2 = 4.11$, $df = 4$ ($P = 0.39$); $I^2 = 3\%$

Test for overall effect: $Z = 3.44$ ($P = 0.0006$)



Intraoperative positive bile leakage and postoperative bile leakage treatment

Study	Intraoperative bile leakage	Postoperative bile leakage	Conservative treatment (n)	Puncture drainage (n)	ENBD (n)	Reoperation (n)
Ijichi <i>et al</i> ^[11] , 2000	41.2% (21/51)	5.9% (3/51)	5	0	0	0
Liu <i>et al</i> ^[13] , 2012	62.3% (33/53)	3.8% (2/53)	7	0	2	1
Li <i>et al</i> ^[12] , 2009	71.4% (45/63)	5.3% (3/57)	No description	No description	1	2
Suehiro <i>et al</i> ^[24] , 2005	No description	2.5% (1/40)	No description	No description	No description	No description
Lam <i>et al</i> ^[2] , 2001	19.7% (60/304)	3.6% (11/304)	7	11	6	10
Leelawat <i>et al</i> ^[26] , 2011	63.3% (19/30)	No description	2	0	0	0
Sakaguchi <i>et al</i> ^[27] , 2010	29.6% (8/27)	0% (0/27)	2	0	0	0
Kaibori <i>et al</i> ^[25] , 2011	80.8% (42/52)	0% (0/52)	2	1	2	0
Total	39.3% (228/580)	3.42% (20/584)	25 (41.0%)	12 (19.8%)	11 (18.0%)	13 (21.2%)

Does the placement of a cystic duct tube after a hepatic resection help reduce the incidence of post-operative bile leak?

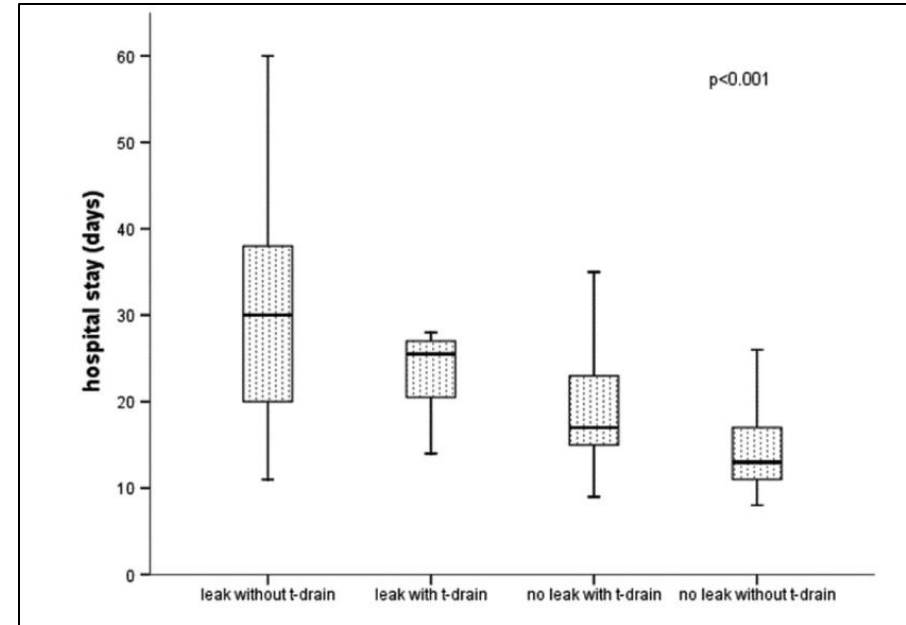
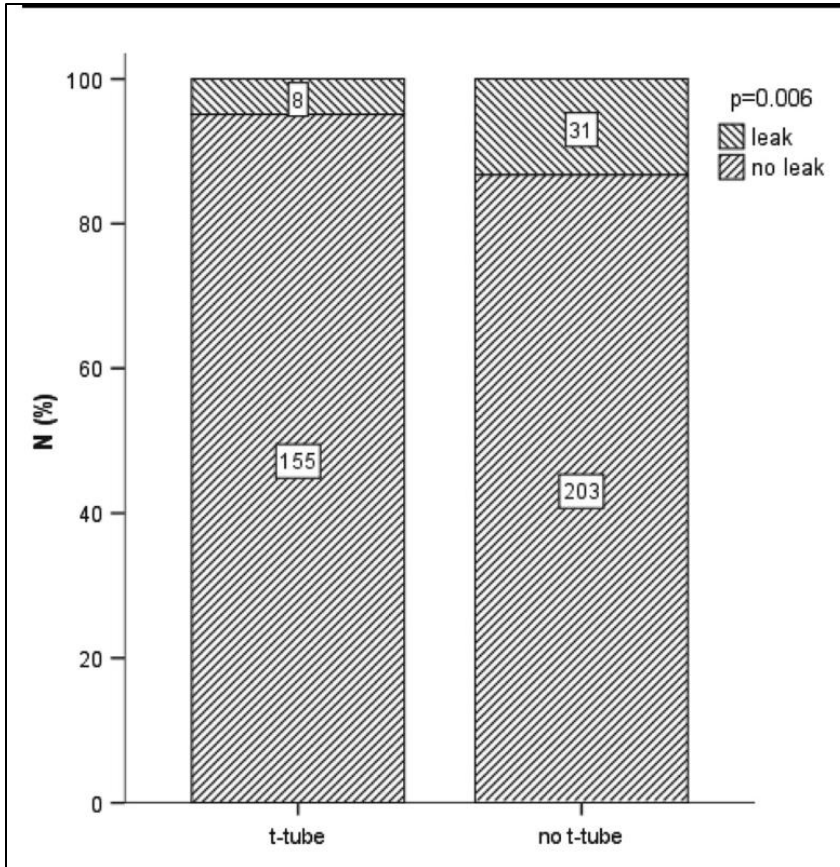
Variables	Post-operative bile leak	
	Risk ratio (95%CI)	P-value
Use of a C tube		
Yes	1.071 (0.397–2.885)	0.893
No	1	

	C tube (-) [n = 467]	C tube (+) [n = 83]	P-value
Age	66 (17–83)	68 (37–87)	0.554
Gender			0.579
Males	327 (86%)	55 (83%)	
Females	140 (14%)	28 (17%)	
Side of hepatectomy			0.0891
Right	262 (56%)	54 (65%)	
Left	205 (44%)	29 (35%)	
Partial hepatectomy			
Bile leak negative	173	9	1.000
Bile leak positive	4	0	
Group A/ B/ C ^a	2/2/0	0/0/0	0.744
Segmentectomy or sectionectomy			
Bile leak negative	123	32	0.448
Bile leak positive	10	5	
Group A/ B/ C ^a	2/8/0	2/3/0	0.560
Hemi-hepatectomy and more hepatectomy			
Bile leak negative	134	35	0.175
Bile leak positive	23	2	
Group A/ B/ C ^a	4/18/5	2/0/0	0.016

	C tube (-)	C tube (+)	P value
Hospital stay after a hepatectomy			0.454
<30 days	4	1	
30–60 days	26	6	
60–90 days	4	0	
>90 days	3 (113, 116, 152 days)	0	

C – Tube
Cannot prevent “isolated” bile leakage!!!

T-drain reduces the incidence of biliary leakage after liver resection



**No inspection of Peripheral leaks!!!
Attention to CHD Damage!!!**

Application of Fibrin Glue Sealant After Hepatectomy Does Not Seem Justified

Results of a Randomized Study in 300 Patients

Postoperative Outcome of Patients in the Fibrin Glue and Control Groups			
	Fibrin Glue (n = 150)	Control (n = 150)	P
Postoperative hospital stay (days) (mean $\pm$ SD)	13.3 $\pm$ 13	12.6 $\pm$ 9	0.57
Patients with postoperative transfusion (%)	27 (18)	19 (12)	0.2
Postoperative transfusion PRBCU (mean $\pm$ SD)	0.15 $\pm$ 0.66	0.17 $\pm$ 0.63	0.72
Overall transfusion (PRBCU) [no. (%) of patients]	40 (27)	29 (19)	0.14
Overall transfusion (mean $\pm$ SD)	0.3 $\pm$ 0.74	0.31 $\pm$ 0.53	0.26
Days of postoperative drainage (mean $\pm$ SD)	7.9 $\pm$ 5	7.13 $\pm$ 47	0.94
Overall drainage volume (mL) (mean $\pm$ SD)	1180 $\pm$ 2528	960 $\pm$ 1253	0.34
Characteristics of fluid from drainage (%)			
Serous	133 (89)	130 (87)	0.64
Hematic	6 (4)	7 (5)	
Bile	11 (7)	13 (8)	

Abdominal drainage is contraindicated after uncomplicated hepatectomy: Results of a meta-analysis of randomized controlled trials

7 PRTs

1064 pts undergoing hepatectomy

533 with 531 without an abdominal drain

Drainage was associated with higher overall complications (RR: 1.37, $P=.0003$)
and wound-related complications (RR:2.29, $P=.01$)

NO SIGNIFICANT DIFFERENCE in:

Bile leak

Intraabdominal collections

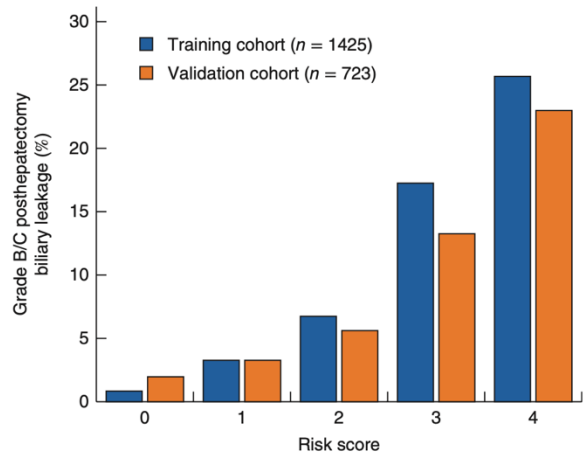
Collections requiring interventions

LOS

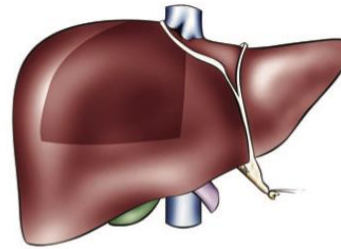
Risk score to predict biliary leakage after elective liver resection

K. Mohkam¹ , O. Farges³, E. Vibert⁴, O. Soubrane³, R. Adam⁴, F.-R. Pruvot⁵, J.-M. Regimbeau⁶ , M. Adham², E. Boleslawski⁵ and J.-Y. Mabrut¹ , on behalf of the Association de Chirurgie Hépatobiliaire et de Transplantation (ACHBT) French Hepatectomy Study Group*

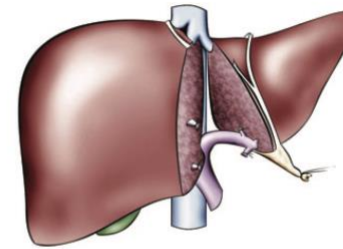
2393 LRs
No HJ
2012-2015



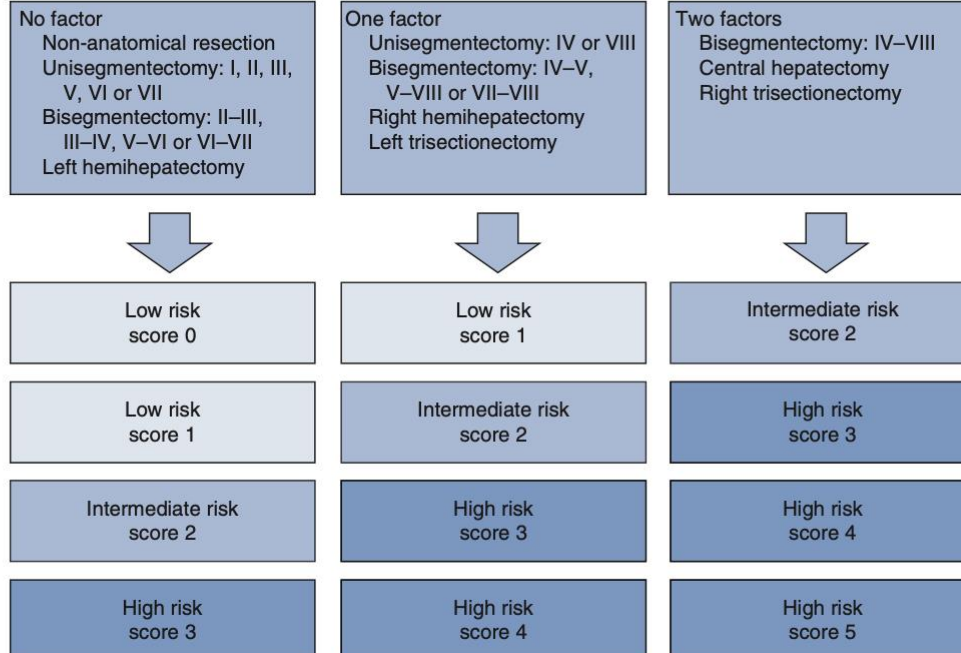
Factors related to type of hepatectomy



Anatomical resection including segment VIII



Transection along right aspect of left intersegmental plane



Operative factors
Blood loss ≥ 500 ml
Remnant ischaemia ≥ 45 min
ALPPS

No factor
One factor
Two factors
Three factors

ΑΝΤΙΜΕΤΩΠΙΣΗ

- Συντηρητική

1. Παροχέτευση - Αντιβίωση
2. Wait-and-see strategy (success rate 75%)

- Επεμβατική

1. Συριγγογραφία – MRCP - scintigraphy
2. ERCP $\pm$ stent or ENBD
3. PTC $\pm$ PTBD
4. Rendezvous technique
5. Alcohol-ablation / Fibrin glue sealing
6. PVE - TAE

- Χειρουργική Επανεπέμβαση

Bile leak after hepatectomy: Predictive factors of spontaneous healing

Drainage Output on Day 10 >100ml

Vigano L, et al. The Am J Surg 2008

Incidence and Management of Bile Leakage after Hepatic Resection for Malignant Hepatic Tumors

“Central” type bile leaks

Drainage >2 wks

Poor QoL

Tanaka S, et al. J Am Coll Surg 2002

Endoscopic management is the treatment of choice for bile leaks after liver resection

Alexander Dechêne, MD,¹ Christoph Jochum, MD,¹ Christian Fingas, MD,^{1,2} Andreas Paul, MD, Prof.,² Dominik Heider, PhD,³ Wing-Kin Syn, MBChB, Prof.,⁴ Guido Gerken, MD, Prof.,¹ Ali Canbay, MD, Prof.,^{*,1} Thomas Zöpfl, MD^{*,1,5}

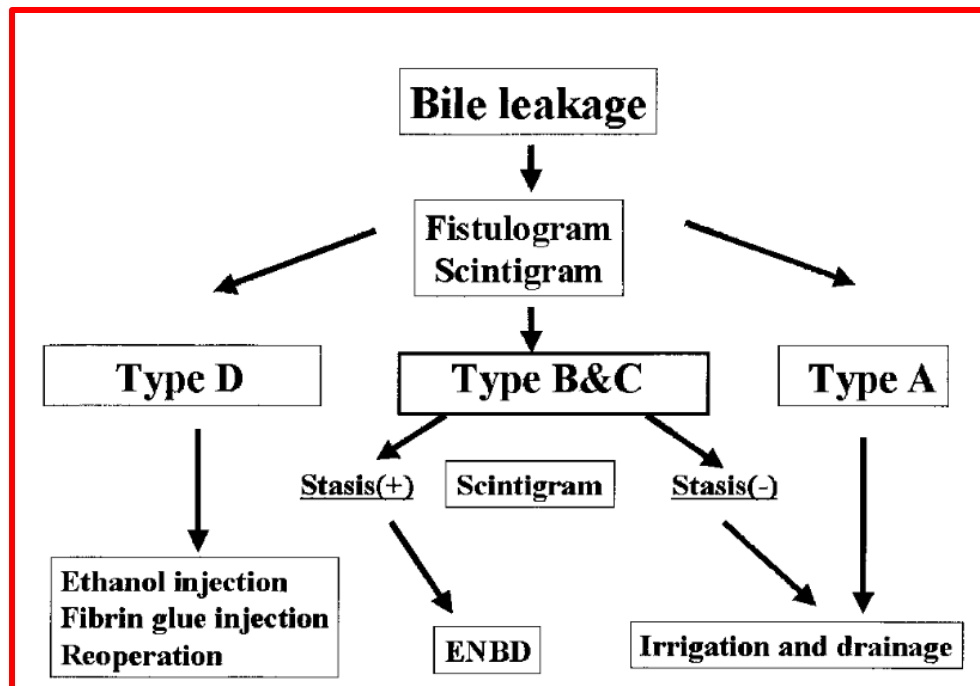
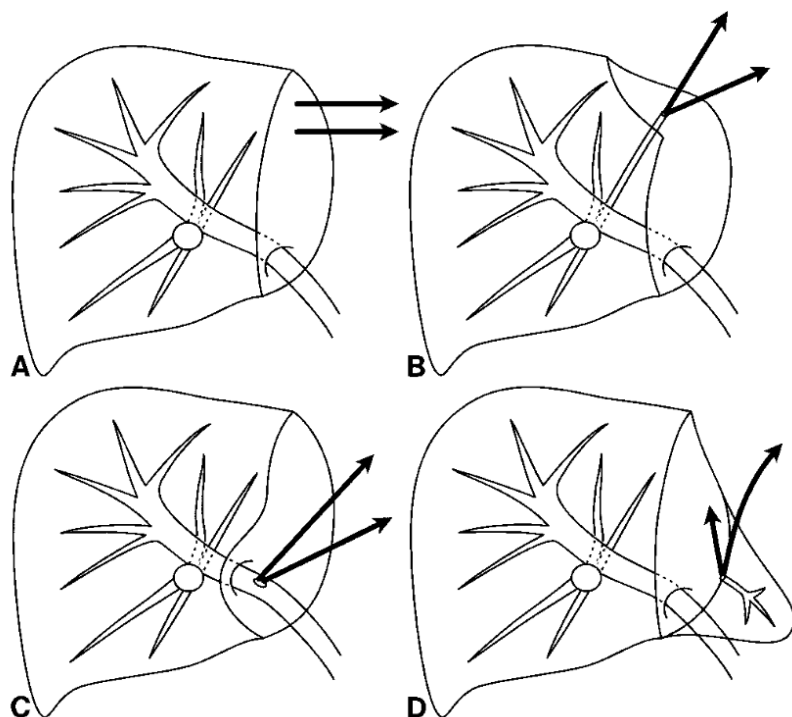
Essen, Karlsruhe, Germany; London, United Kingdom

ERCP + stent **ALWAYS!!!**

7-10 days

Gastrointest Endosc 2015

Risk Factors and Management of Bile Leakage after Hepatic Resection

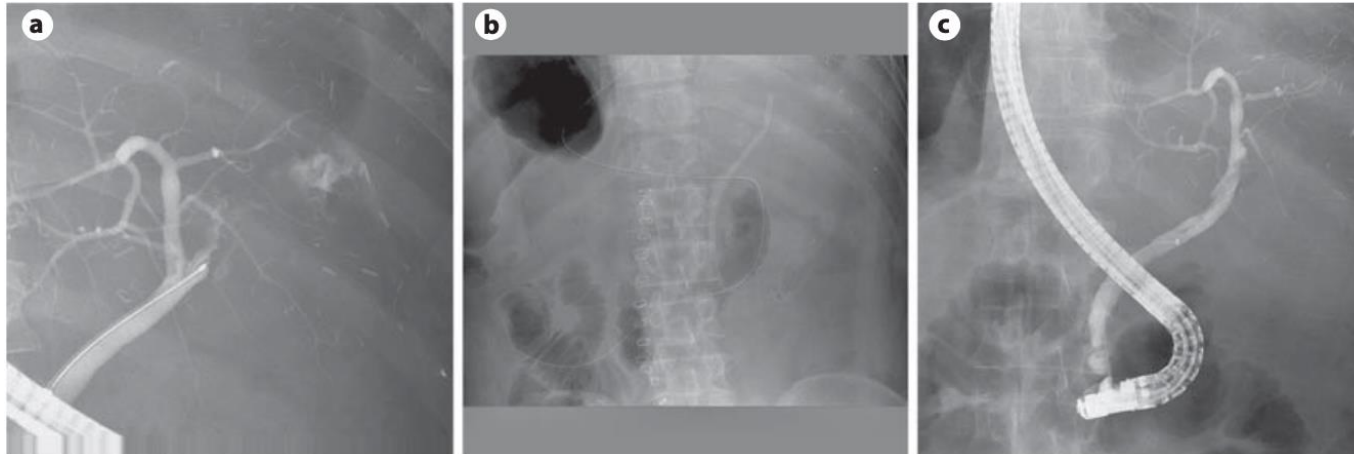


Fistulographic findings and outcome.

Presence of bile duct	No. (n = 11)	Duration of treatment after hepatectomy (days)
Negative	5	37.8
Positive	6	91.3
Proximal bile duct	5	102.6
Bile drainage (+)	2	30.0
Bile drainage (-)	3	179.2
Distal bile duct (+)	1	80.0 (Operation)

Posthepatectomy Bile Leakage: How to Manage

Hoekstra L, et al. Dig Surg 2012



R Hepatectomy
Stent
6 wks

Treatment bile leakage	Without HJ (n = 10)	With HJ (n = 9)
Percutaneous radiological drainage	6 (60%)	9 (100%)
Endoscopic drainage	3 (30%)	0
Conservative treatment	0	0
Sutured	1 (10%)	0

Diagnosis	Initial treatment	Complication	Relaparotomy
Colorectal metastases	Percutaneous drainage	Subphrenic abscess	Drainage
Focal nodular hyperplasia	ERCP	Persistent bile leakage	HJ
Choledochal cyst	ERCP	Persistent bile leakage	Right hemihepatectomy
Traumatic laceration	Suture	Persistent bile leakage	Left hemihepatectomy

Location of a biliary leak after liver resection determines success of endoscopic treatment



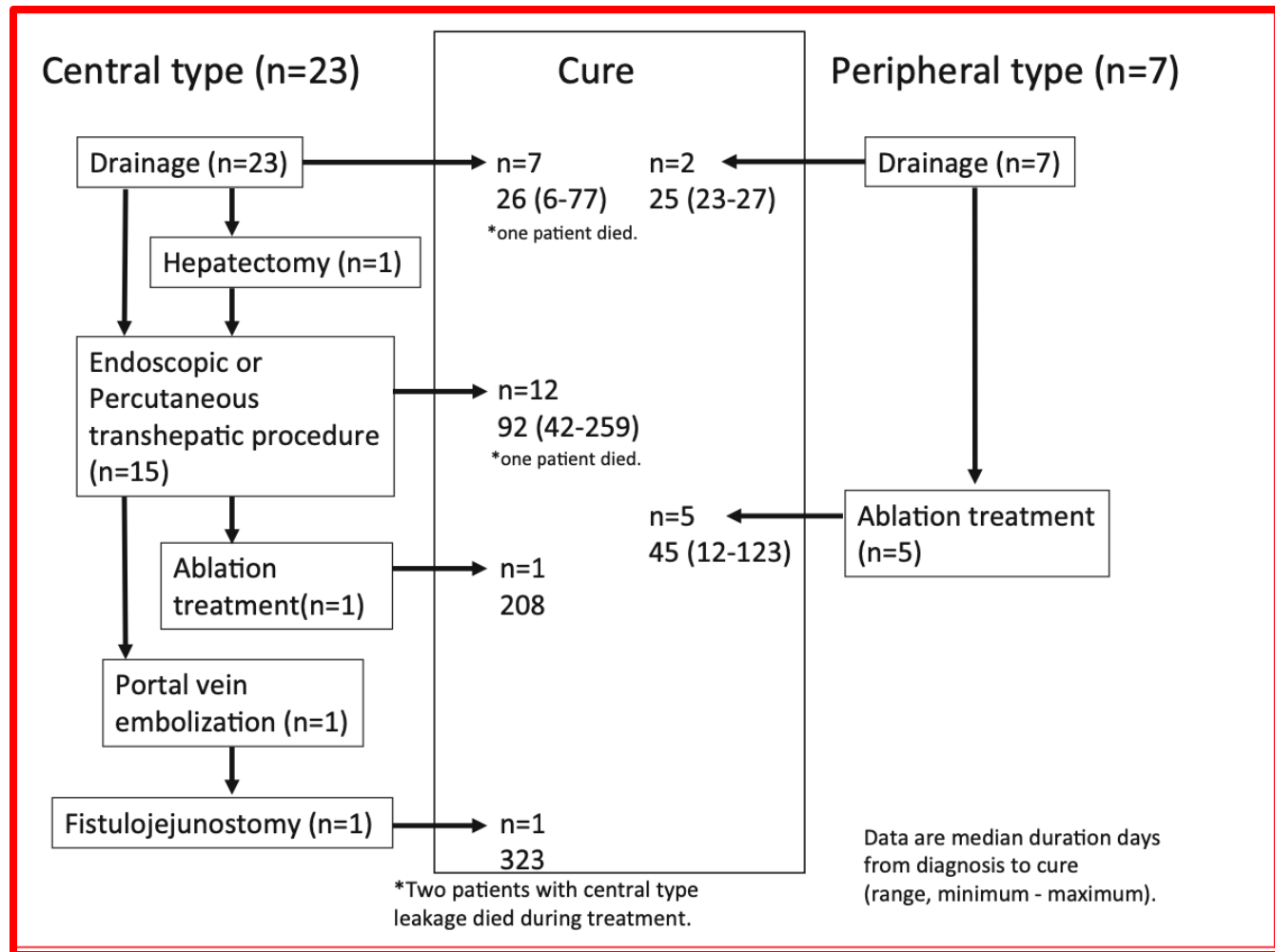
"Central" type bile leaks: Success rate 70-80%

Location of a biliary leak after liver resection determines success of endoscopic treatment

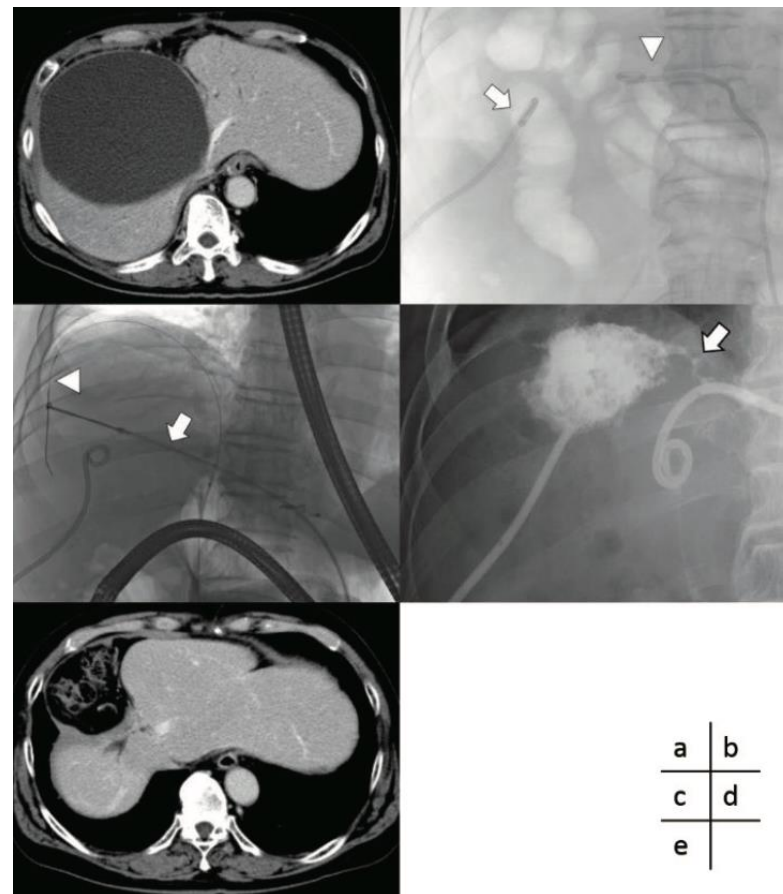
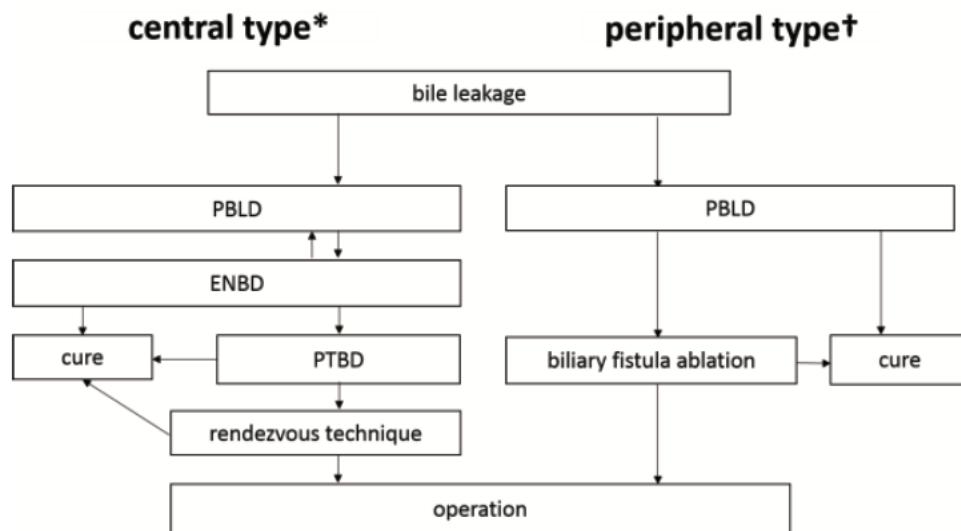


"Peripheral" type bile leaks: Success rate <50%

Risk Factors and Managements of Bile Leakage After Hepatectomy



Non-Surgical Management of Bile Leakage After Hepatectomy: A Single-Center Study



a	b
c	d
e	

Central type: 315 days

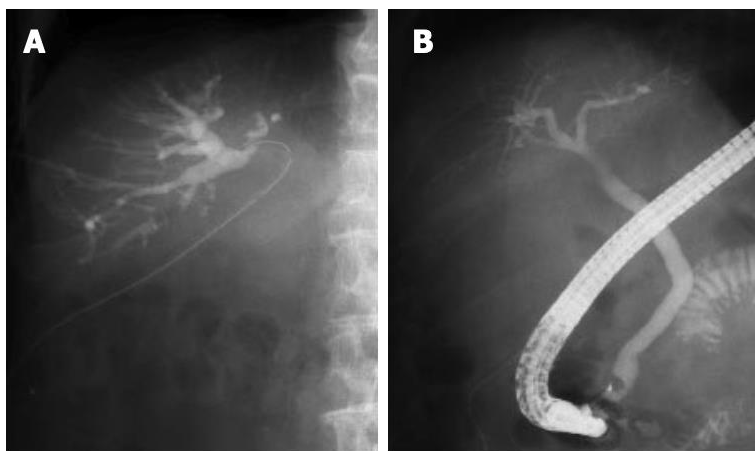
Peripheral type: 156 days

Postoperative bile leakage managed successfully by intrahepatic biliary ablation with ethanol

HCC
6.3×5.9cm
S1, S5-S7



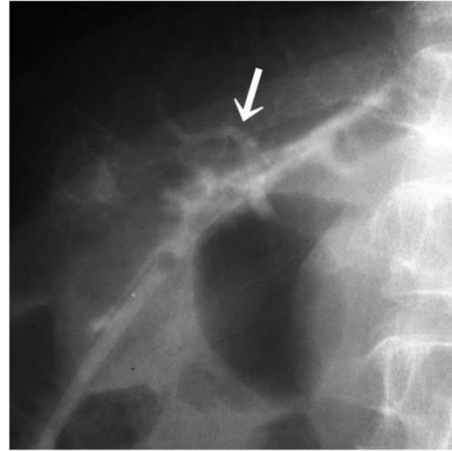
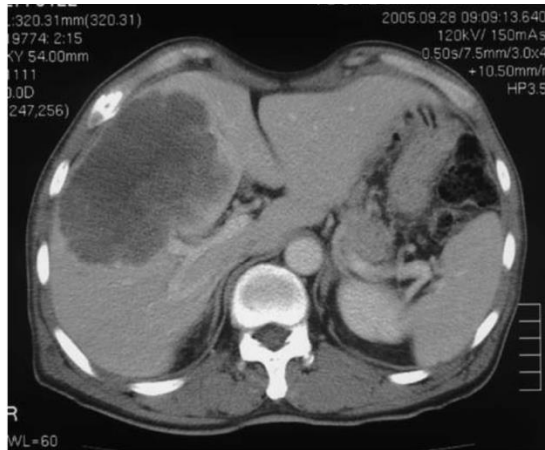
R Ant Bile duct



5-Fr balloon occlusion cath
4-ml pure alcohol injection 5 times/ wk
Sealing after 25 sessions / 9 months after surgery

Percutaneous Transhepatic Portal Embolization for Persistent Bile Leakage After Hepatic Resection: Report of a Case

YOSHIHIKO SADAKARI¹, ATSUSHI MIYOSHI¹, TAKAO OHTSUKA¹, NAOHIKO KOHYA¹, TOMOHIDE TAKAHASHI¹, KOICHI MATSUMOTO², and KOHJI MIYAZAKI¹



Successful treatment of isolated bile leakage after hepatectomy combination therapy with percutaneous transhepatic portal embolization and bile duct ablation with ethanol: a case report

Surg Case Reports 2018

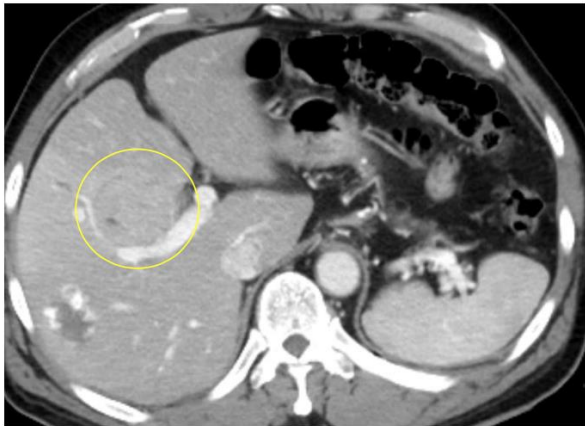
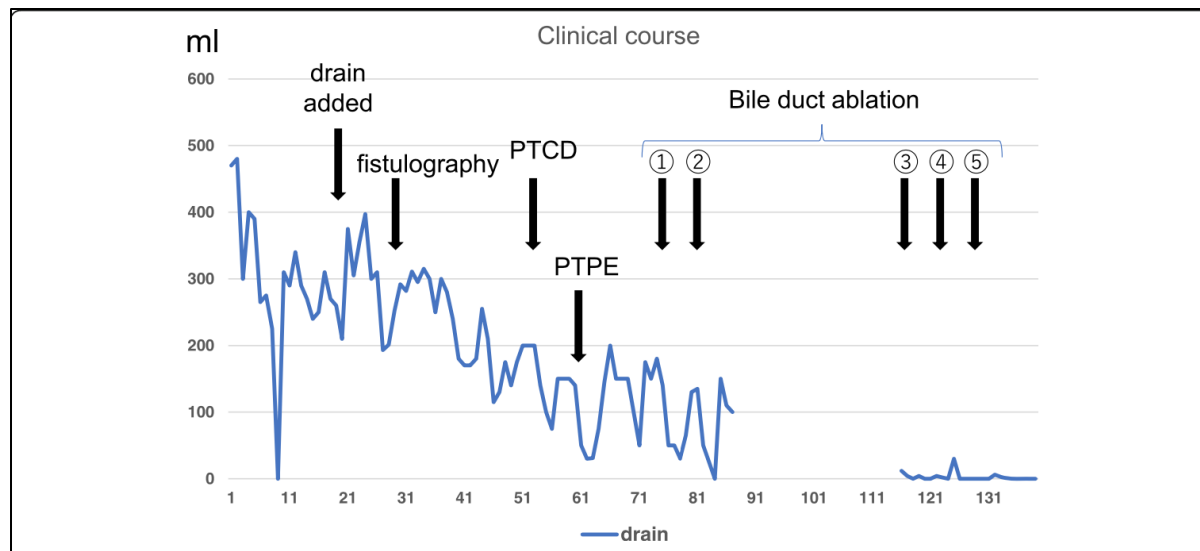
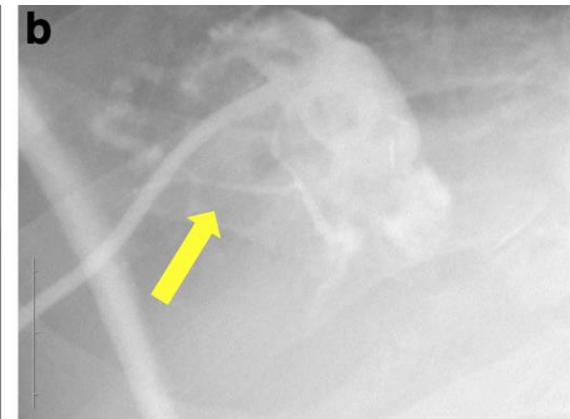
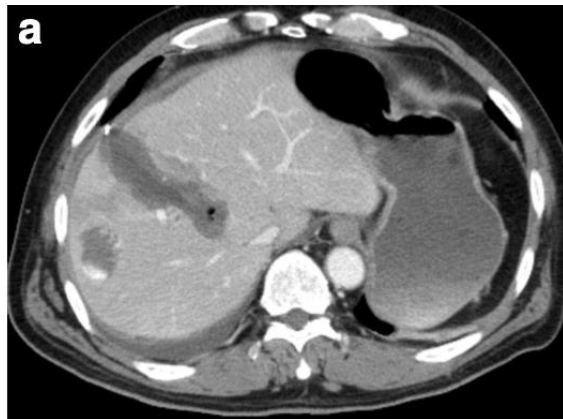
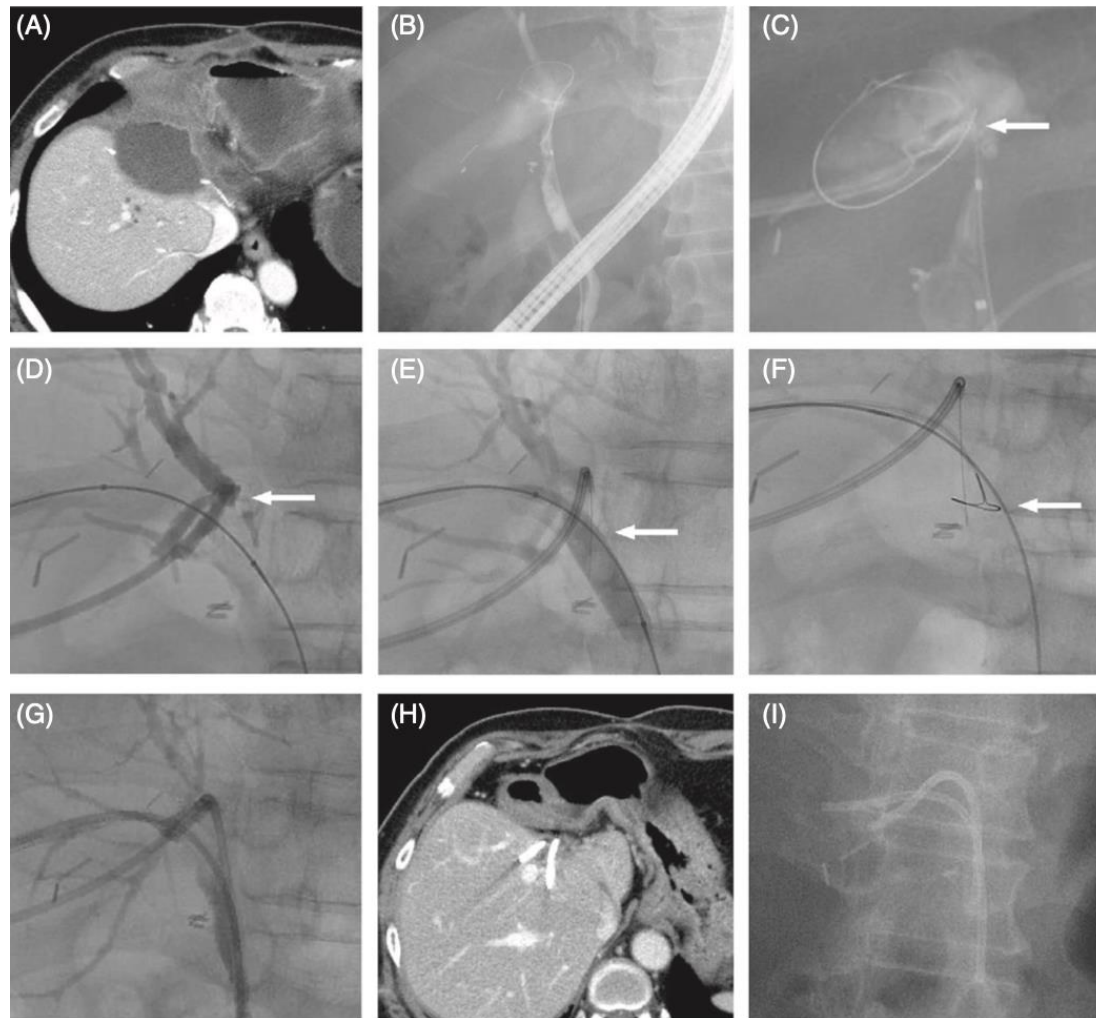


Fig. 1 Preoperative abdominal computed tomography showed a 50 × 48 mm hepatic mass that pressed the anterior Glisson



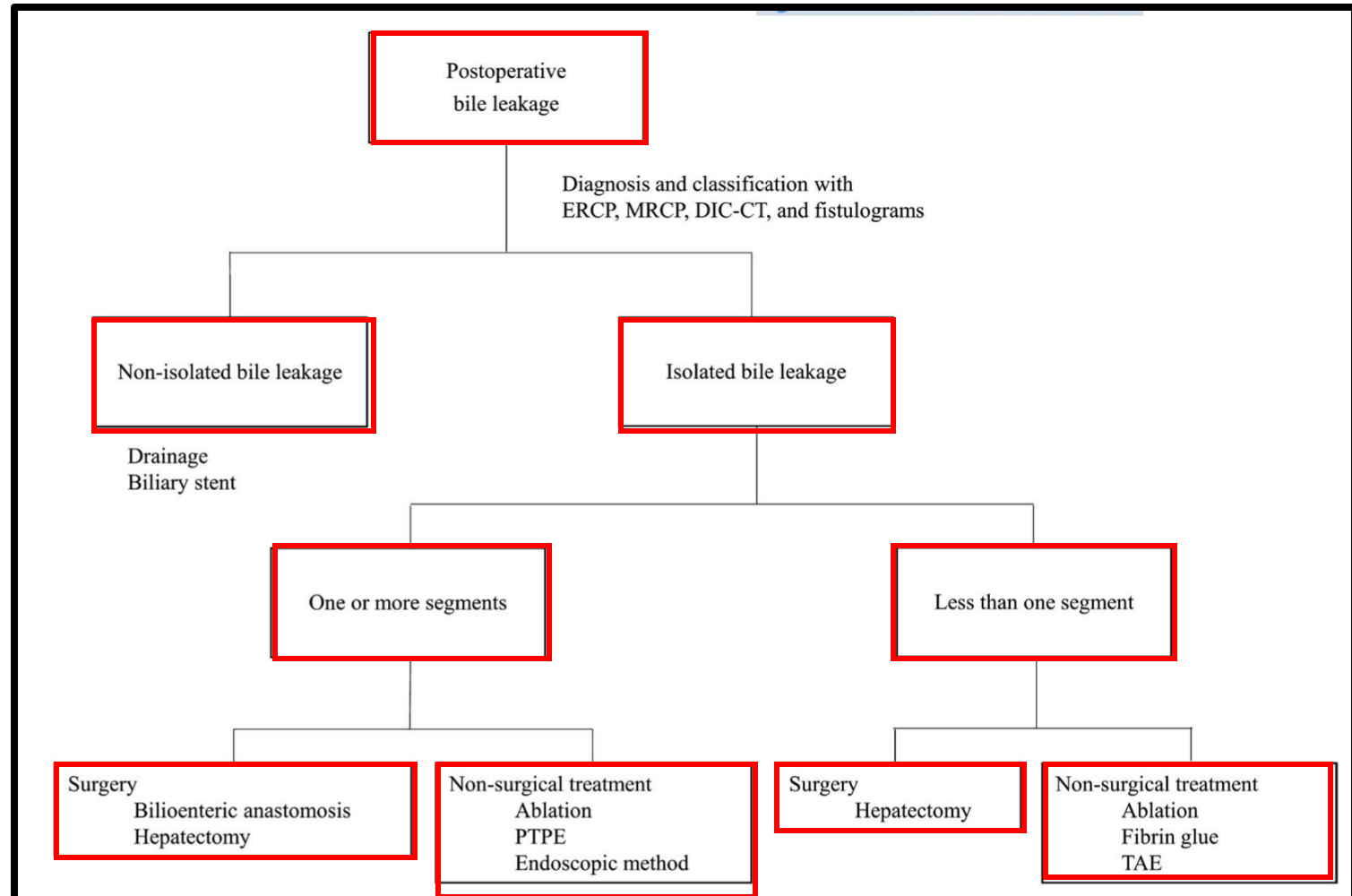
Sharp recanalization and rendezvous technique for biliary occlusions due to spacious bile leak after hepatectomy



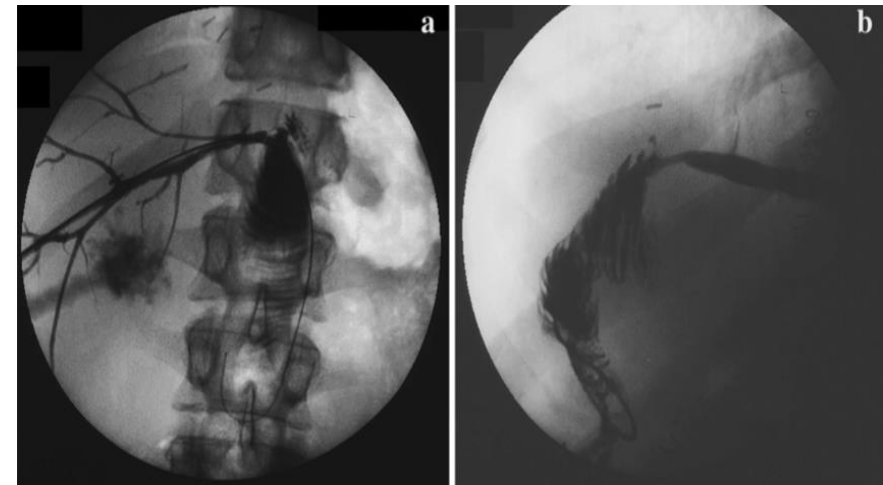
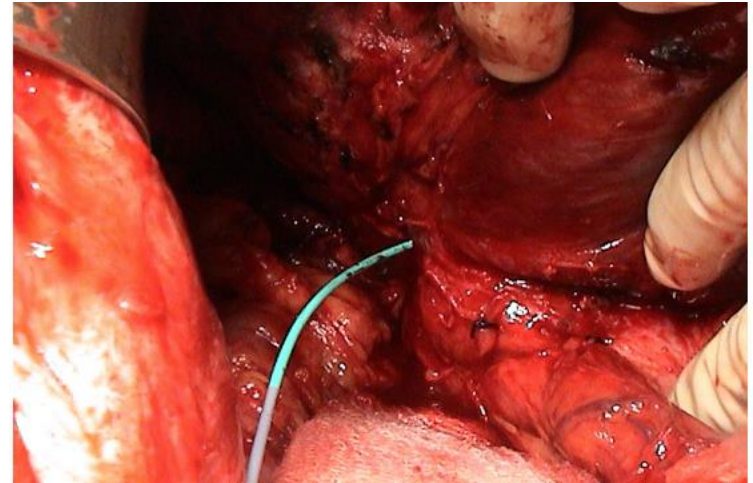
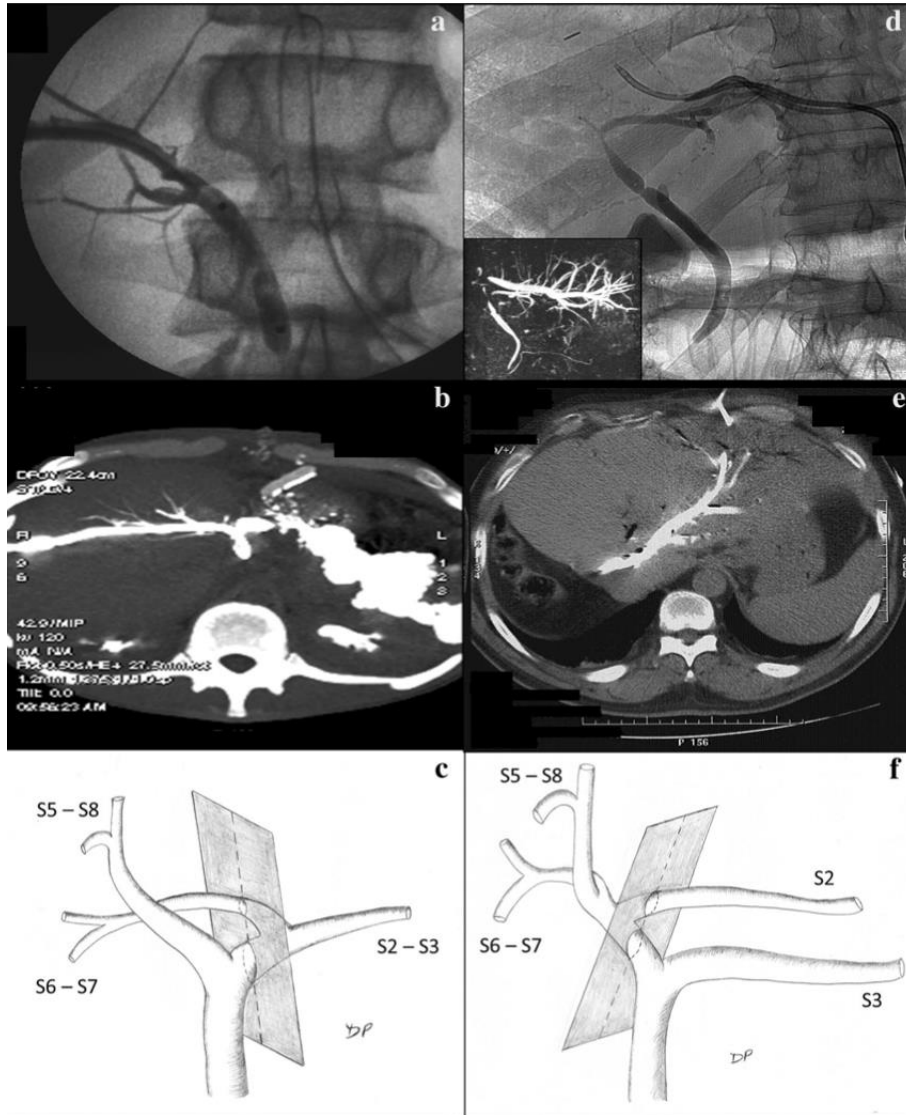
Treatment strategy for isolated bile leakage after hepatectomy: Literature review

Norio Kubo  | Ken Shirabe

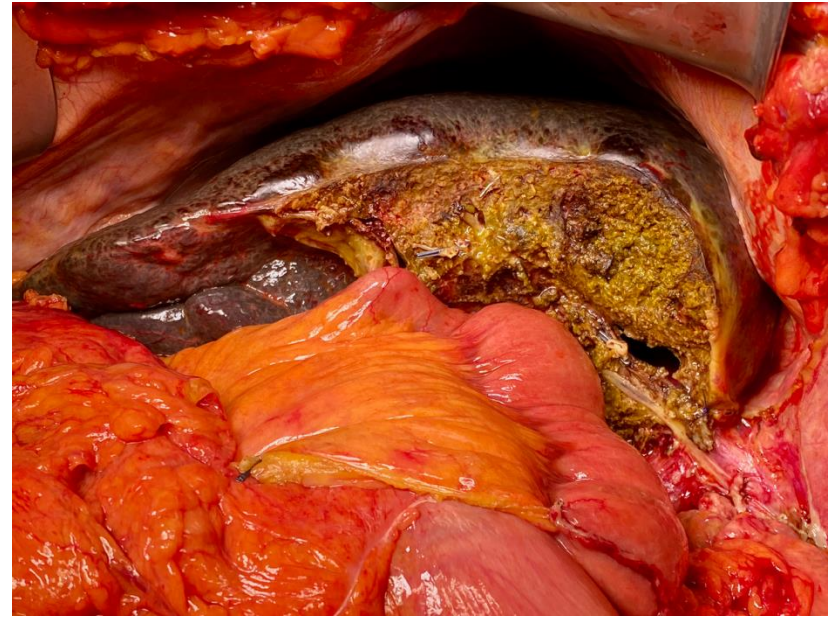
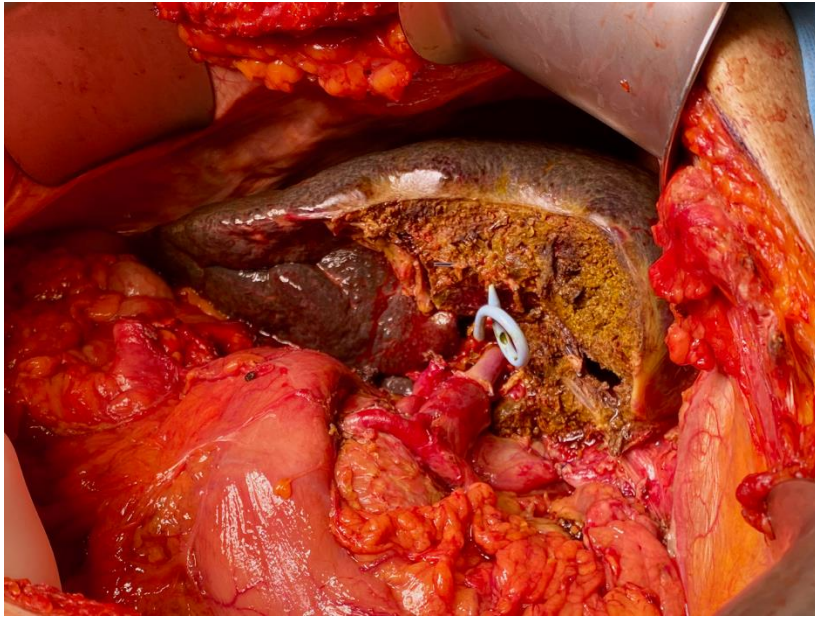
Ann Gastroenterol Surg 2020



Excluded segmental duct bile leakage: the case for bilio-enteric anastomosis



Surgical Treatment of IBL



Incidence and risk factors for anastomotic bile leakage in hepatic resection with bilioenteric reconstruction - A international multicenter study

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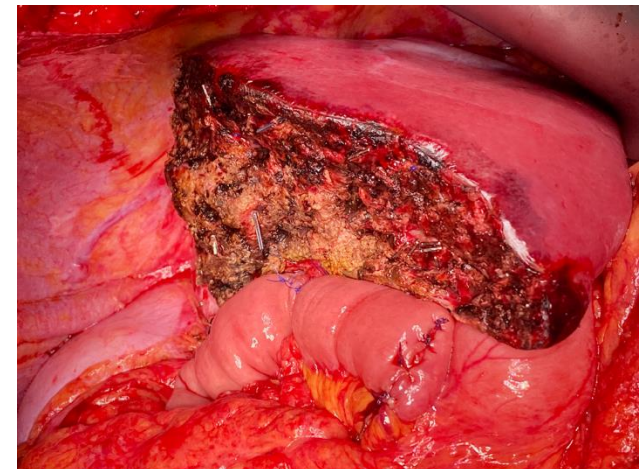
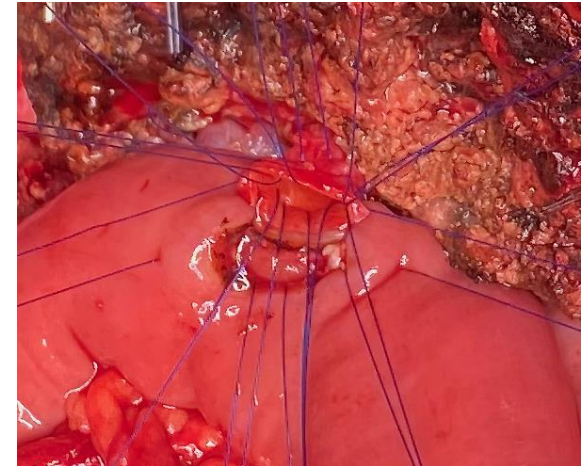
11 Academic Institutions in Europe
921 pts

AL 5.4%
30d mortality 9.6%

-Risk factors for B/C bile leak:
Pringle
Postop external biliary drainage
Abdominal drainage
-Protective factors and TREATMENT
PTC-PTBD

AL + ENBD	76.5%
AL + PTBD	17.6%
AL + PTBD + stent	5.9% ($p < .001$)

AL → liver failure, cholangitis,
hemorrhage, sepsis



D. KORKOLIS, MD

Incidence and risk factors for anastomotic bile leakage in hepatic resection with bilioenteric reconstruction - A international multicenter study

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D. KORKOLIS, MD

ΣΥΜΠΕΡΑΣΜΑ

- Η χολόρροια μετά ηπατεκτομή παραμένει συχνή επιπλοκή
- Σήψη, ηπατική ανεπάρκεια, θνητότης
- Προγνωστικά μοντέλα
- Πρόληψη
- Αυτόματη επούλωση
- Συντηρητική αντιμετώπιση: Πρώτη Επιλογή!!!
- Ενδοσκοπική Παροχέτευση
- Διαδερμική Διηπατική προσέγγιση
- Έγχυση αλκοόλης – Fibrin sealant – PVE
- Χειρουργική επέμβαση