GI Malignancy – Surgical Option

Pancreas, Colorectal, Anal Canal

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31st AROI-ICRO, Faridkot, April 27-28th, 2019



GI Malignancy – Surgical Options

• Pancreas

- Standard resection pancreaticoduodenectomy
- Boderline resectable disease options

Colorectal

- Colon:
 - Standard colectomy
 - Complete mesocolic excision
 - Multivisceral resection

GI Malignancy – Surgical Options



- Colorectal
 - Rectum:
 - Total Mesorectal excision (TME)
 - Sphincter Preservation
 - Abdomino perineal resection (APR)
 - Extralevator APR
 - Rectal resection Beyond TME

- Colorectal peritoneal metastasis - CRS+HIPEC



Pancreas – Surgical Options

- Very poor prognosis, 5 yrs survival 6%
- Late stage of presentation
- Only 20 % are eligible for initial resection
- 5 yrs survival of R0 resected patients 25%

Tumour biology of pancreatic cancer contributes to early recurrence and metastasis, and resistance to chemotherapy / radiotherapy



PANCREATIC CANCER

Resectable

Boderline resectable

Unresectable



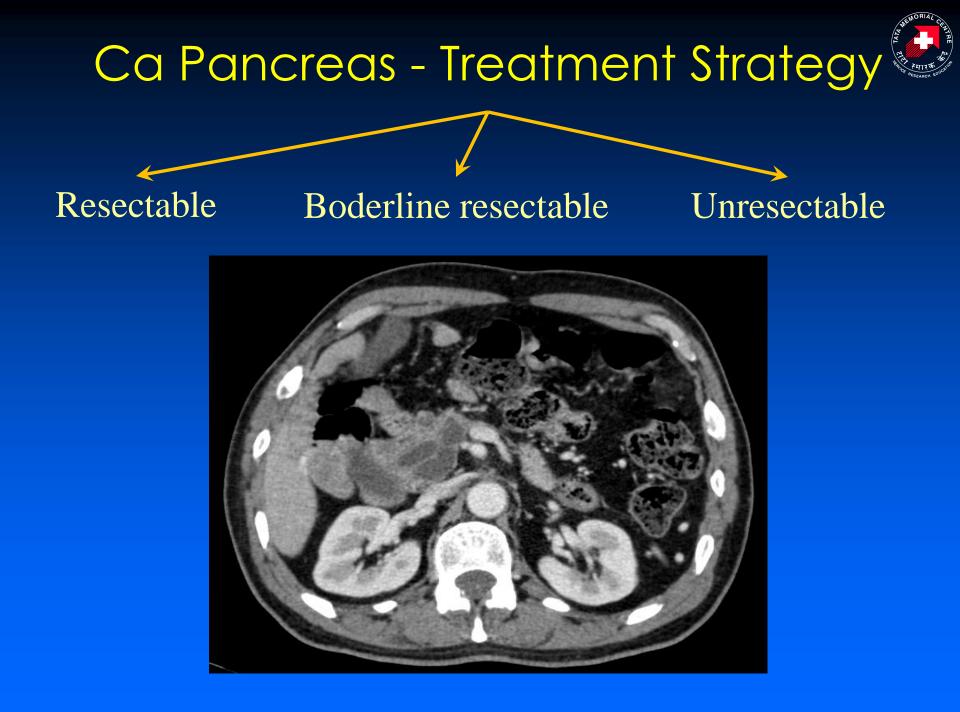
Boderline resectable

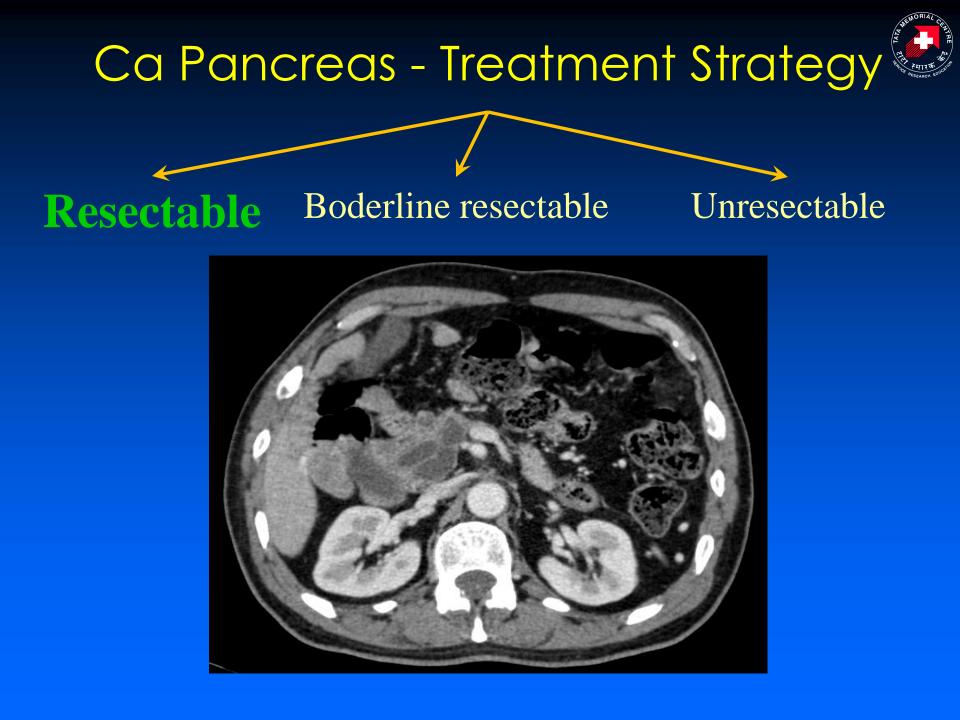
- For tumors of the head or uncinate process. Solid tumor contact
 - With the **SMV or portal vein of >180 degrees**, with vein deformity / thrombosis but reconstructable.
 - With the inferior vena cava.
 - With the common hepatic artery without extension to the celiac axis or hepatic artery bifurcation, allowing for safe and complete resection and reconstruction.
 - With the SMA ≤ 180 degrees.
 - With variable anatomy (eg, accessory right hepatic artery, replaced right hepatic artery, replaced common hepatic artery, and the origin of replaced or accessory artery).

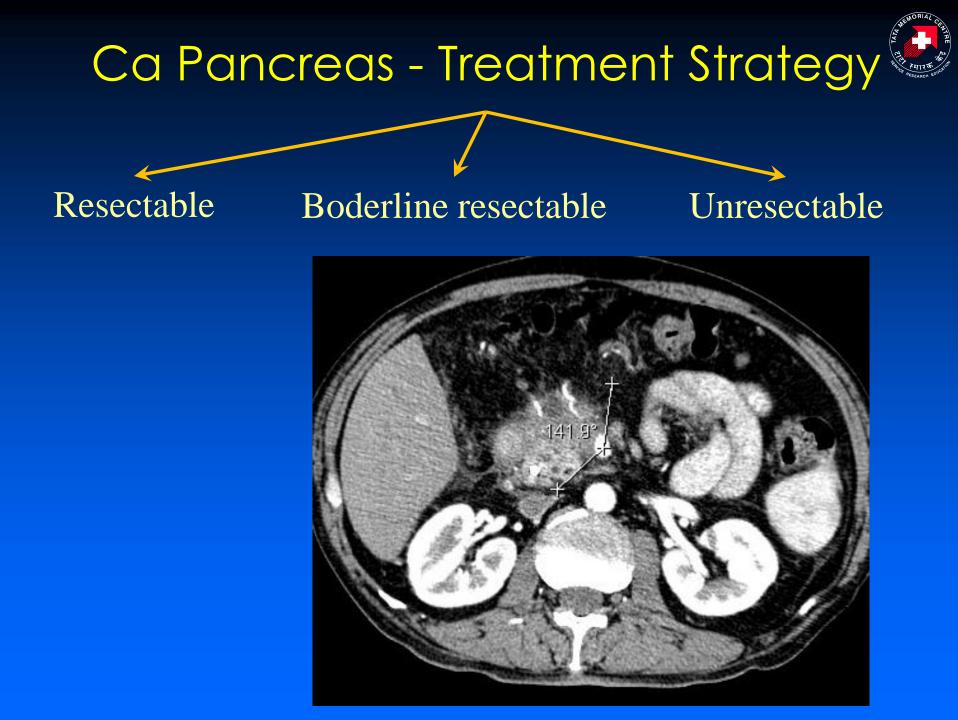
• For tumors of the body/tail: Solid tumor contact

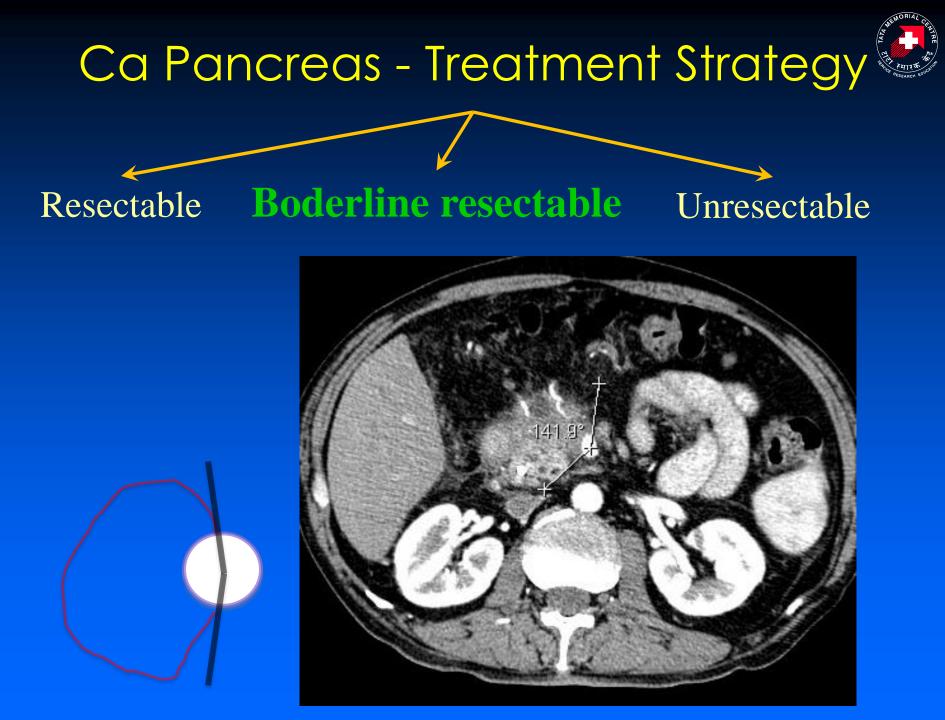
- With the celiac axis of ≤ 180 degrees.
- With the celiac axis >180 degrees without involvement of the aorta and with an intact and uninvolved gastroduodenal artery.

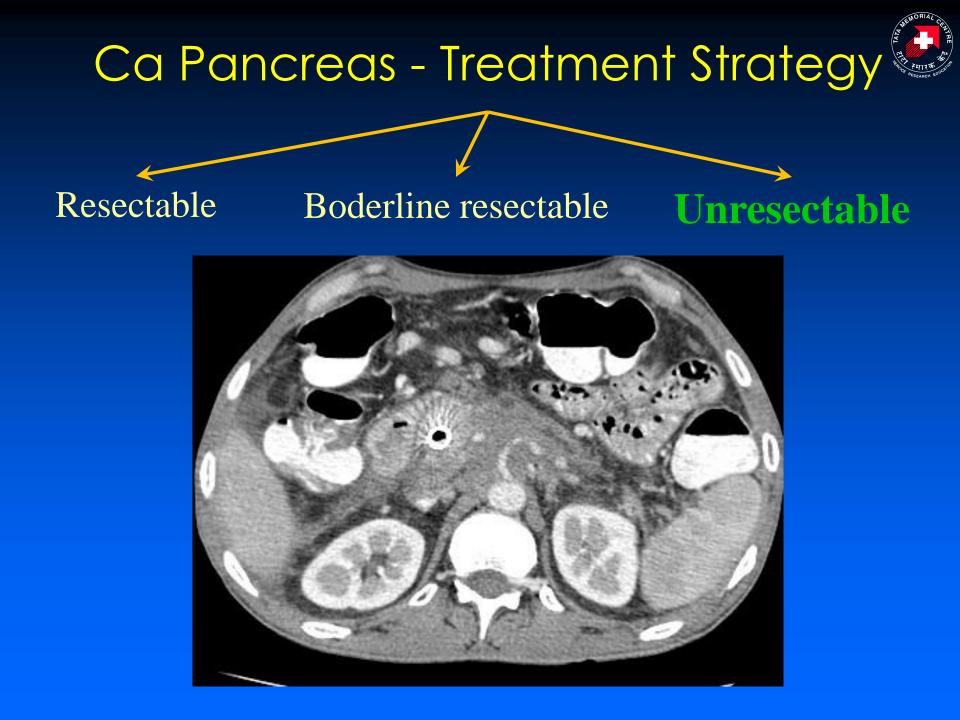
- Head of pancreas/uncinate lesions: Solid tumor contact
 - With the **SMA >180 degrees**
 - With the celiac axis >180 degrees
 - With the first jejunal SMA branch
 - Non reconstructable SMV or portal vein due to tumor involvement or occlusion (can be due to tumor or bland thrombus)
 - With the most proximal draining jejunal branch into the SMV
- Body and tail lesions: Solid tumor contact
 - Of >180 degrees with the SMA or celiac axis
 - With the celiac axis and aortic involvement
 - Unreconstructable SMV or portal vein due to tumor involvement or occlusion (can be due to tumor or bland thrombus)
- For all sites:
 - Distant metastases
 - Metastases to lymph nodes beyond the field of resection





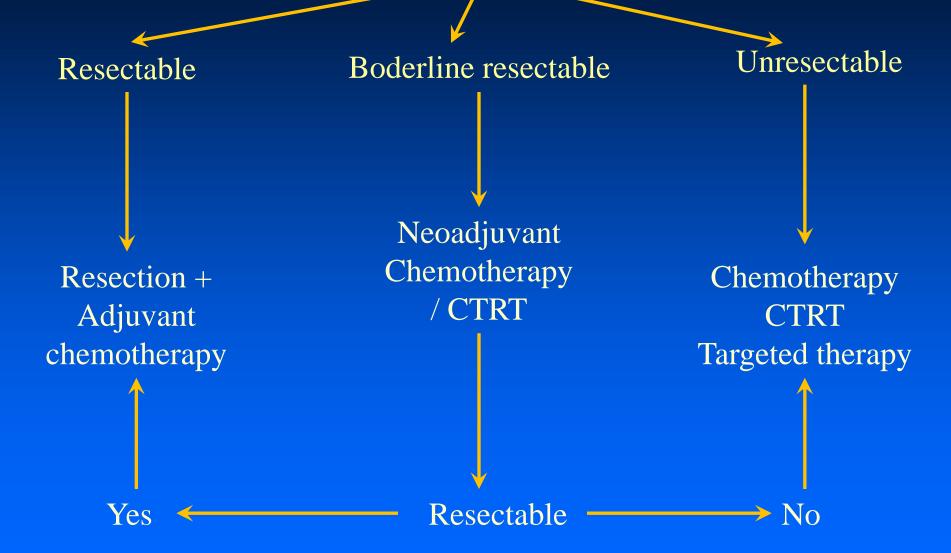






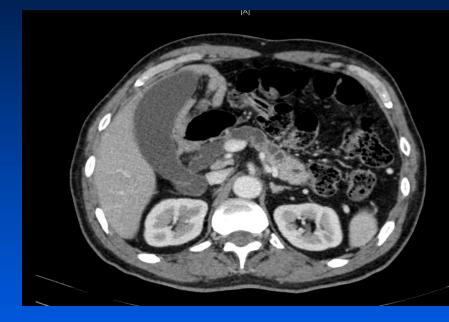


Treatment Strategy



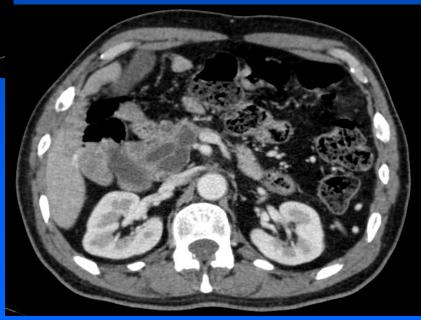


65 year old male; Obstructive jaundice; bilirubin of 13 mg%



RESECTABLE Pre-Op Issues

Normal LFT's BT / CT / INR...WNL's Next step? ? Role of stenting



Treatment Strategy – Preop Issues



Preoperative biliary drainage - Not routinely indicated

The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

Preoperative Biliary Drainage for Cancer of the Head of the Pancreas

Niels A. van der Gaag, M.D., Erik A.J. Rauws, M.D., Ph.D., Casper H.J. van Eijck, M.D., Ph.D., Marco J. Bruno, M.D., Ph.D., Erwin van der Harst, M.D., Ph.D., Frank J.G.M. Kubben, M.D., Ph.D., Josephus J.G.M. Gerritsen, M.D., Ph.D., Jan Willem Greve, M.D., Ph.D., Michael F. Gerhards, M.D., Ph.D., Ignace H.J.T. de Hingh, M.D., Ph.D., Jean H. Klinkenbijl, M.D., Ph.D., Chung Y. Nio, M.D., Steve M.M. de Castro, M.D., Ph.D., Olivier R.C. Busch, M.D., Ph.D., Thomas M. van Gulik, M.D., Ph.D., Patrick M.M. Bossuyt, Ph.D., and Dirk J. Gouma, M.D., Ph.D.*

CONCLUSIONS

Routine preoperative biliary drainage in patients undergoing surgery for cancer of the pancreatic head increases the rate of complications. (Current Controlled Trials number, ISRCTN31939699.)

Treatment Strategy – Preop Issues



Preoperative biliary drainage - Not routinely indicated

Original article

Effect of preoperative biliary stenting on immediate outcome after pancreaticoduodenectomy

P. Jagannath³, V. Dhir³, S. Shrikhande¹, R. C. Shah³, P. Mullerpatan³ and K. M. Mohandas²

Departments of ¹Gastrointestinal Surgery and ²Digestive Diseases and Clinical Nutrition, Tata Memorial Hospital and ³Lilavati Hospital and Research Centre, Mumbai, India

Conclusions

Stent only in symptomatic jaundice Very high hyperbilirubinemia >20mg%

Positive bile culture - Higher morbidity and mortality Uncomplicated stenting – no increase Wait for 3 - 6 weeks post stenting

Br J Surg 2005

Treatment Strategy – Preop Issues 🕻



Need for tissue diagnosis

Obstructive jaundice with a mass lesion in the pancreas on imaging does **NOT** require tissue diagnosis

• Indications:

- if there is evidence of systemic spread of disease,
- if there is local evidence of unresectability on staging studies,
- if the patient is unfit for major surgery,
- if neoadjuvant treatment is being contemplated (eg, for a borderline resectable lesion)
- if alternative diagnoses need to be excluded (eg, metastatic disease to the pancreas).



Need for tissue diagnosis

Obstructive jaundice with a mass lesion in the pancreas on imaging does not require tissue diagnosis

• Indications:

- if Surgery is not the first line of management

if the diagnosis is not clear (eg, metastatic disease to the pancreas).

Treatment Strategy – Preop Issues

- CT, MRI, USG rarely picks up peritoneal metastasis <1 cm in diameter.
- Potentially resectable lesions in the body or tail of the pancreas
 50% will have occult peritoneal metastases.

- Indications:
 - Primary tumour >3cm,
 - Initial CA 19-9 level >100 units/mL
 - Imaging suspicious for peritoneal disease.

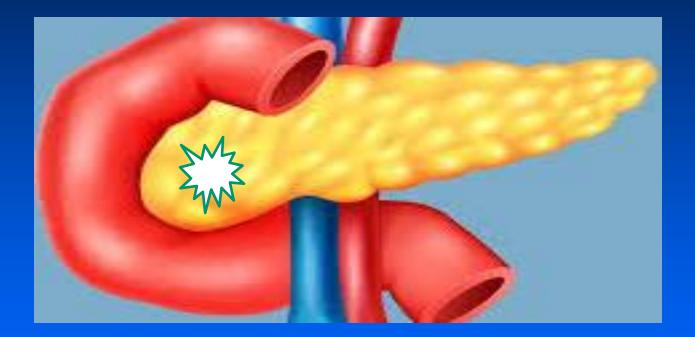
Pancreatic Tumors – Surgical Options



- Depending on location
 - Pancreaticoduodenectomy Classic / PPPD
 - Distal/Subtotal Pancreatectomy \pm splenectomy
 - Total Pancreatectomy

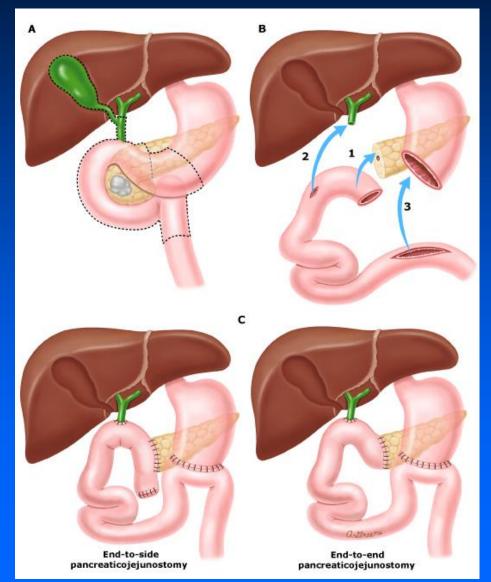


Pancreatico-duodenectomy (PD)



Periampullary tumours Lesions in the head, neck and uncinate process

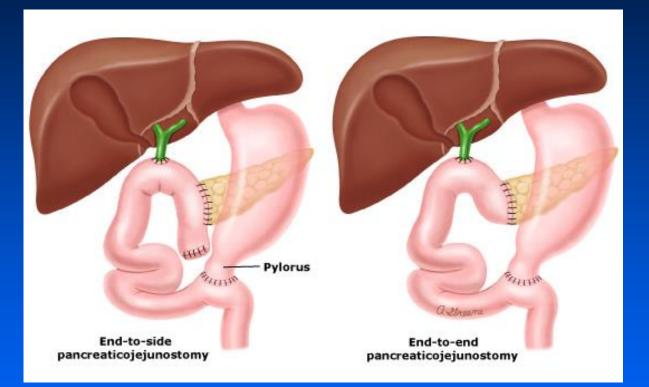
Surgery for Pancreatic Tumors Classical Pancreatico-duodenectomy (PD) (Whipple)



Resection template

- pancreatic head
- duodenum
- first 15 cm of the jejunum
- common bile duct
- gallbladder
- partial gastrectomy

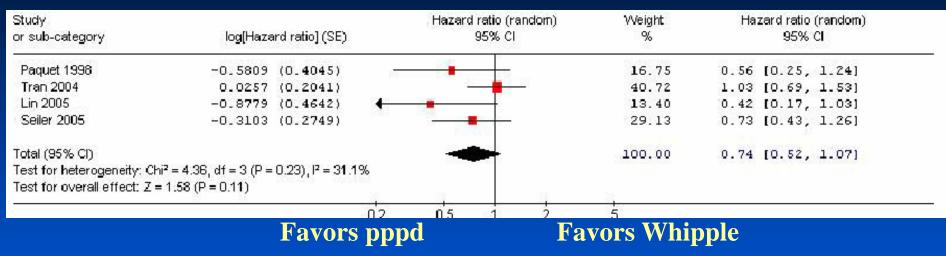
Surgery for Pancreatic Tumors Pylorus Preserving Pancreatico-duodenectomy (PPPD)



 decrease the incidence of postoperative dumping, marginal ulceration, and bile reflux gastritis associated with partial gastrectomy



Survival



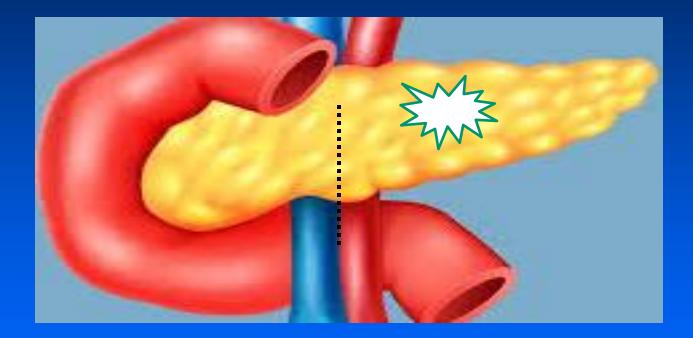
RCT's 6

Equally radical operations No difference in survival No difference in morbidity / mortality Similar QOL Wenger et al., Chirurg 1999 Tran et al., Ann. Surg. 2004 Lin et al., Hepatogastroenterology 2005 Seiler et al., Br. J. Surg. 2005 Paquet et al., Chir. Gastroenterol. 1998 Bloechle et al., DGCh Forumband 1999

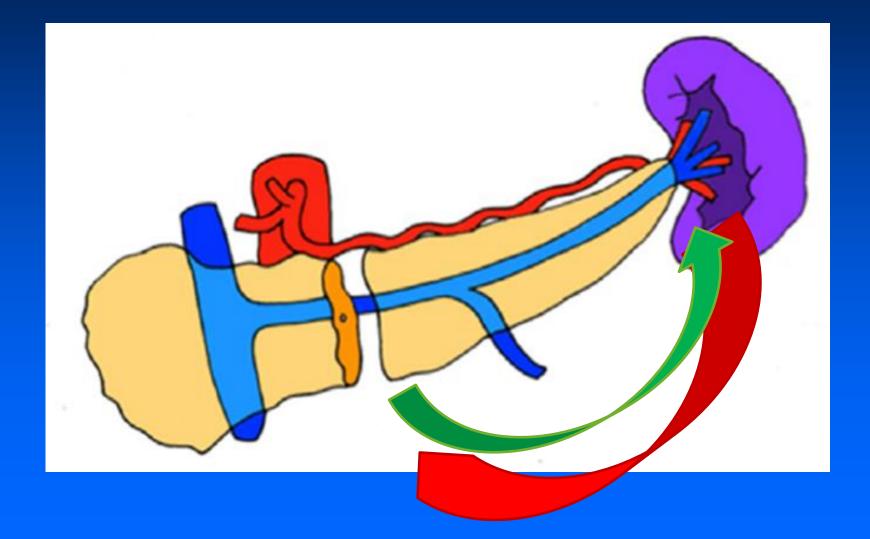
Diener M et al., Ann. Surg. 2007

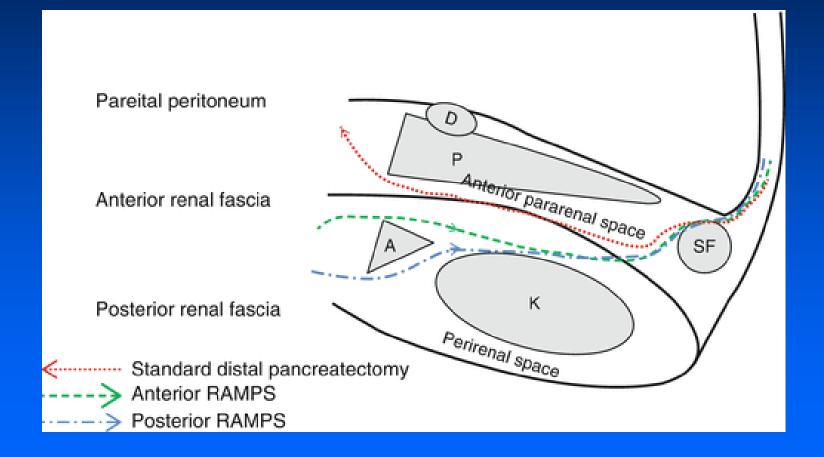


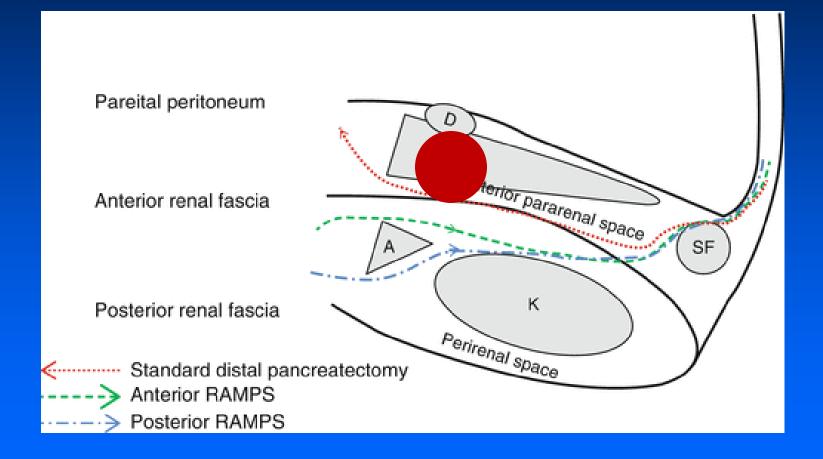
Distal Pancreatectomy +/- Splenectomy

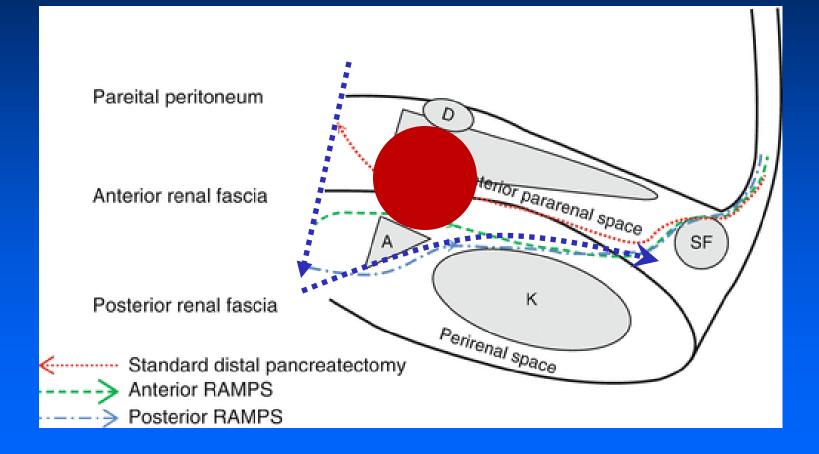


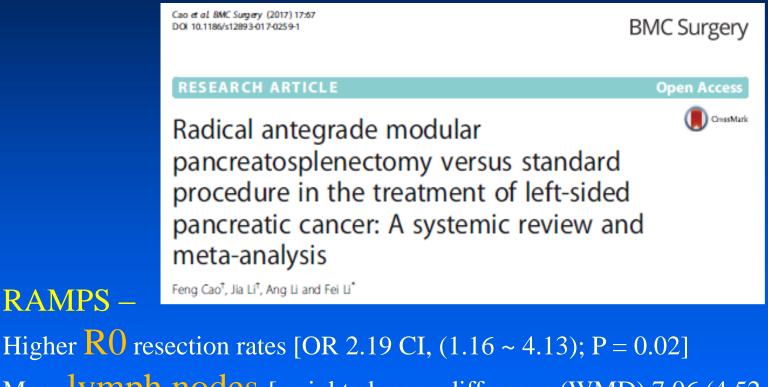












RAMPS

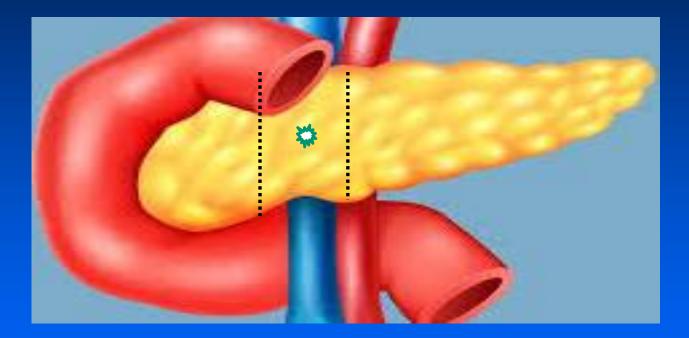
More lymph nodes [weighted mean difference (WMD) 7.06 (4.52 ~ 9.60); P < 0.01]

No statistically significant difference in recurrence rates [P = 0.10], OS [P = 0.05] or DFS [P = 0.93].

BMC Surgery 2017



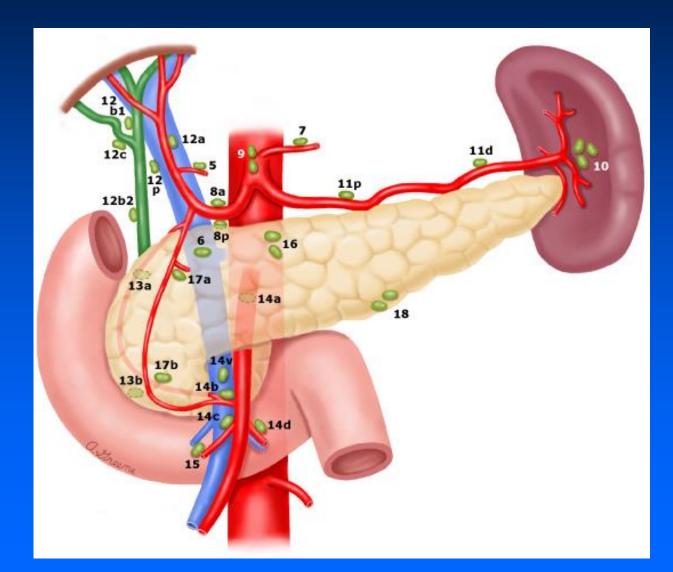
Central/Median Pancreatectomy



Very small lesions in the neck Neuroendocrine tumours

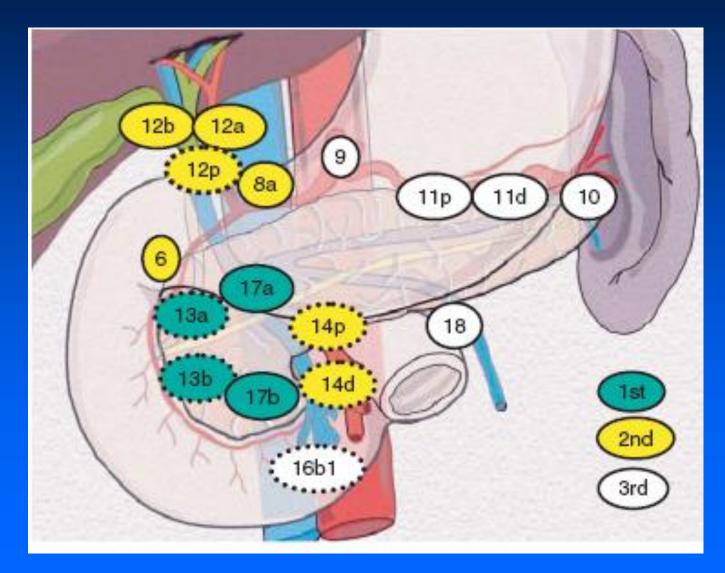


Extent of Lymphadenectomy





Extent of Lymphadenectomy





Extent of Lymphadenectomy

Review: Comparison: Outcome:	LA in PDAC 01 Extended LA vs Standard LA 02 3 YSR	3-yea	3-year survival			
Study or sub-category	Extended LA v n/N	Standard LA n/N	OR (random) 95% Cl	Weight %	OR (random) 95% Cl	
Pedrazzoli 199 Yeo 2002 Farnell 2005 Nimura 2005	98 9/41 65/148 16/39 5/50	9/40 64/146 10/40 9/51		14.71 56.03 17.28 11.98	0.97 [0.34, 2.76] 1.00 [0.63, 1.59] 2.09 [0.80, 5.44] 0.52 [0.16, 1.67]	
Test for hetero	278 5 (Extended LA), 92 (Standard LA) geneity: Chi ² = 3.42, df = 3 (P = 0.33), I ² effect: Z = 0.21 (P = 0.83)	277 = 12.4%		100.00	1.05 [0.69, 1.59]	
		0.1	0.2 0.5 1 2 5			
		Favou	rs Extended LA Favours Star	idard LA		
Review: Comparison: Outcome:	LA in PDAC 01 Extended LA vs Standard LA 04 Overall Complications	m	orbidity			
Study or sub-category	Extended LA / n/N	Standard LA n/N	OR (random) 95% Cl	Weight %	OR (random) 95% Cl	
Pedrazzoli 19 Yeo 2002 Farnell 2005 Nimura 2005	98 8/41 79/148 39/39 34/50	11/40 36/146 25/40 7/51		27.46 31.34 3.37 27.83	0.64 [0.23, 1.81] 3.50 [2.13, 5.74] 48.02 [2.75, 838.36] 13.36 [4.94, 36.11]	
Test for hetero	278 60 (Extended LA), 79 (Standard LA) geneity: Chi ² = 20.64, df = 3 (P = 0.0001 l effect: Z = 2.17 (P = 0.03)	•		100.00	4.52 [1.16, 17.61]	
		0.1 Eavou	0.2 0.5 1 2 5 rs Extended LA Favours Star			
4		Favou	IS LAIGHUGU LA FAVOUIS SIAI	iuaiu LA		

Michalski CW et al., Br J Surg 2007



Reconstruction: Management of pancreatic stump after PD

Pancreatico-enteric anastomotic breakdown still remains a life-threatening complication



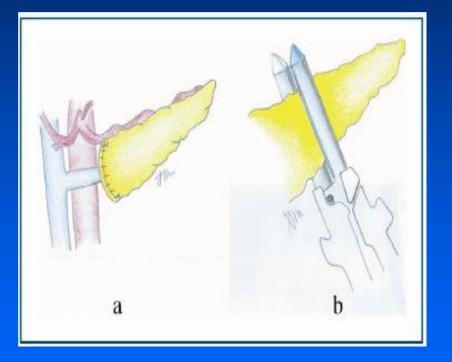


Reconstruction: Management of pancreatic stump after PD

- Closure of the pancreatic stump
- Pancreatico gastrostomy
- Pancreatico duodenostomy



Surgery for Pancreatic Tumors Closure of the pancreatic stump



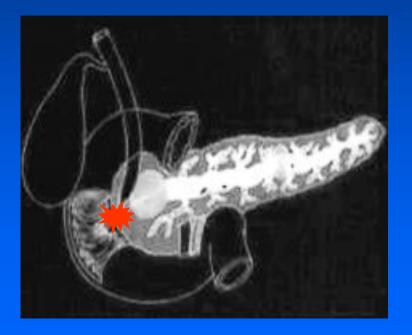
by suture (a) or stapler(b).

- Non-physiological
- High POPF rates
- Exocrine pancreatic insufficiency
- Islet cell dysfunction



Surgery for Pancreatic Tumors Pancreatic Anastomosis: Most demanding

- small duct
- soft texture





Pancreatic Cancer

Distal Bile Duct Cancer



Surgery for Pancreatic Tumors Post Operative Pancreatic Fistula Risk Factors

> **Disease related** (Texture / Location of tumor / Juice output)

> > **Patient related** (MPD location / Age / Obesity etc)

Operative procedure related (Type of anastomosis / High volume centre / Surgeon / blood loss etc)



Surgery for Pancreatic Tumors Pancreatico-jejunostomy

- Most commonly used option
- Various techniques proposed

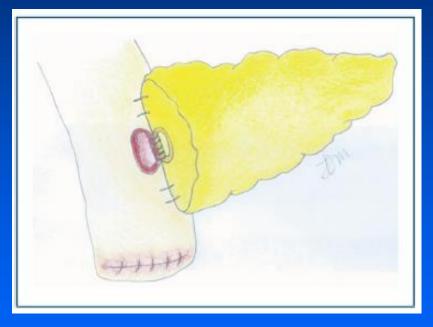
 Trans-mesocolic or antecolic
 Roux-en-Y limb, an omega jejunal loop
- There are three main types of PJ:
 - Duct-to-mucosa anastomosis
 - Invagination anastomosis
 - Binding pancreatico-jejunostomy.

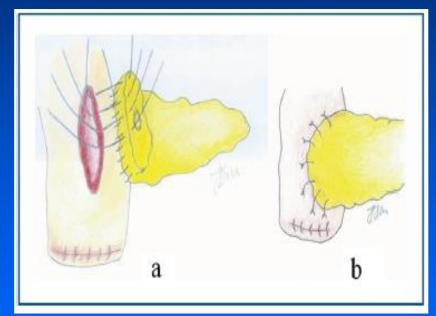


Surgery for Pancreatic Tumors Pancreatico-jejunostomy

Duct to Mucosa-PJ

Invagination -PJ

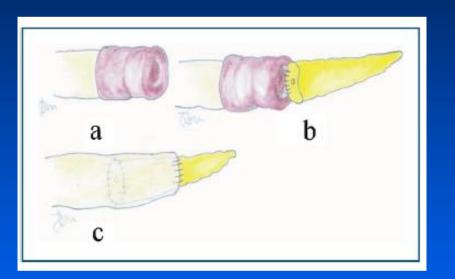




At least 6 RCT's comparing both techniques, 4 showed no difference, 2 in favour of duct to mucosa.



Surgery for Pancreatic Tumors Binding Pancreatico-jejunostomy



a: everted jejunal mucosa;b: suture between the jejunal mucosa and pancreatic stump;c: completed binding pancreatico-jejunostomy.

Proposed by Peng et al

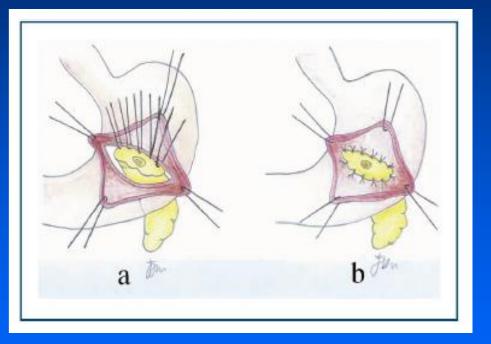
RCT - Conventional Versus Binding Pancreaticojejunostomy, *Ann Surg 2007*

Conclusion – Binding PJ was associated with significantly decreased postoperative complication

However, these results were not re-produced.



Surgery for Pancreatic Tumors Pancreatico-gastrostomy



Advantages – -Thick gastric wall, -profusely vascularized, -close to the pancreas, -anastomosis is performed in a field where no enterokinase is present

Disadvantages--High incidence of postoperative anastomotic bleeding. -Pancreatic duct obstruction with gland atrophy and exocrine insufficiency



The American Journal of Surgery 193 (2007) 171–183 Clinical surgery–International

Pancreaticojejunostomy versus pancreaticogastrostomy: systematic review and meta-analysis

Moritz N. Wente, M.D., M.Sc.^a, Shailesh V. Shrikhande, M.D.^{a,b}, Michael W. Müller, M.D.^a, Markus K. Diener, M.D.^a, Christoph M. Seiler, M.D., M.Sc.^a, Helmut Friess, M.D.^a, Markus W. Büchler, M.D.^{a,*}

> *Department of General, Visceral and Trauma Surgery, University of Heidelberg, Heidelberg, Germany *Department of Gastrointestinal Surgical Oncology, Tata Memorial Hospital, Mumbai, India

> > Manuscript received August 10, 2006; revised manuscript October 11, 2006

- 13 nonrandomized observational clinical studies, 3 RCTs
- Observational studies reported superiority of PG over PJ, most likely influenced by publication bias
- All three RCTs suggested both PJ and PG provide equally good results



Pancreatico-gastrostomy Vs Pancreatico-jejunostomy

- A meta-analysis of seven randomized trials
 - PG resulted in significantly lower POPF(11% Vs 19%)
 - Biliary fistulas (2% Vs 5%)

Ann Surg. 2015;261(5):882.

- Cochrane review of 10 trials:- No difference in
 - POPF rates (21.4% PG Vs 24.3% PJ),
 - Clinically significant POPF (12.8 % PG Vs 19.3% PJ)
 - Postoperative mortality (4.8% PG Vs 3.9% PJ)

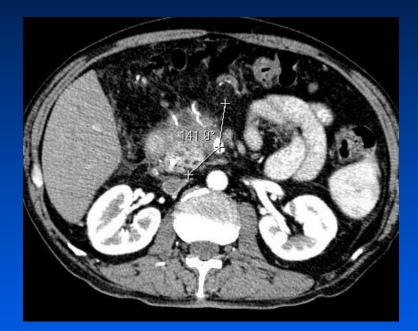
Cochrane Database Syst Rev. 2017;9:CD012257

Conclusion

Pancreatico-gastrostomy is equivalent to Pancreatico-jejunostomy



Boderline resectable pancreatic cancer

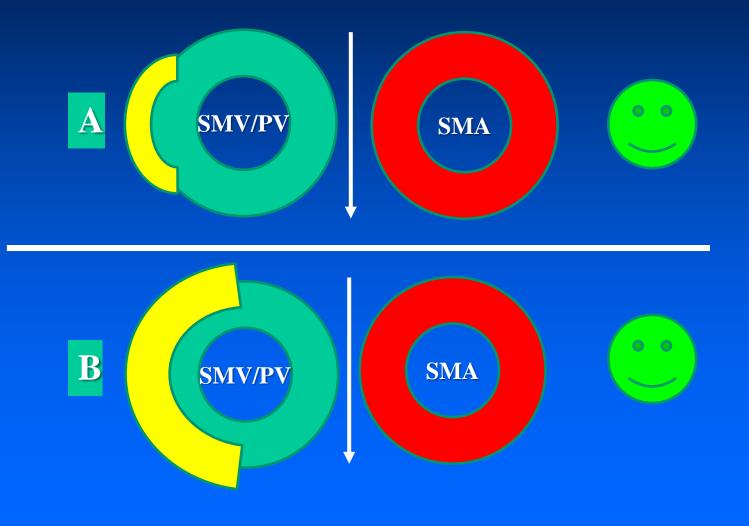


Surgical Options ? Upfront resection ? Neo-adjuvant chemotherapy ? Neo-adjuvant chemoradiotherapy



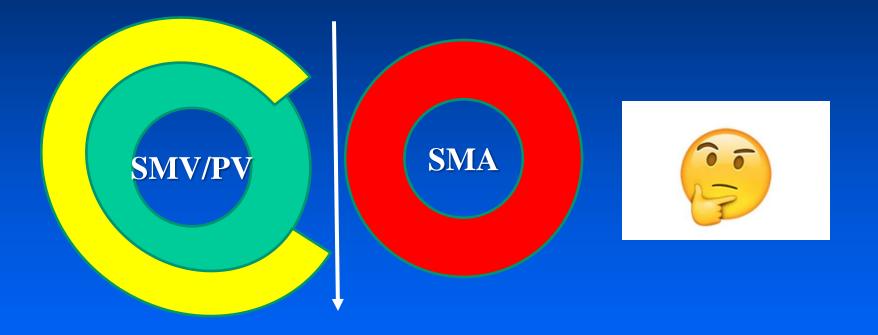


Boderline resectable pancreatic cancer





Boderline resectable pancreatic cancer





Neo-adjuvant therapy for patients with BRPC:

A systematic review and meta-analysis of response and

resection percentages

Tang K, Lu W et al. Pancreatology 2016;16: 28-37

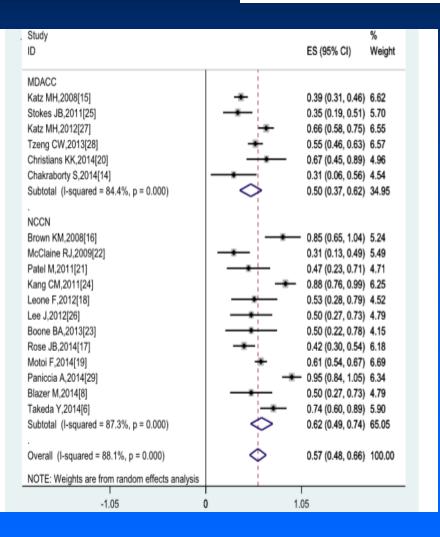
18 trials; N=959

13 trials chemo + RT

5 chemo alone



R0 resection estimates



63% pts resected87% R0Median OS 25.9 months

	FOLFIRINOX (n=64)	Gem-based
Resection rate	72%	67%
RO	60%	58%
G3 /4 Toxicity	53%	30%

Tang K, Lu W et al. Pancreatology 2016;16: 28-37



Does CT RT have higher response rate than chemo alone ?

- Very little evidence for this
- Even in the combined analysis , the definition of response varied over years
- Primary pancreatic cancer
 - \rightarrow appears less responsive than metastatic diseases
 - \rightarrow difficult to measure even in high quality scan



Surgery for Pancreatic Tumors Boderline resectable pancreatic cancer

For borderline resectable diseases, NACT or neo-adjuvant CTRT is recommended.

Selected cases when R0 resection is possible can undergo upfront resection.



Surgery for Pancreatic Tumors Boderline resectable pancreatic cancer Approaching the BR tumor.....

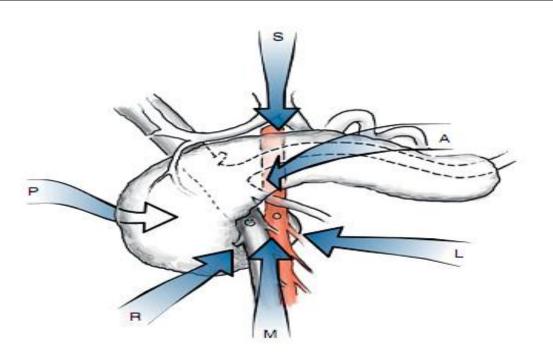
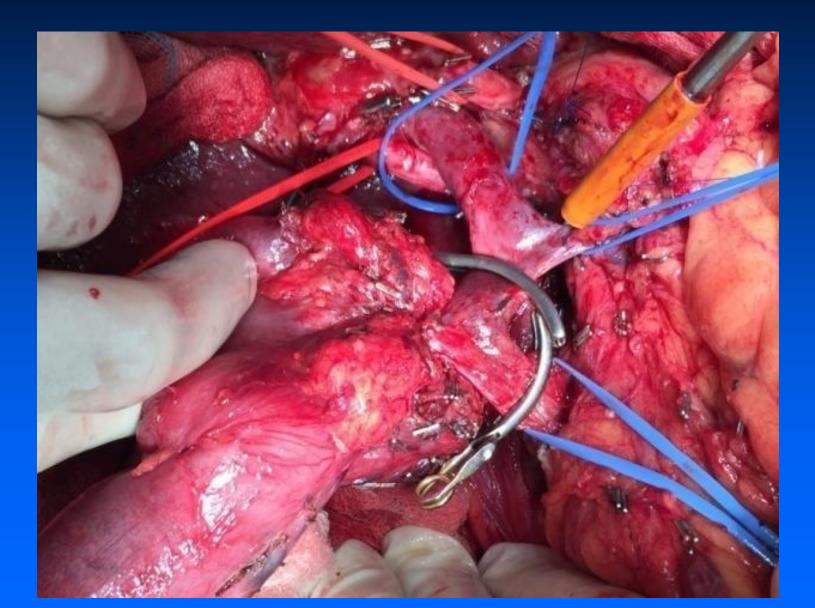
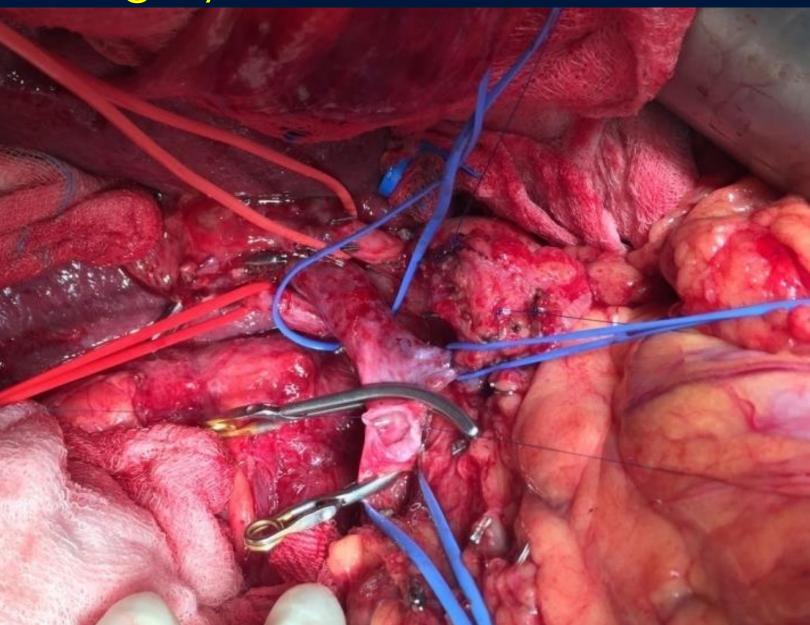


Fig. 1 Diagram showing the six approaches to the superior mesenteric artery: S, superior approach; A, anterior approach; P, posterior approach; L, left posterior approach; R, right/medial uncinate approach; M, mesenteric approach

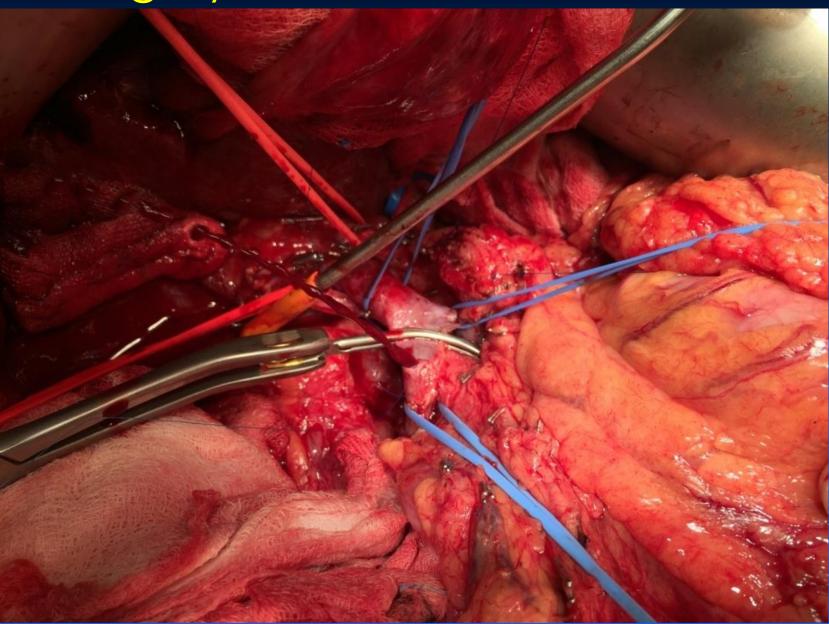




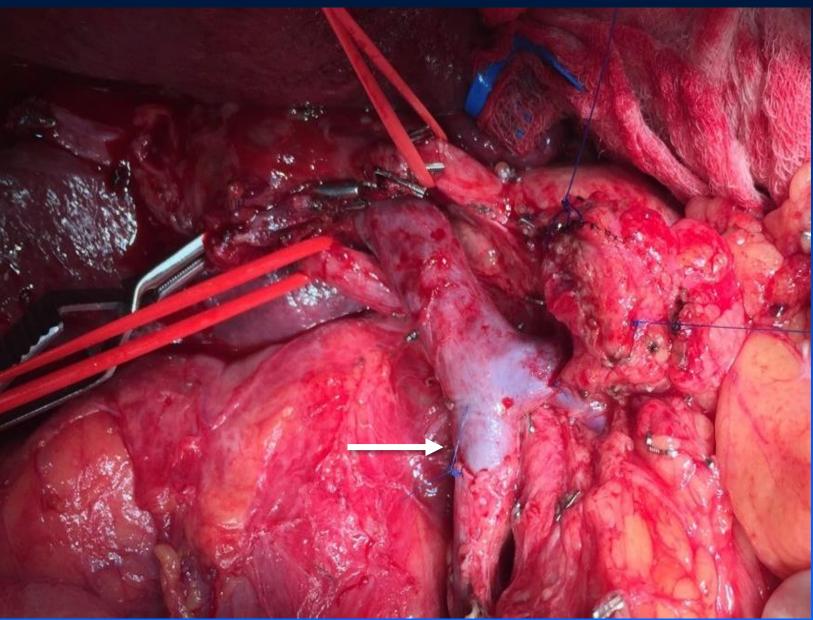














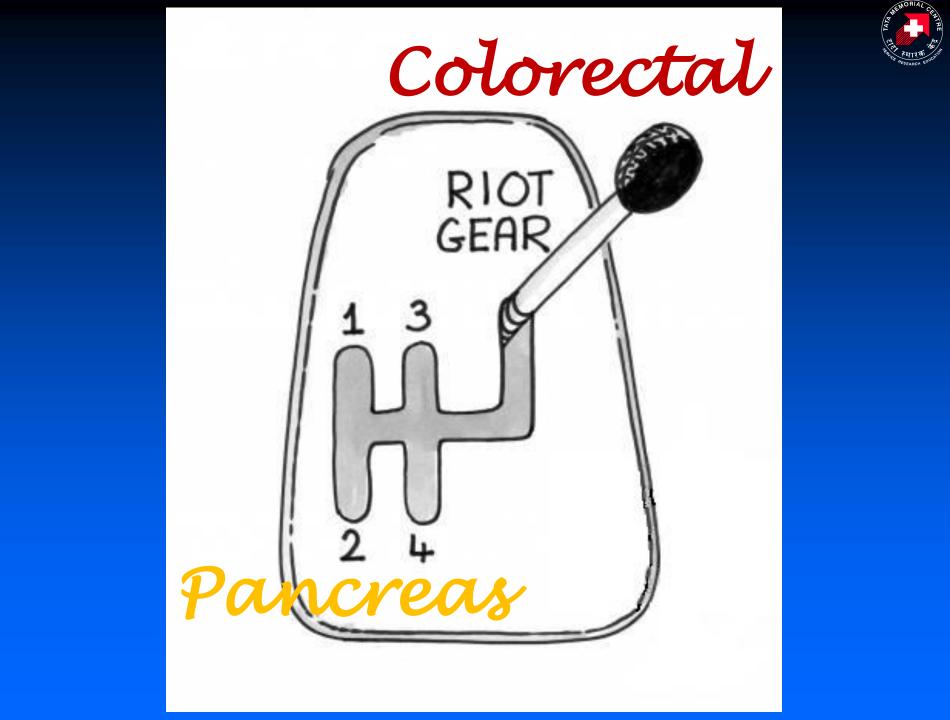
Period	A (1992-2001)	B (Jan 2003- July 2009)	C* (Aug 2009- Dec 2011)	D (Jan 2012- Sept 2016)	E (Oct 2016 - Dec 2017)	Total
Ν	144	206	150	516	196	1212
Resections/yr	16	34	60	110	160	
Median Age	50	53 (18-74)	53 (8-82)	55(10-85)	53	53 (8 - 85)
POPF	16% (23)	8% (16)	10.7% (16)	16.08% (83)	13.2% (26)	13.3% (162)
Bile leaks	6.3% (9)	3.4% (7)	0.7% (1)	0.7%(4)	2.5% (5)	2% (25)
DGE	6.9% (10)	2.4% (3)	2% (3)	6.25%(32)	6.6% (13)	4.8% (59)
РРН	11.1% (16)	5% (10)	2% (3)	3.68%(19)	2.5% (5)	4.3% (53)
Median hospital stay	16	15	12	12	12	12
Morbidity	41.7% (60)	30% (61)	29% (44)	26.74% (138)	25% (49)	29.1% (353)
Mortality	6.3% (9)	4.8% (10)	5.3% (8)	2.71% (14)	1.5% (3)	3.6% (44)

*Pancreatology 2013



Pancreas – Surgical Options Summary

- Very poor prognosis, 5 yrs survival 6%
- Surgery only curative option
- Classify patients resectable/borderline/unresectable
- Selective preop biliary drainage
- Staging lap (occult metastasis) selected cases
- Extended lymphadenectomy No role
- Type of resection location of tumour
- RAMPS for body and tail lesions
- PPPD procedure of choice of head and periamp
- Pancreatico-Jejunostomy = Pancreatico-Gastrostomy
- BRPC NACT/RT → Surgery.





GI Malignancy – Surgical Options

- Colorectal
 - Colon:
 - Standard colectomy
 - Complete mesocolic excision
 - Multivisceral resection
 - Rectum:
 - Total Mesorectal excision (TME)
 - Sphincter Preservation
 - Abdomino perineal resection (APR)
 - Extralevator APR
 - Rectal resection Beyond TME
 - Colorectal peritoneal metastasis CRS+HIPEC





Colon Cancer

Rectal Cancer

• Stage I – T1/T2, N0, M0

Upfront SURGERY, no adjuvant treatment, surveillance



• Stage IV – Any T, Any N, M1

Chemotherapy (backbone), SURGERY in selected cases

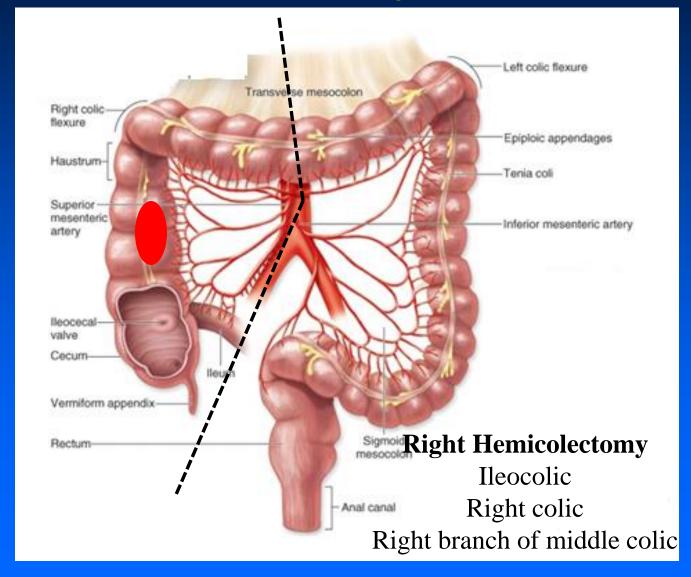


Colon Cancer - Localised disease (Stage I, II, III)



Radical colectomy IS NOT Resection anastomosis of the colon



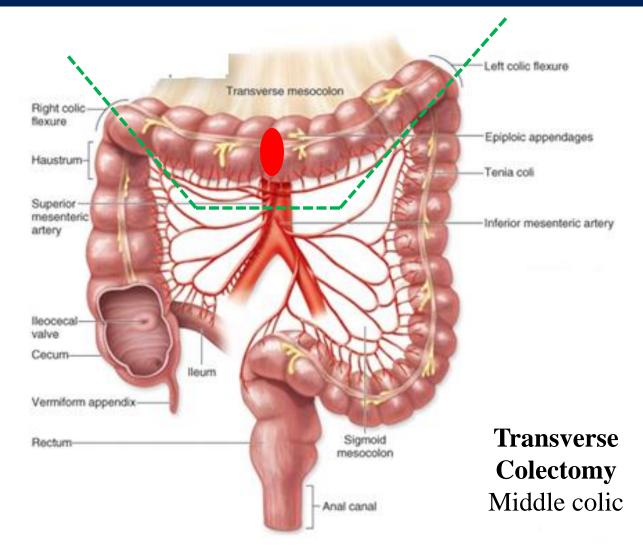




Left colic flexure Transverse mesocolon Right colic flexure Epiploic appendages Haustrum Tenia coli Superior mesenteric Inferior mesenteric artery artery lleocecal valve Cecum leun Vermiform appendix **Right Extended** Hemicolectomy Sigmoid Rectum mesocolon Ileocolic Right colic Anal canal Middle colic

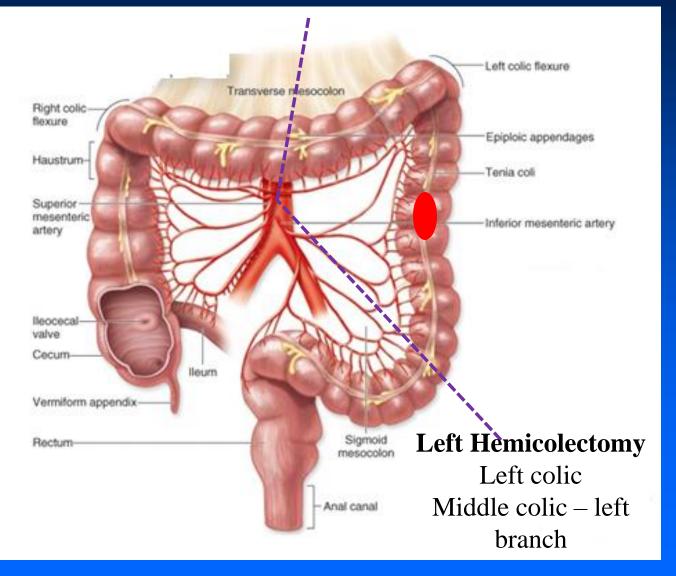


Colon Cancer - Localised disease (Stage I, II, III) – SURGERY



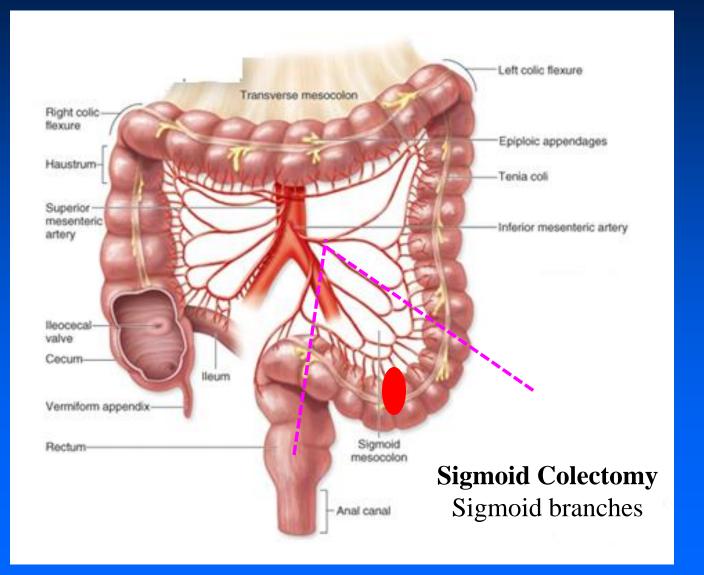


Colon Cancer - Localised disease (Stage I, II, III) – SURGERY

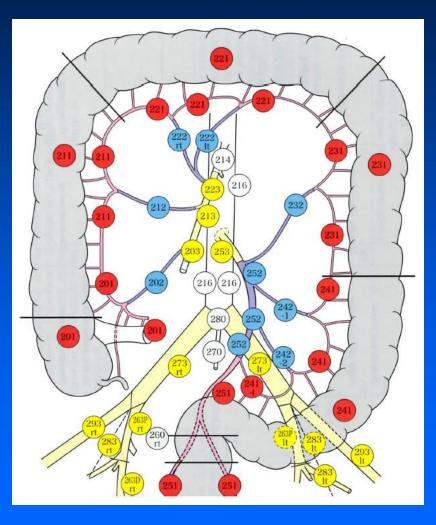




Colon Cancer - Localised disease (Stage I, II, III) – **SURGERY**



Colon Cancer – Surgical Options Colectomy – Lymph node stations



Lymph node classification according to the Japanese Society for Cancer of the Colon and Rectum (JSCCR).

Level 1 lymph node stationsLevel 2 lymph node stations

Level 3 lymph node





Colectomy – Lymph node stations

	Cecum	Ascending colon	Hepatic flexure	Proximal transverse colon
N1	51%, 55%	52%, 57%	46%, NA	46%, 43%
N2	33%, 11%	48%, 27%	56%, NA	59%, 21%
N3	11%, 10%	7%, 16%	17%, NA	15%, 36%

Park IJ, Choi GS, Kang BM, Lim KH, Jun SH (2009) Lymph node metastasis patterns in right-sided colon cancers: is segmental resection of these tumors oncologically safe? Ann Surg Oncol 16:1501–1506 Kobayashi H, Enomoto M, Higuchi T, Uetake H, Iida S, Ishikawa T et al (2011) Clinical significance of lymph node ratio and location of nodal involvement in patients with right colon cancer. Dig Surg 28: 190–197



Colon Cancer – Surgical Options Skip Metastasis – Right Colon Cancer

Table 2 Nodal Status of this case series $(n = 244)$				
Total number of dissected lymph nodes (mean \pm SD)	34.4 ± 8.4			
Number of harvested lymph node (mean, range)				
N1	15.4 (4-28)			
N2	12.6 (6-35)			
N3	6.4 (4-16)			
Level of lymph-node involvement (Number of patients)				
NO	42			
Orderly metastasis	80.2 % (162/202)			
N1	80			
N1 + N2	55			
N1 + N2 + N3	27			
Skip metastasis	19.8 % (40/202)			
N2 only	19			
N1 + N3	4			
N2 + N3	6			
N3 only	11			

D3 dissection - stage migration (stage II to III) in 4.5 % classified as N0 lesions after conventional D2 dissection.

Liang JT, Lai HS, Huang J, Sun CT (2014) Longterm oncologic results of laparoscopic D3 lymphadenectomy with complete mesocolic excision for right-sided colon cancer with clinically positive lymph nodes. Surg Endosc 29:2394–2401



Colon Cancer – Surgical Options Complete Meoscolic Excision

Principle of CME:

Removal of all lymphatic, vascular and neural tissue in the drainage area of the tumour in a complete mesocolic envelope with intact mesentery, peritoneum and encasing fascia

Three main components to CME:

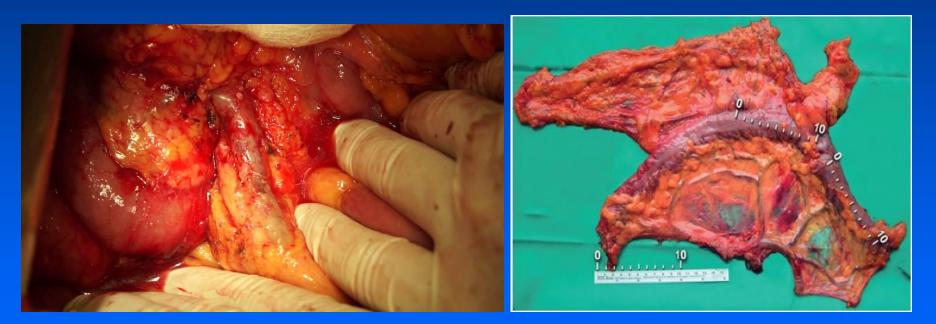
a. Dissection in the embryological plane - lymphaticsb. Central vascular tie – Lymph nodes at the rootc. Resection of a sufficient length of bowel (10cm on each side)

Hohenberger W, Weber K, Matzel K, Papadopoulos T, Merkel S (2009) Standardized surgery for colonic cancer: complete mesocolic excision and central ligation—technical notes and outcome. Color Dis : Off J Assoc Coloproctology Great Britain Ireland 11(4):354–364 .



Colon Cancer - Localised disease (Stage I, II, III) – **SURGERY**

Complete Mesocolic Excision (CME) Resection within fascial envelop Central vascular ligation



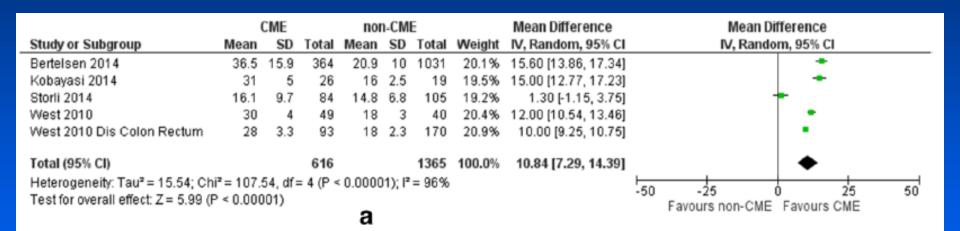
10% improvement in disease free survival

Lancet Oncol 2015; 16: 161–68



CME Vs Standard Colectomy

Lymph node yield



Gouvas N et al. Surgery along the embryological planes for colon cancer: a systematic review of complete mesocolic excision. Int J Colorectal Dis (2016) 31:1577–1594



Colon Cancer – Surgical Options CME Vs Standard Colectomy Lymph node yield – Prognostic Significance

Higher lymph node yield – better survival

Le Voyer et al (2003) Colon cancer survival is associated with increasing number of lymph nodes analyzed: a secondary survey of intergroup trial INT-0089. J Clin Oncol 21(15):2912–2919

Chen SL, Bilchik AJ (2006) More extensive nodal dissection improves survival for stages I to III of colon cancer: a population based study. Ann Surg 244(4):602–610

Chang GJ et al(2007) Lymph node evaluation and survival after curative resection of colon cancer: systematic review. J Natl Cancer Inst 99(6):433–441



CME Vs Standard Colectomy Lymph node yield – Prognostic Significance

Lymph node ratio - better prognostic indicator than the number of involved lymph nodes or pN status Greater the negative nodes : metastatic nodes - better prognosis

Parnaby CN et al (2015) Prognostic value of lymph node ratio and extramural vascular invasion on survival for patients undergoing curative colon cancer resection. Br J Cancer 2015 Jul 14;113(2):212-9

Lykke Jet al (2013) The relation between lymph node status and survival in Stage I-III colon cancer: results from a prospective nationwide cohort study. Colorectal Dis 15(5):559–565

Rosenberg R et al(2008) Prognosis of patients with colorectal cancer is associated with lymph node ratio: a single-center analysis of 3,026 patients over a 25-year time period. Ann Surg 248(6):968–978



Colon Cancer – Surgical Options CME Vs Standard Colectomy CME – Lymph node yield

Survival benefit with more extensive lymphadenectomy / higher No. of -ve nodes,

- Stage migration
- Removal of nodes with micrometastases, if left in situ, significantly affect survival

Færden AE et al. (2011) Lymph node micrometastases and isolated tumor cells influence survival in stage I and II colon cancer. Dis Colon Rectum 54(2):200–206



Colon Cancer – Surgical Options Standard Vs CME Colectomy

	CME (n= 529)	Standard (n=1071)	р
Morbidity (60 day)	30.6%	28.5%	0.351
Injury to other organs (Spleen, SMV, colon)	9.1%	3.6%	<0.01
Surgical complications	20.8%	19.3%	0.491
Anastomotic leak	8.5%	7.1%	0.327
Non Surgical complications	18.9%	16.2%	0.163
Mortality (90 day)	6.2%	4.9%	0.219

Bertelsen CA et al. Short-term outcomes after complete mesocolic excision compared with 'conventional' colonic cancer surgery. Br J Surg 2016 Apr;103(5)



CME Vs Standard Colectomy Recurrence Rates

Study	Recurrence	Standard	CME	р
Bertelsen et al (2015)	Local + distant	16.8%	11.3%	0.028
Galizia et al (2014)	Local	20.7%	0%	0.034
Storli et al (2013)	Local	2.9%	1.2%	0.19
	Distant	8.6%	2.4%	0.19

Gouvas N et al. Surgery along the embryological planes for colon cancer: a systematic review of complete mesocolic excision. Int J Colorectal Dis (2016) 31:1577–1594



CME Vs Standard Colectomy Survival Rates

Study	Survival	Standard	CME	р
Galizia et al (2014) Right colon	OS	74.1%	91%	0.055
Storli et al (2013) Stage I/II	OS	79%	88.1%	0.003
Bertelsen et al (2015)	DFS	75.9%	85.7%	0.001
Storli et al (2013) Stage I/II	DFS	74.3%	82.1%	0.026

Gouvas N et al. Surgery along the embryological planes for colon cancer: a systematic review of complete mesocolic excision. Int J Colorectal Dis (2016) 31:1577–1594



Colon Cancer – Surgical Options Standard / Conventional Colectomy

Open compared with laparoscopic complete mesocolic excision with central lymphadenectomy for colon cancer: a systematic review and meta-analysis

C. D. Athanasiou*, G. A. Markides*, A. Kotb*, X. Jia*, S. Gonsalves* and D. Miskovic*†

*John Goligher Colorectal Unit, St James' University Hospital, The Leeds Teaching Hospitals, Leeds, UK and †The Leeds Institute of Biomedical and Clinical Sciences, University of Leeds, Leeds, UK

Colorectal Dis. 2016 Jul;18(7):O224-35.

Conclusion: Based on the current evidence, the laparoscopic technique appears to be **at least as safe** as the open technique when used in performing ELTs for colonic cancer, with similar morbidity and oncological outcomes.



Colon Cancer – Surgical Options Standard / Conventional Colectomy

Int J Colorectal Dis (2014) 29:419–428 DOI 10.1007/s00384-013-1818-2

REVIEW

The rationale behind complete mesocolic excision (CME) and a central vascular ligation for colon cancer in open and laparoscopic surgery

Proceedings of a consensus conference

K. Søndenaa • P. Quirke • W. Hohenberger • K. Sugihara • H. Kobayashi • H. Kessler • G. Brown • V. Tudyka • A. D'Hoore • R. H. Kennedy • N. P. West • S. H. Kim • R. Heald • K. E. Storli • A. Nesbakken • B. Moran Norway, UK, Germany, Japan, USA, Belgium, Korea

Conclusion: The consensus conference agreed that there are sound oncological hypotheses for a more radical approach than has been common up to now. However, this may not necessarily apply in early stages of the tumour stage. Laparoscopic resection appears to be equally well suited for resection as open surgery.



Colon Cancer – Surgical Options CME – TMH Experience

244 patients (CME n=88; NCME n=156) met the inclusion criteria

Parameter	CME (n=88)	Non-CME (n=156)	р
Age (mean, yrs)	52.08	50.59	0.38 ^a
Sex male	55 (62.5)	106 (67.9)	0.40 ^b
ASA 1	46 (52.3)	85 (54.5)	0.75 ^b
ASA 2	39 (44.3)	68 (43.6)	
ASA 3	3 (3.4)	3 (1.9)	
Site			
Caecum	27 (30.7)	50 (32.0)	0.83 ^b
Ascending colon	34 (38.6)	59 (37.8)	
Transverse colon	7 (8.0)	17 (11.0)	
Hepatic Flexure	20 (22.7)	30 (19.2)	
Lap	31 (35.2)	14 (8.9)	<0.001 ^b

a – Student's t test b – Chi-Square test

Numbers in parenthesis indicate percentage



Colon Cancer – Surgical Options CME – TMH Experience

Parameter	CME (n=88)	Non-CME (n=156)	р
Sx Type Rt Hemicolectomy	63 (71.6)	124 (79.5)	0.21 ^b
Rt Extended Hemicolectomy	25 (28.4)	32 (20.5)	
BMI (mean, Kg/m ²)	22.97	22.53	0.46 ^a
Blood Loss (mean, ml)	218.6	295.0	0.005 ^a
Anastomotic leak	7 (7.9)	11 (7.1)	0.80 ^b
Clavien-dindo 0-IIIa IIIb – V	81 (92) 7 (8)	141 (90.4) 15 (9.6)	0.82 ^b
Hospital stay (mean, days)	7.41	7.56	0.82 ^a

a - Student's t test = b - Chi-Square test

Numbers in parenthesis indicate percentage



Colon Cancer – Surgical Options CME – TMH Experience

Adjuvant chemotherapy – CME (58%) NCME (52.6%) [p=0.79] Median follow up duration - 20.8 months

Parameter	CME (n=88)	Non-CME (n=156)	р
pT Stage			
T2	10 (11.4)	24 (15.4)	0.12 ^b
T3	58 (65.9)	112 (71.8)	
T4a	20 (22.7)	20 (12.8)	
pN Stage			
N0	51	102	0.45 ^b
N1	22	35	
N2	15	19	
Total Nodes (mean)	32.73	27.35	0.003 ^a
90-day Mortality	1.13%	1.28%	0.921 ^b
3-yr OS	93.6%	95.7%	0.56 ^c
3-yr DFS	85.3%	80.2%	0.15 °

a:Student's t test; b:Chi-Square test; c: Kaplan Meier. Numbers in parenthesis indicate percentage



29yr male, ECOG 0

Colonoscopy – polypoid lesion at rectosigmoid + hepatic flexure mass

Exploratory laparotomy: Bulky mass adherent to pancreatic head, ileotransverse anastomosis done.

Patient was given 6# FOLFIRINOX + 6# Cisplatin & 5FU



Lancet Oncol. 2012 Nov; 13(11): 1152–1160. doi: <u>10.1016/S1470-2045(12)70348-0</u> PMCID: PMC3488188

Feasibility of preoperative chemotherapy for locally advanced, operable colon cancer: the pilot phase of a randomised controlled trial

FOxTROT Collaborative Group^{†*}

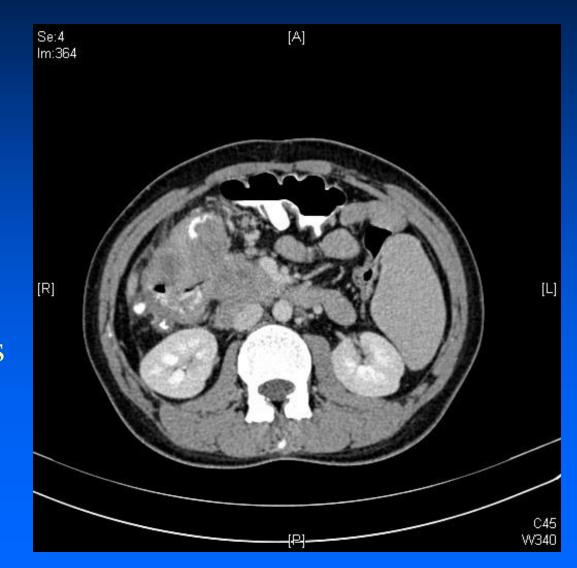
Results: Feasible with acceptable toxicity and perioperative morbidity

• FOxTROT phase 3 results awaited.

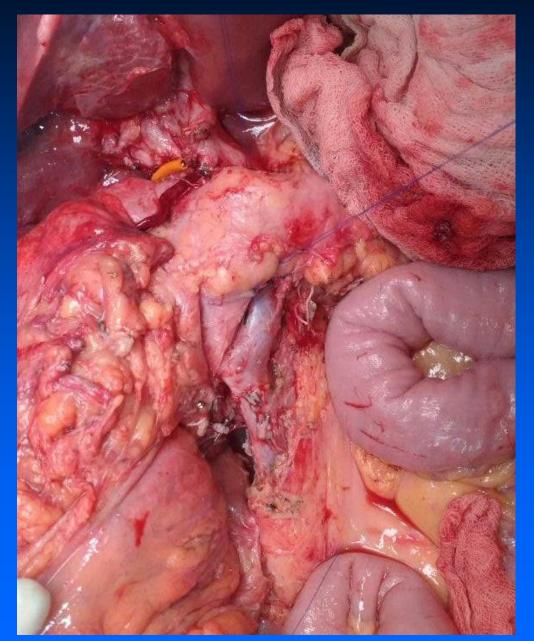


Post 6# FOLFIRINOX + 6# Cisplatin & 5FU

CECT (T+A+P) – non meastatic - Bulky hepatic flexure mass with infiltration into pancreatic head + loss of plane with SMV







Total colectomy + en masse PPPD (SMV sleeve resection) + ilesorectal anastomosis



HPR: MDAC ascending colon infiltrating into pancreas and duodenum (yT4). LVE+ PNI + rectum : 2 polyps ---tubulovillous adenoma with low grade dysplasia Nodes : peripancreatic + hepatic 0/25 middle colic 0/2 colonic nodes :0/53. Total nodes: 0/80

Stage: ypT4N0

Patient is alive without disease at 1 year





Annals of Surgical Oncology

September 2013, Volume 20, <u>Issue 9</u>, pp 2929–2936

Multivisceral Resection in Colorectal Cancer: A Systematic Review

- 22 studies comprising 1575 patients
- Most common organs resected bladder and reproductive organs
- Perioperative mortality was 4.2 % with morbidity of 41.5 %
- Overall 5-year survival rate was 50.3 %
- R0 resection was the strongest factor associated with long-term survival.



Hepatobiliary Pancreat Dis Int. 2015 Jun;14(3):320-4. Combined right hemicolectomy and pancreaticoduodenectomy for locally advanced right hemicolon cancer. Sheng QS¹, Chen WB, Li MJ, Cheng XB, Wang WB, Lin JJ.



Long term survival after right hemicolectomy and pancreatoduodenectomy for locally advanced colonic cancer: Case report

Iraklis Perysinakis*, Alexander Nixon, Aggeliki Katopodi, Emmanouil Tzirakis, Despoina Georgiadou, Spyridon Avlonitis, Ilias Margaris

3rd Surgical Department, "George Gennimatas" General Hospital of Athens, Mesogeion Av. 154, 15669, Greece



Langenbeck's Archives of Surgery

January 2014, Volume 399, <u>Issue 1</u>, pp 33–40

Clinical review: surgical management of locally advanced and recurrent colorectal cancer

1,470 patients with recurrent or locally advanced primary colorectal cancer - 22 studies.
R0 resection offers best prognosis with a 5-year survival of up to 70 %
MVR needed in approx. 10 % with the most commonly involved organ being bladder
Mean post-operative morbidity is 40 %



GI Malignancy – Surgical Options Colon - Summary

- Surgery offer the only possibility for long term control
- Upfront surgery is usually the initial treatment option
- Radical colectomy is not resection-anastomosis of colon
- Complete mesocolic excision in suspected node positive
- T4b lesions Multivisceral resection with R0
- Neoadjuvant chemotherapy is feasible more data required.



GI Malignancy – Surgical Options

- Colrectal
 - Rectum:
 - Total Mesorectal excision (TME)
 - Sphincter Preservation
 - Abdomino perineal resection (APR)
 - Extralevator APR
 - Rectal resection Beyond TME

- Colorectal Peritoneal Metastasis - CRS+HIPEC



Rectal Cancer – Surgical options

MRI – Rectal Cancer

Diagnostic accuracy of preoperative magnetic resonance imaging in predicting curative resection of rectal cancer: prospective observational study

MERCURY Study Group

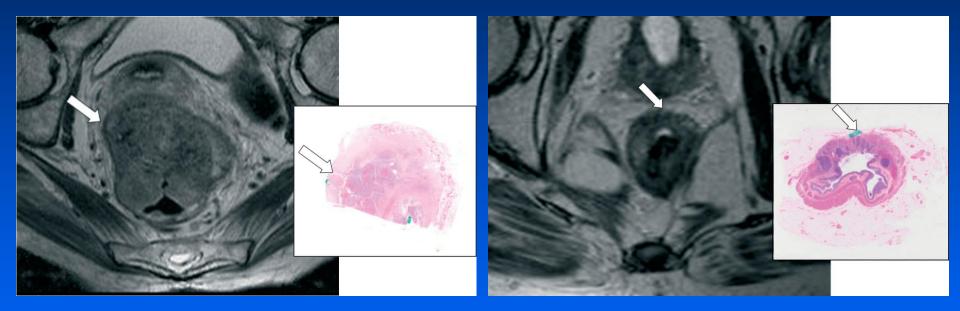
Accuracy for predicting involved CRM – 92%

BMJ. 2006 Oct 14;333(7572):779



Rectal Cancer – Surgical options

MRI – Rectal Cancer



Involved CRM

Uninvolved CRM

British Journal of Surgery 2011; 98: 872-879



Rectal Cancer – Surgical options MRI – Rectal Cancer

One millimetre is the safe cut-off for magnetic resonance imaging prediction of surgical margin status in rectal cancer

F. G. M. Taylor¹, P. Quirke², R. J. Heald⁴, B. Moran⁴, L. Blomqvist⁶, I. Swift¹, S. St Rose⁵, D. J. Sebag-Montefiore³, P. Tekkis⁵ and G. Brown⁵, on behalf of the MERCURY study group

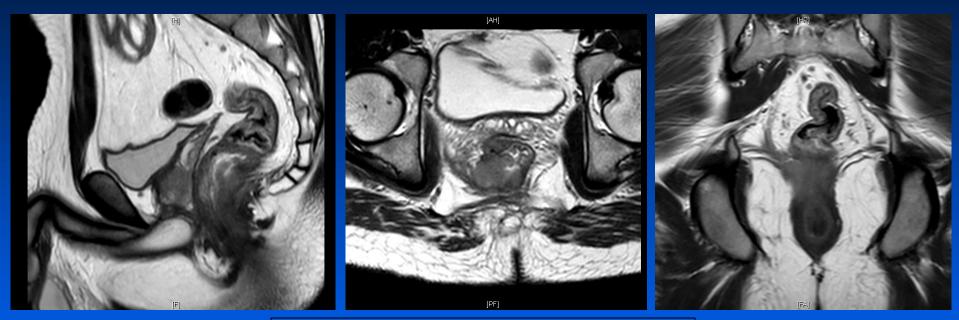
¹Mayday University Hospital, Croydon, ²Pathology and Tumour Biology, Leeds Institute of Molecular Medicine, University of Leeds, and ³St James's Institute of Oncology, St James's University Hospital, Leeds, ⁴Pelican Cancer Foundation, North Hampshire Hospital, Basingstoke, and ⁵Royal Marsden Hospital, Sutton, UK, and ⁶Karolinska University Hospital and Karolinska Institute, Stockholm, Sweden *Correspondence to:* Dr G. Brown, Royal Marsden Hospital, Downs Road, Sutton SM2 5PT, UK (e-mail: gina.brown@rmh.nhs.uk)

British Journal of Surgery 2011; 98: 872–879

MRI (Rectal Protocol) = Preoperative histopathology assessment of margins



Rectal Cancer – Surgical options Rectal Cancer – T3/T4 / N+ – NACTRT



Standard of Care

Decreased local recurrence

Better compliance, higher sphincter preservation

German rectal cancer study group trial, EORTC 22921, NSABP R03

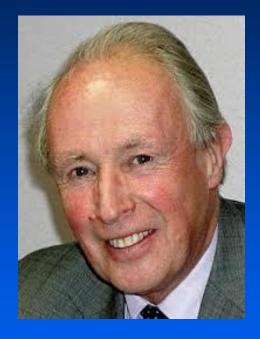
Rectal Cancer – Surgical options Total Mesorectal Excision



1982 - Total mesorectal excision (TME) was introduced as a new surgical technique for rectal cancer.

TME reduced **local recurrence to <5%** and increased **overall survival to 80%** with surgery alone

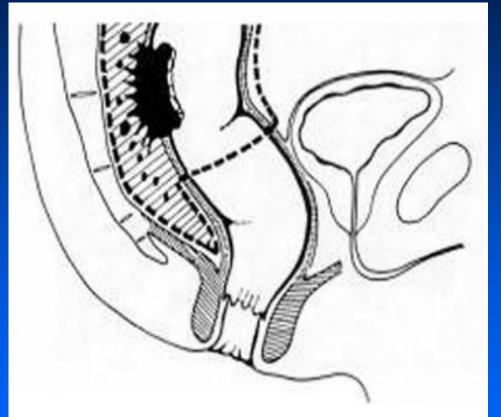
This was much better than any comparable studies even with adjuvant therapy at that time.

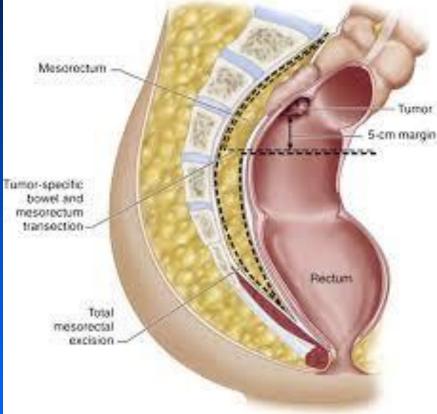


Heald RJ, Ryall RDH. Recurrence and survival after total mesorectal excision for rectal cancer. Lancet1986; i:1479–82.

Rectal Cancer – Surgical options Total Mesorectal Excision – Standard of Care

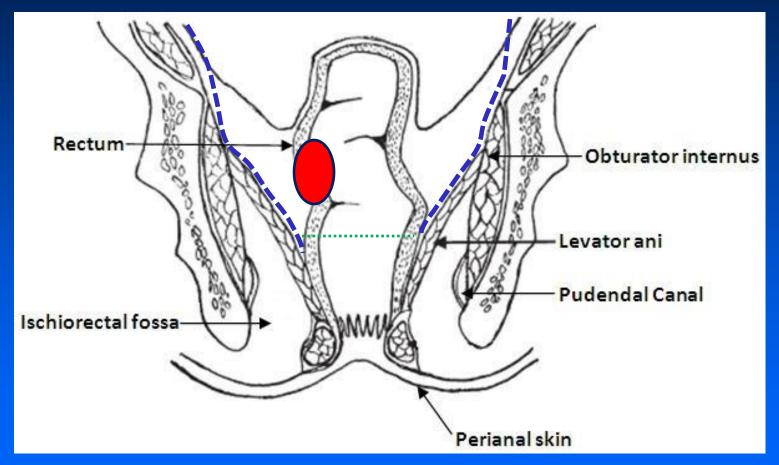






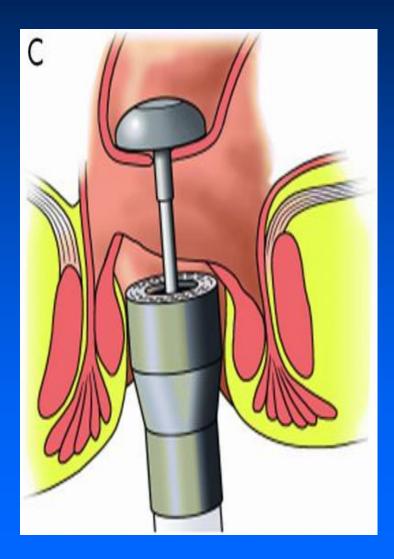
Total mesorectal excision (Mid / Low rectal cancer) Tumour specific mesorectal excision (Upper rectal cancer)



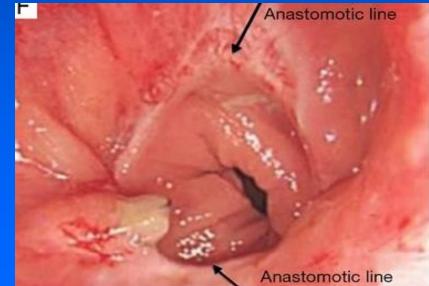


Total Mesorectal Excision (TME) and Anterior resection (Low / Ultralow – double stapling technique)

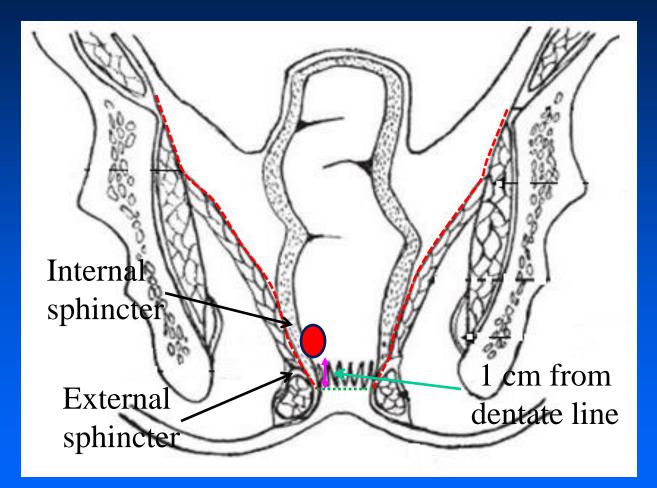












Intersphincteric Resection (ISR)



Intersphincteric resection and hand-sewn coloanal anastomosis for low rectal cancer: Short-term outcomes in the Indian setting

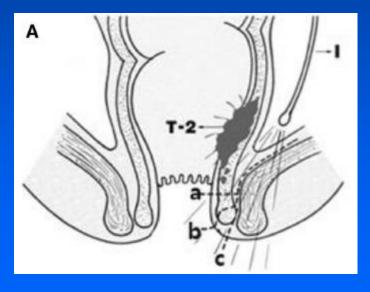
Vishwas D. Pai • Ashwin De Souza • Prachi Patil • Reena Engineer • Supreeta Arya • Avanish Saklani

- First 33 patients of ISR (July 2013 Dec 2013)
- 70% open (91aparoscopic cases, no conversion)
- All distal margins free
- CRM positivity 2 patients.
- Complications 6% (2 patients, ill fashioned ileostomy, urinary retention)

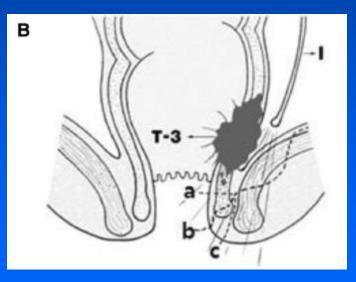


Long-term results of extended intersphincteric resection for very low rectal cancer: a retrospective study

Hyun Sung Kim, Sanghwa Ko and Nahm-gun Oh*



Standard ISR



Extended ISR

Kim et al. BMC Surgery (2016) 16:21



Long-term results of extended intersphincteric resection for very low rectal cancer: a retrospective study

Hyun Sung Kim, Sanghwa Ko and Nahm-gun Oh*

Table 6 Functional results at different times after stoma closure (12 months, 24 months)

	12 months	12 months			24 months		
	Group I	Group II	P value	Group I	Group II	P value	
Stoolfrequency (per day) ^a	3.54 (1.38)	4.29 (1.46)	<0.05	2.21 (1.03)	2.39 (1.12)	0.31	
Kirwan classification ^b			0.86			0.91	
1	14	22		19	25		
Ш	6	10		3	8		
Ш	3	3		1	3		
IV	1	3		1	2		
V	o	0		0	0		
Wexner score ^c	7.33 (2.84)	8.18 (2.91)	0.26	5.21 (1.67)	5.82 (1.93)	0.21	



Sphincter-Preserving Surgery for Low Rectal Cancer: Do We Overshoot the Mark?

Johannes Klose¹ · Ignazio Tarantino¹ · Yakup Kulu¹ · Thomas Bruckner² · Stefan Trefz¹ · Thomas Schmidt¹ · Martin Schneider¹ · Thilo Hackert¹ · Markus W. Büchler¹ · Alexis Ulrich¹

Conclusions ISR is technically feasible with acceptable postoperative morbidity rates. Functional results following ISR are compromised by incontinence as the most important complication. However, long-term quality of life is superior to APR, which should be considered when selecting patients for ISR.

J Gastrointest Surg , Dec 2016



Review

Ann Coloproctol 2018;34(4):167-174 https://doi.org/10.3393/ac.2018.08.02



Annals of Coloproctology pISSN 2287-9714 eISSN 2287-9722 www.coloproctol.org

Intersphincteric Resection for Patients With Low-Lying Rectal Cancer: Oncological and Functional Outcomes

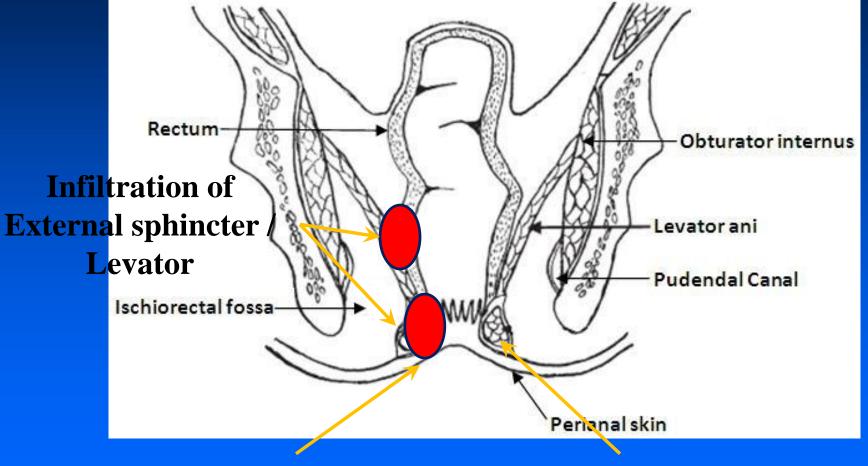
In Ja Park, Jin Cheon Kim

Department of Colon and Rectal Surgery, Asan Medical Center, University of Ulsan College of Medicine, Seoul, Korea

Various researchers have reported diverse continence levels after an ISR:

- a. normal continence (29% to 86.3%)
- b. major incontinence (0% to 25.8%)
- c. need-for-colos-tomy (0% to 0.8%)





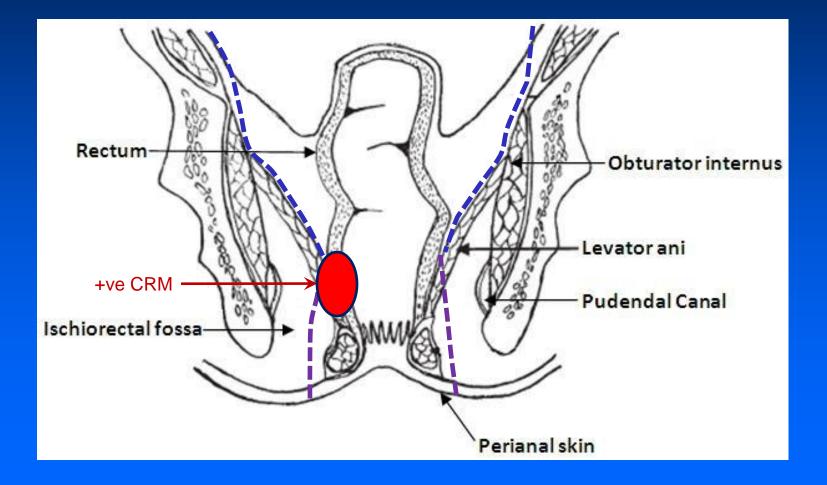
No clear distal margin

Incompetent sphincter



Rectal Cancer – Surgical options

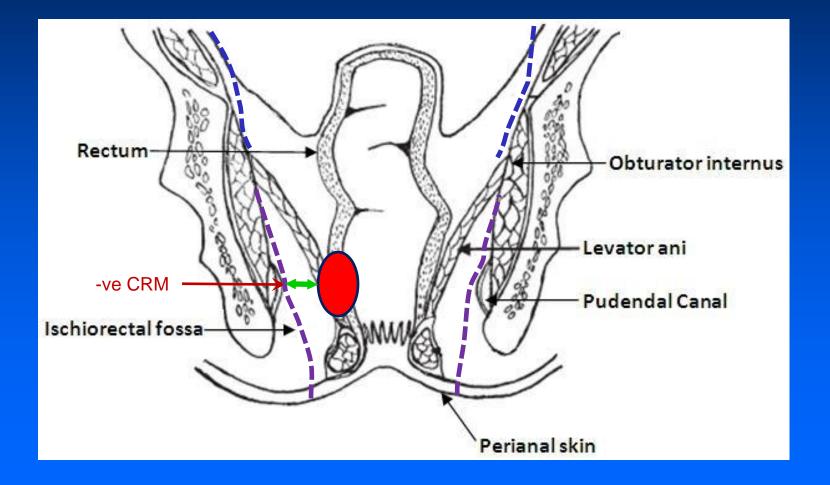
Conventional APR – Potential Issues

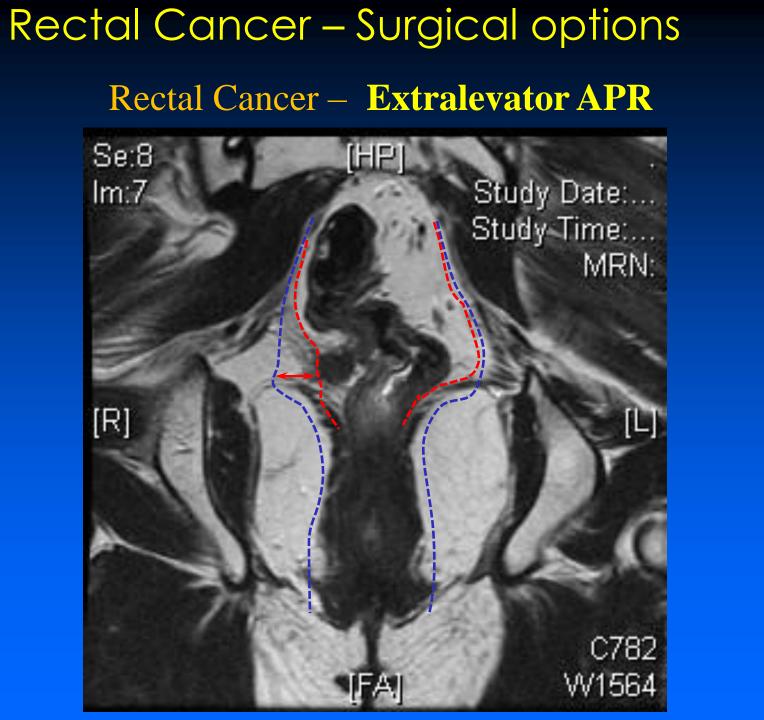


Rectal Cancer – Surgical options



Extralevator APR





Extralevator APR – TMH (Jul 2013 – Jan 2015)

Demographics	Conventional(n = 78)	ELAPER (n= 42)	p value
Age [Median]	47yrs	46 yrs	0.971
Sex			
• Male	53 [2:1]	37 [7:1]	0.011
• Female	25	5	
Histology			
• Adenocarcinoma	67 [86%]	38 [90%]	0.732
• Melanoma	8 [10%]	2[5%]	
• SCC	2 [2.5%]	1[2.5%]	
• GIST	1 [1.5%]	1[2.5%]	
Levator involvement			
• Involved	13 [17%]	22 [52%]	0.000
• Not involved	65 [83%]	20[48%]	
NACTRT			
• Yes	62 [79%]	37[88%]	0.315
• No	16 [21%]	5 [12%]	
Type of surgery			
• Open	44 [56%]	26 [62%]	0.333
• Laparoscopic	30 [39%]	16 [38%]	
Robotic	4 [5%]	0	



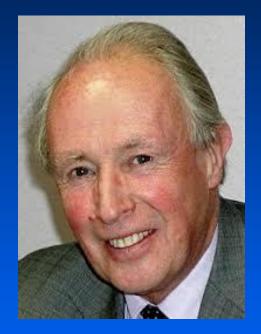
Clinical outcome	Conventional (n = 78)	ELAPER(n= 42)	p value
Blood loss	400 ml	500 ml	0.412
Plastic reconstruction			
• No	75 [96%]	35 [83%]	0.032
• Yes	3 [4%]	7 [17%]	
Mesh placement			
• Yes	0	1 [2.4%]	0.329
• No	0	41	
Wound complications			
• Yes	25 [32%]	8 [19%]	0.141
• No	53 [68%]	34 [81%]	
Hospital stay [Median]	8 days	9 days	0.024

Rectal Cancer – Surgical options Total Mesorectal Excision



1982 - Total mesorectal excision (TME) was introduced as a new surgical technique for rectal cancer.

TME reduced **local recurrence to <5%** and increased **overall survival to 80%** with surgery alone



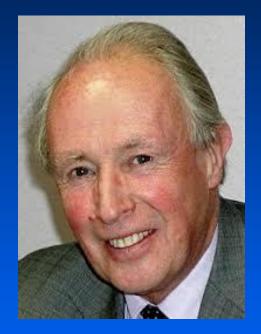
STANDARD OF CARE MINIMUM SURGICAL REQUIREMENT

Rectal Cancer – Surgical options Total Mesorectal Excision



1982 - Total mesorectal excision (TME) was introduced as a new surgical technique for rectal cancer.

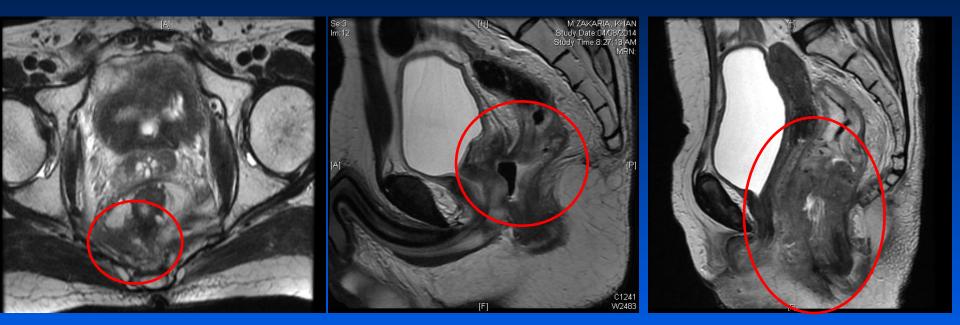
TME reduced **local recurrence to <5%** and increased **overall survival to 80%** with surgery alone



IS TME ENOUGH FOR ALL RECTAL CANCERS ?



Rectal Cancer - IS TME ENOUGH ?



• 5-10% of rectal cancers are T4b at diagnosis

Beyond TME collaborative. Consensus statement on the multidisciplinary management of patients with recurrent and primary rectal cancer beyond total mesorectal excision planes. Br J Surg 2013;100: 1009–14.



Rectal Cancer - IS TME ENOUGH ?

- Involved mesorectal fascia / T4b disease **R0 resection** cannot be achieved with conventional TME.
- For a negative CRM (>1 mm) a multivisceral resection involving en bloc removal of the tumour and adjacent infiltrated organs (beyond-TME)

Nagtegaal ID, Quirke P. What is the role for the circumferential margin in the modern treatment of rectal cancer? J Clin Oncol 2008;26:303–12.

Quirke P, Steele R, Monson J, et al. Effect of the plane of surgery achieved on local recurrence in patients with operable rectal cancer:a prospective study using data from the MRC CR07 and NCIC-CTG C016 randomised clinical trial. Lancet 2009;373:821–8.

Rectal Cancer - IS TME ENOUGH ? Recurrent rectal cancers - 6 to 13% disease recurrence in the pelvis Lopez-Kostner F, Fazio VW, Vignali A, Rybicki LA, Lavery IC. Locally



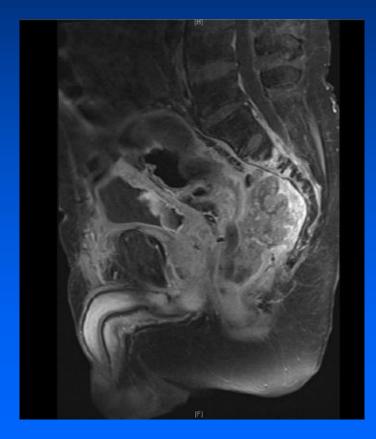
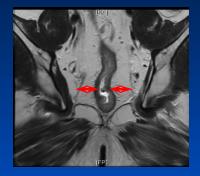
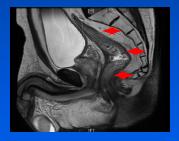


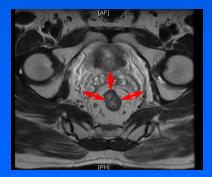
Table 1 Different classifications of recurrent rectal cancers		
Wanebo classific	ation	
TR1-2	Intraluminal recurrency with subserosal infiltration of the colon wall	
TR3	Anastomotic recurrency with infiltration of perirectal soft tissue	
TR4	Invasion in local tissue without fixation	
TR5	Invasion of bony ligaments and structures	
Suzuki-Gunderso	on Classifikation (Mayo Clinic)	
F0	No invasion of local structures	
F1	Invasion of local structures in one direction	
F2	Invasion of local structures in two directions	
F3	Invasion of local structures in three directions	
Memorial Sloan	Kettering Classifikation	
Axial	Anastomotic recurrency, invasion of perirectal tissue and perineum	
Anterior	Invasion of the urogenital tract	
Posterior	Invasion of Os sacrum and presacral fascia	
Lateral	Invasion of the lateral pelvic wall and bony pelvis	



Rectal Cancer - Beyond TME Beyond TME– Surgical Options



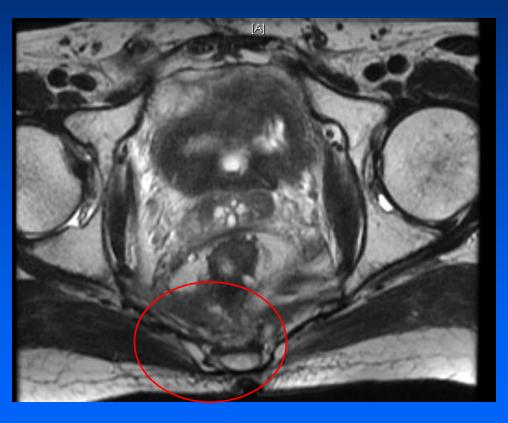


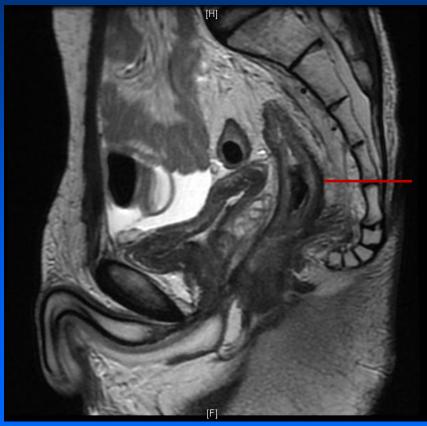


- a. Lateral disease
 - a. Extralevator APR
 - b. Extended Lateral Pelvic Sidewall Excision (ELSiE)
- b. Posterior disease
 a. Sacrectomy High / Low
- c. Anterior disease
 - a. Seminal vesicle / Posterior vagina
 - **b.** Pelvic exenteration
 - c. Pubic bone



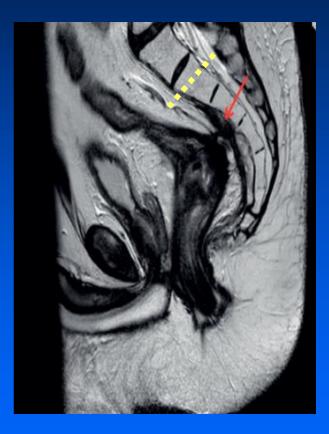
Rectal Cancer - Beyond TME Technical challenges – Posterior Sacral Bone involvement











High sacrectomy - S2/S3, S3



Sacral Resection With Pelvic Exenteration for Advanced Primary and Recurrent Pelvic Cancer: A Single-Institution Experience of 100 Sacrectomies

Tony Milne, B.Sc., M.B.B.S. (Hons.)^{1,2} • Michael J. Solomon, M.B.B.C.H. (Hons.), M.Sc., F.R.A.C.S., F.R.C.S.I.^{1,2,3} • Peter Lee, M.B.B.S., B.Sc., F.R.A.C.S.^{1,2} • Jane M. Young, M.B.B.S., M.P.H, Ph.D., F.A.F.P.H.M.^{1,4} • Paul Stalley, M.B.B.S. (Hons.), F.R.A.C.S., F.A.Orth.A.⁵ James D. Harrison, M.P.H., Ph.D.¹ • Kirk K. S. Austin, B.Sc., A.F.R.C.S.I., F.R.A.C.S.^{1,2}



Rectal Cancer - Beyond TME

Technical challenges – Posterior

Sacrectomy + surgical procedure	
Total pelvic exenteration	68
Bladder-sparing procedure	32
Proximal level sacrectomy	
S2 and above	28
S3 and below	72
Additional bone resection (n = 25)	
Pubis	10
lschium	23
llium	2
Exenteration margin status	
RO	72
R1	22
R2	5



Rectal Cancer - Beyond TME

Technical challenges – Posterior

- Overall complication rate 74%,
 - Major (43%)
 - Minor (67%)
- Bladder-sparing procedure,
 - Urinary retention (28%)
 - Incontinence (19%).
- There was no 30-day or in-hospital mortality.
- Median length of hospital stay 25 days (9–190)



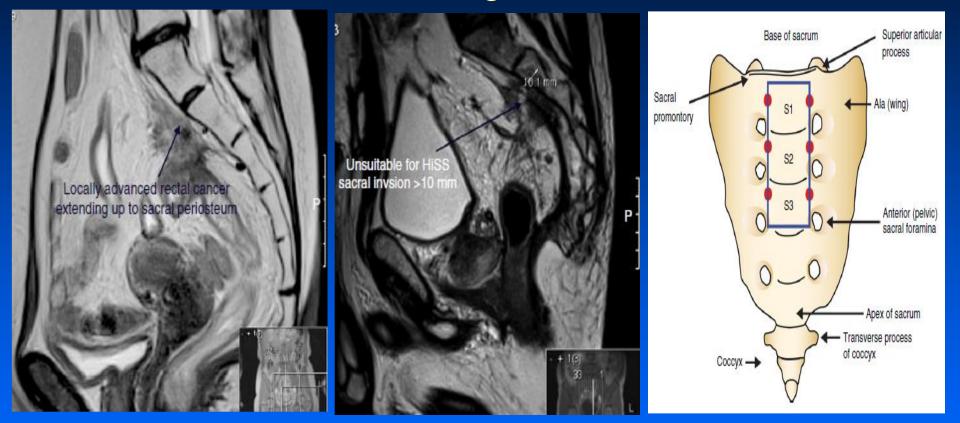
	Low n(%)	High n(%)	р
Median operating time (min)	674	785	0.026
Median blood loss (ml)	3000	7000	< 0.001
R0 margin status	51(71)	21(75)	0.677
30 day mortality	0	0	
Any complication	55(76)	19(68)	0.382
Neurologic deficit	14(19)	12(43)	0.017



High subcortical sacrectomy: a novel approach to facilitate complete resection of locally advanced and recurrent rectal cancer with high (S1–S2) sacral extension

I. Shaikh*, I. Holloway⁺, W. Aston⁺, S. Littler[§], D. Burling[¶], A. Antoniou^{**}, J. T. Jenkins^{**} and On behalf of Complex Cancer Clinic St Mark's Hospital London

Colorectal Dis. 2016 Apr;18(4):386-92.



Contraindications:

- cancellous bone invasion > 10 mm
- disease lateral to the sacral foramina

Colorectal Dis. 2016 Apr;18(4):386-92.

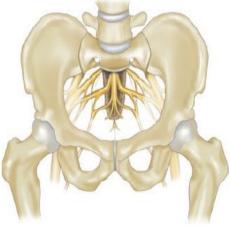


Partial anterior sacrectomy with nerve preservation to treat locally advanced rectal cancer

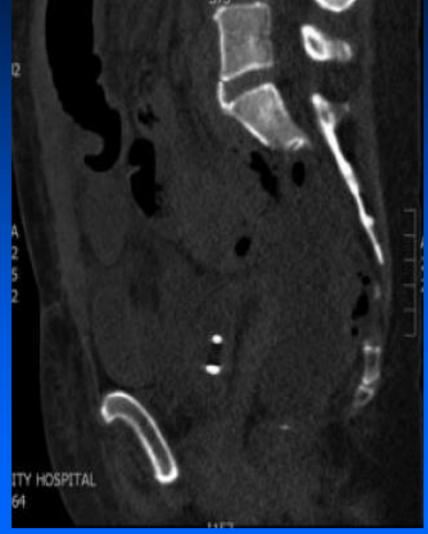
M. D. Evans*, D. P. Harji*, P. M. Sagar*, J. Wilson*, A. Koshy*, J. Timothy† and P. V. Giannoudis‡

*The John Goligher Department of Colorectal Surgery, St James University Hospital, Leeds, UK, †Department of Neurosurgery, The General Infirmary at Leeds, UK and ‡Department of Trauma and Orthopaedic Surgery, The General Infirmary at Leeds, Leeds, UK

Colorectal Dis. 2013 Jun;15(6):e336-9







Colorectal Dis. 2013 Jun;15(6):e336-9

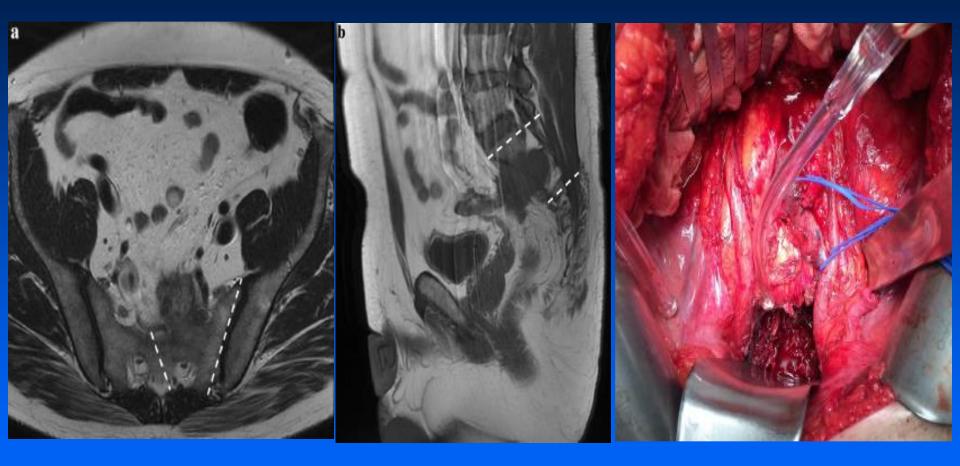




Posterior high sacral segmental disconnection prior to anterior en bloc exenteration for recurrent rectal cancer

K. G. M. Brown^{1,2} · M. J. Solomon^{1,2,3,4} · K. K. S. Austin^{1,3} · P. J. Lee^{1,3} · P. Stalley⁵







Rectal Cancer - Beyond TME Rectal Cancer - Sacrectomy

Outcome of abdominosacral resection for locally advanced primary and recurrent rectal cancer

A. Bhangu^{1,3}, G. Brown², M. Akmal^{1,4} and P. Tekkis^{1,3}

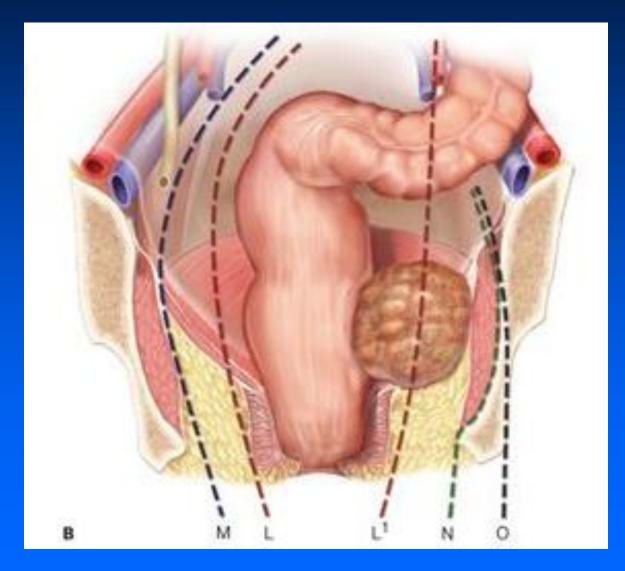
Departments of ¹Colorectal Surgery and ²Radiology, The Royal Marsden Hospital, ³Division of Surgery, Imperial College, Chelsea and Westminster Campus, and ⁴Department of Orthopaedic and Trauma Surgery, Imperial College NHS Trust, London, UK

R0 resection in 23/30 pts – All positive margins in recurrent disease

	RO	R+ (Recurrent)	р
3yr LRFS	87%	0%	<0.001
3yr DFS	71%	0%	0.033

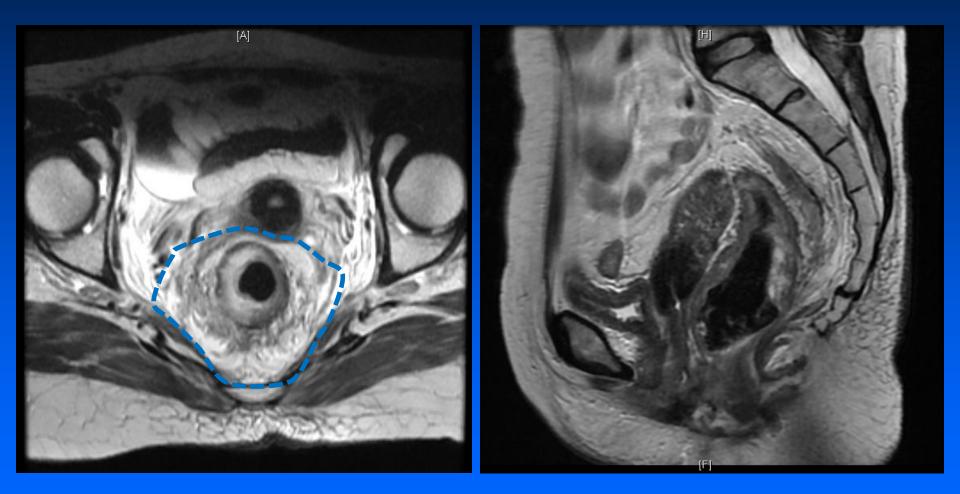
British Journal of Surgery 2012; 99: 1453-1461



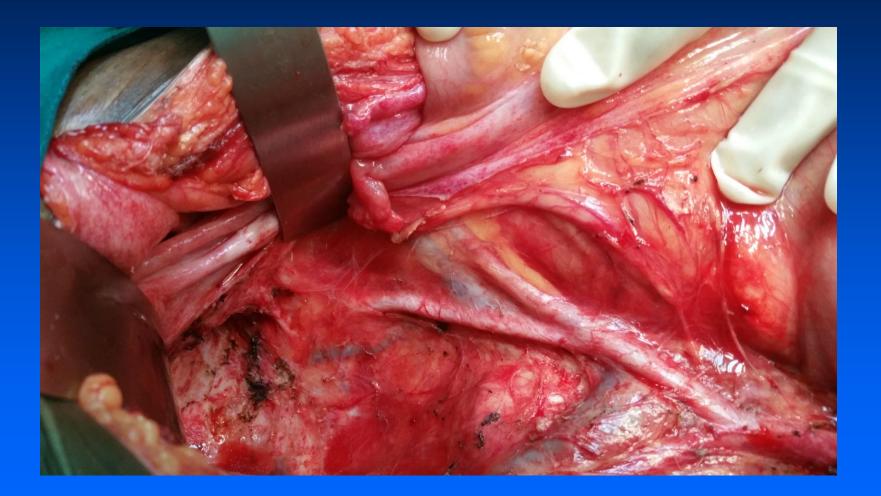


Cancer spread to iliac vessels, pelvic autonomic nerves and ureters, which extends through the greater sciatic foramen with or without invading sciatic nerve





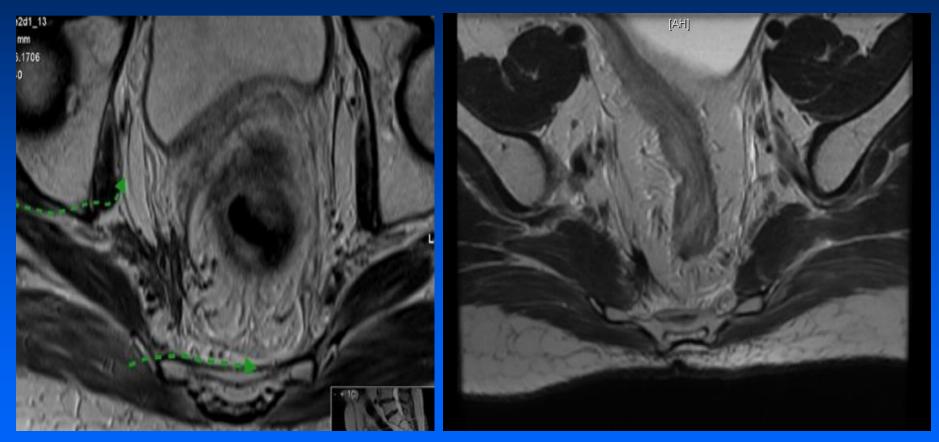






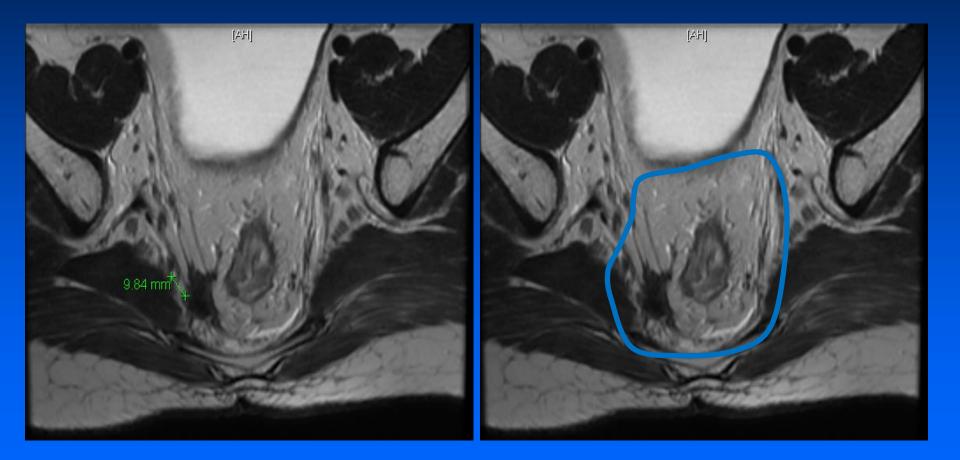




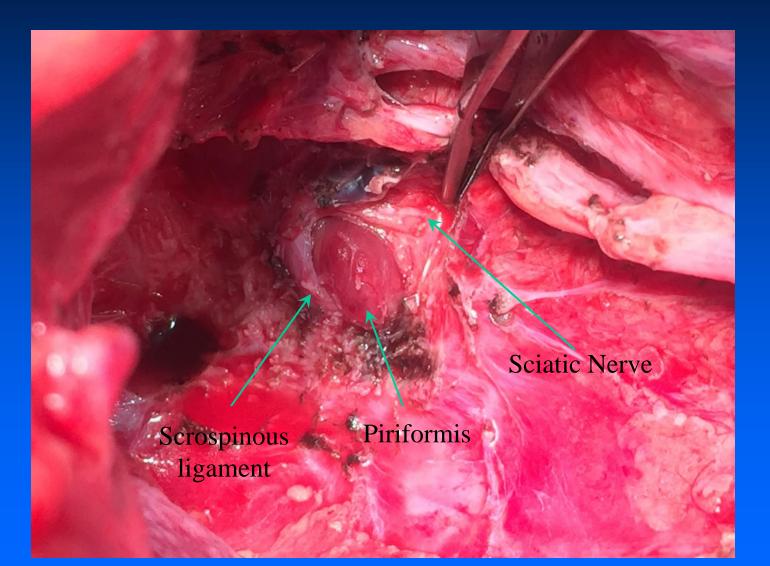


Shaikh et al. Tech Coloproctol (2014) 18:1161–1168











Surgical Option - Pelvic Exenteration

- Pelvic exenteration was first described by Alexander Brunschwig in 1948 in New York as a palliative procedure for recurrent carcinoma of the cervix.
- Thompson and Howe reported the first case of complete pelvic evisceration for locally advanced rectal cancer in 1950



Rectal Cancer - Pelvic Exenteration

The primary justification of such radical surgery is the reasonable chance of cure, which is now achievable in up to 63% of patients

You YN, Roses RE, Chang GJ, et al. Multimodality salvage of recurrent disease after local excision for rectal cancer. Dis Colon Rectum. 2012;55:1213–1219.

Harris CA, Solomon MJ, Heriot AG, et al. The outcomes and patterns of treatment failure after surgery for locally recurrent rectal cancer. Ann Surg. 2016;264:323–329.

Hansen MH, Balteskard L, Dørum LM, Eriksen MT, Vonen B; Norwegian Colorectal Cancer Group. Locally recurrent rectal cancer in Norway. Br J Surg. 2009;96:1176–1182.

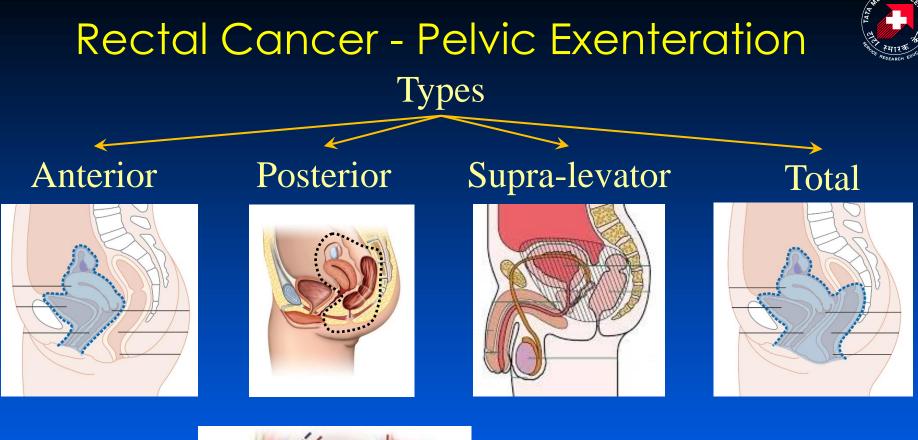


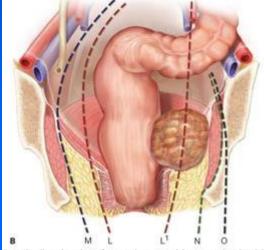


Determinants of survival following pelvic exenteration for primary rectal cancer

R. W. Radwan, H. G. Jones, N. Rawat, M. Davies, M. D. Evans, D. A. Harris and J. Beynon, on behalf of Swansea Pelvic Oncology Group

	Median OS (months)	5 yr survival
R0 Resection	121	59.3%
R1 Resection	24	23%
R0 (local) in Resectable metastatic disease	18	0%





± pelvic sidewall / Sacrum / Bone pelvis



Reconstruction

- VRAM Vertical Rectum Abdominis myocutaneous flap
- Inferior gluteal artery myocutaneous flap
- Gracilis flap
- Anterior-lateral thigh flap



Rectal Cancer - Pelvic Exenteration Reconstruction



Bilateral Gluteus VY advancement flap



Surgical and Survival Outcomes Following Pelvic Exenteration for Locally Advanced Primary Rectal Cancer

Results from an International Collaboration

The PelvEx Collaborative

(Ann Surg 2017;xx:xxx-xxx)

Factors affecting outcomes following pelvic exenteration for locally recurrent rectal cancer

The PelvEx Collaborative*

*Members of the PelvEx Collaborative are co-authors of this article and can be found under the heading Collaborators Correspondence to: Dr M. E. Kelly, Centre for Colorectal Disease, Department of Surgery, St Vincent's University Hospital, Elm Park, Dublin 4, Ireland (e-mail: kellym11@tcd.ie; @@PelvExGroup) 2018 BJS Society Ltd



- A retrospective international observational cohort study to assess the outcomes of patients undergoing pelvic exenteration for LARC in a 10-year period (from 2004 to 2014)
- Twenty-seven international institutions provided data, with each center being a tertiary referral unit with specialist expertise in advanced colorectal cancer.





TABLE 1. Characteristics of Patients Included in the Study

Characteristics

Age (y)	
Median (IQR)	63 (17)
Gender: N (%)	
Male	778 (60.3%)
Female	513 (39.7%)
BMI	
Median (IQR)	24 (6)
Neoadjuvant therapy: N (%)	
Yes	1008 (78.1%)
No	129 (10.0%)
Unknown	154 (11.9%)



TABLE 1. Characteristics of Patients Included in the Study		
Characteristics		
Type of exenteration: N (%)		
Total	551 (42.6%)	
Posterior	529 (41.0%)	
Anterior	30 (2.3%)	
Modified	139 (10.8%)	
Unknown	42 (3.3%)	
Duration of surgery (min)		
Mean (SD)	433.2 (184.7)	
Nodal yield		
Median (IQR)	14 (14)	
Margin status: N (%)		
RÔ	1030 (79.8%)	
R1	172 (13.4%)	
R2	29 (2.2%)	
Unknown	60 (4.6%)	



TABLE 2. Post Exenteration Length ofRates, Morbidity, and Mortality	Stay,	Readmission
Postoperative Characteristics		
Length of hospital stay (d)		
Median (IQR)		16 (14)
Readmission within 30 d: n (%)		
Yes		95 (7.4%)
No		1196 (92.6%)
Major complications within 30 d: n (%)		
Yes		488 (37.8%)
No		803 (62.2%)
Surgical reintervention: n (%)		
Yes		111 (8.6%)
No		1180 (91.4%)
Radiological reintervention: n (%)		
Yes		78 (6.0%)
No		1213 (94.0%)
30-d mortality: n (%)		
Yes		19 (1.5%)
No		1272 (98.5%)



	RO	R1	R2	р
Median survival (months)	43	21	10	< 0.001
3yr overall survival	56.4	29.6	8.1	< 0.001

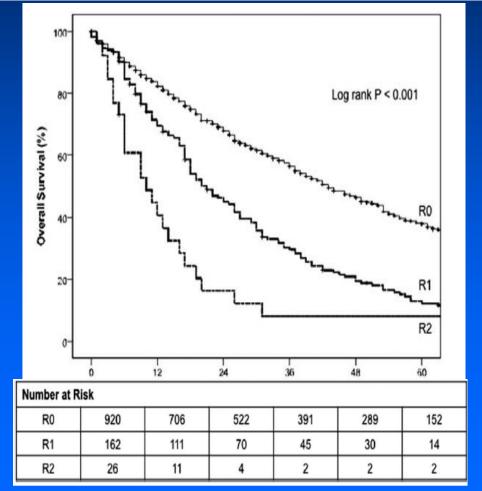




TABLE 4. Univariable and Multivariable Analyses of Factors That Influence Survival

						variable Cox Model
	Median OS (mo)	3-y OS (%)	5-y OS (%)	Univariable P	HR	Р
Margin status (N = 1147)						
R0 (n = 956)	43	56.4	37.8	< 0.001		
R1 ($n = 165$)	21	29.6	12.3		1.80^{*}	< 0.001
R2 (n = 26)	10	8.1	<8.1		3.1**	< 0.001
Neoadjuvant (N = 1029)						
Yes $(n = 910)$	36	49.6	31.3	0.189		NS
No $(n = 119)$	26	39.9	23.5			
Bone resection $(N = 855)$						
Yes $(n = 90)$	29	40.3	29.4	0.383		NS
No $(n = 765)$	37	50.3	31.6			
Nodal status ($N = 856$)						
Positive $(n = 302)$	31	44.3	28.9	< 0.001	1.27	0.009
Negative $(n = 554)$	46	58.0	39.7			



Factors affecting outcomes following pelvic exenteration for locally recurrent rectal cancer

The PelvEx Collaborative*

*Members of the PelvEx Collaborative are co-authors of this article and can be found under the heading Collaborators Correspondence to: Dr M. E. Kelly, Centre for Colorectal Disease, Department of Surgery, St Vincent's University Hospital, Elm Park, Dublin 4, Ireland (e-mail: kellym11@tcd.ie;) @PelvExGroup)

Rectal Cancer - Pelvic Exenteration

No. of patients* (n = 1184)



Age (years)† 63 (56-69) Sex ratio (M:F) 752:432 BMI (kg/m²)† 25(22-28)Neoadjuvant therapy Yes 614 (51.9) No 515 (43.5) Unknown 55 (4.6) Type of neoadjuvant therapy Chemoradiotherapy 463 (75.4) Radiotherapy alone 54 (8.8) Chemotherapy alone 61 (9.9) Unknown 36 (5.9) Type of exenteration Total <u>418 (35·3)</u> Posterior <u>395 (33·4)</u> Anterior 80 (6.8) Modified 91 (7.7) Unknown 200 (16.9)

BJS 2018



	No. of patients* ($n = 1184$)
Bone resection	
Yes	240 (20.3)
No	944 (79.7)
Duration of surgery (min)‡	509(201)
Blood transfusion	
Yes	372 (31.4)
No	812 (68·6)
No. of units transfused†	5 (1-8)
Nodal yield†	5 (1–10)
Margin status	
R0	656 (55.4)
R1	365 (30.8)
R2	87 (7.3)
Unknown	76 (6.4)



 Table 3 Postoperative duration of hospital stay and complications

	No. of patients* ($n = 1184$)
Duration of hospital stay (days)†	15 (10–26)
Readmission within 30 days	
Yes	70 (5.9)
No	1114 (94.1)
Major complications within 30 days	
Yes	380 (32.1)
No	804 (67.9)
Inpatient at 30 days	
Yes	179 (15.1)
No	867 (73.2)
Unknown	138 (11.7)
Reintervention	
Yes	118 (10.0)
Only surgical	63
Only radiological	33
Both surgical and radiological	22
No	1066 (90.0)
30-day mortality	
Yes	21 (1.8)
No	1163 (98.2)



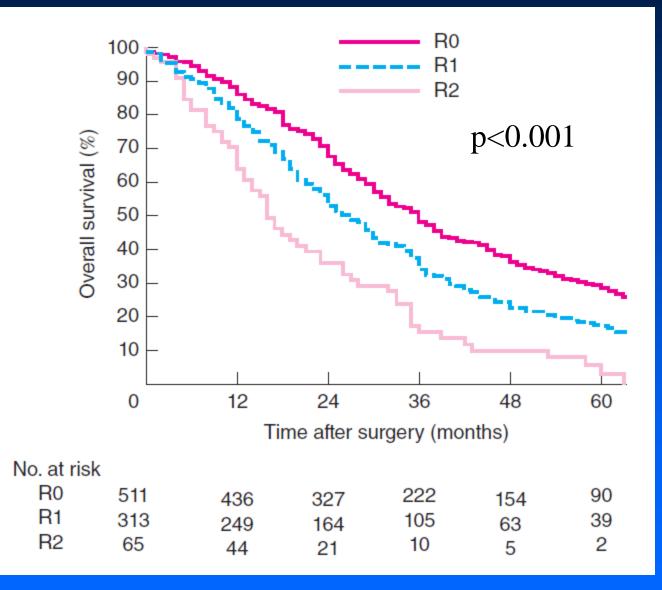




Table 4 Univariable and multivariable analyses of factors that influenced survival of pelvic exenteration for locally recurrent rectal cancer

	Median overall	3-year overall	5-year overall	Univariable	Multivariable Co	x regression
	survival (months)	survival (%)	survival (%)	P*	Hazard ratio	Р
Margin status ($n = 889$)				< 0.001		
R0 (n = 511)	36	48.1	28.2		1.00 (reference)	
R1 (n = 313)	27	33.9	17.3		1·28 (0·97, 1·69)	0.076
R2 $(n = 65)$	16	15	3		4.84 (2.77, 8.48)	< 0.001
Neoadjuvant therapy ($n = 913$)				0.008		
Yes (n = 530)	32	43.5	25.6			0.260
No (<i>n</i> = 383)	27	34.1	16.4			
Bone resection ($n = 825$)				< 0.001		
Yes (n = 184)	36	48.9	35.3		0.74 (0.55, 0.99)	0.049
No (<i>n</i> = 641)	29	38.8	19.0		1.00 (reference)	
Node status ($n = 337$)				0.014		
Positive $(n = 76)$	22	21	11			0.164
Negative ($n = 261$)	29	38.0	22.8			





- There was a significant difference in margin status according to whether patients underwent bone resection (where required)
- R0 resection rate was 67.4% among patients who had bone resection and 56.2% in those who did not (*P*=0.006).



- Patterns of failure following Sx for recurrent rectal cancer
- Local recurrence alone in 14%
- Systemic metastases with or without local recurrence in 42%.
- Chemoradiotherapy before exenteration was associated with a significant (P < 0.05) improvement in overall 5-year cancer-specific survival for patients with an R0 resection.

Harris Ca et al. Ann Surg 2016 Aug;264(2):323-9.



A 10-year experience of total pelvic exenteration for primary advanced and locally recurrent rectal cancer based on a prospective database

M. B. Nielsen*, P. C. Rasmussen*, J. C. Lindegaard⁺ and S. Laurberg^{*}

*Departments of Surgery and †Oncology, Aarhus University Hospital, Aarhus, Denmark



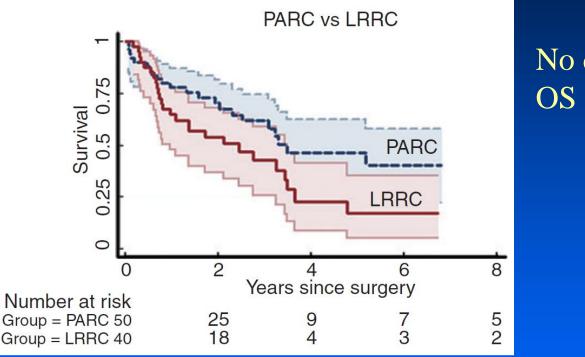
	Primary $(n = 50)$	Recurrent $(n = 40)$	Р
Desertion			
Resection			
TPE with sacral resection	5	15	0.002
Reconstruction			
VRAM	28	30	
Gluteal	2	2	
Radicality			
Complete resection (R0)	33	15	0.007
Microscopic incomplete (R1)	17	20	
Macroscopic incomplete (R2)	0	5	
Duration of surgery (min) (median, range)	296 (129-495)	395 (210-730)	0.000
Hospital stay (days) (median, range)	13 (4–51)	15 (9–71)	0.16



		Primary $(n = 50)$	Recurrent $(n = 40)$	Р
No complication	44	26	18	0.51
Any complication	46	24	22	

The 5-year DFS was 25.9% (11.4–43.2) for PARC and 22.0% (10.2–36.6) for LRRC (P = 0.02).





No difference in OS (p=0.16)

There was no statistically significant difference in OS between PARC and LRRC when comparing R0 resections (P = 0.20) or R1/R2 resections (P = 0.96)Colorectal disease 2011



Outcomes of pelvic exenteration for recurrent and primary locally advanced rectal cancer

Matteo Rottoli*, Carlo Vallicelli, Luca Boschi, Gilberto Poggioli

Surgery of the Alimentary Tract, Sant'Orsola - Malpighi Hospital, Alma Mater Studiorum University of Bologna, Bologna, Italy

International Journal of Surgery 48 (2017) 69-73



Variable	ARC (28)	RRC (18)	р
Male gender	12 (42.9%)	12 (66.7%)	0.12
Age	59 (29-86)	55 (31-76)	0.71
Squamous cell carcinoma	6 (21.4%)	2 (11.1%)	0.41
ASA score 3	17 (60.7%)	12 (66.7%)	0.90
Neoadjuvant therapy	20 (71.4%)	10 (55.5%)	0.25
Intraoperative blood loss (mL)	600 (300-4000)	750 (265-2700)	0.74
Number of resected compartments			0.43
2	22 (78.6%)	13 (72.2%)	
3	6 (21.4%)	4 (22.2%)	
4	0	1 (5.6%)	
Sacrectomy	5 (17.9%)	4 (22.2%)	0.74
Flap reconstruction	9 (32.1%)	2 (11.1%)	0.14
Duration of surgery (min)	310 (180-612)	305 (175-745)	0.73
Radicality of resection			0.41
R0	20 (71.4%)	10 (55.6%)	
R1	7 (25%)	6 (33.3%)	
R2	1 (3.6%)	2 (11.1%)	

International Journal of Surgery 48 (2017) 69–73



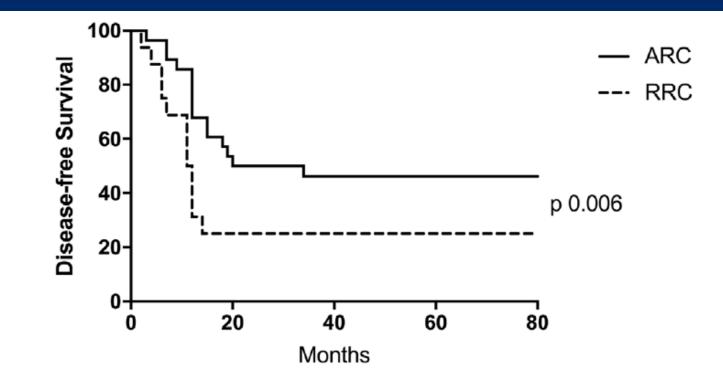


Fig. 1. Comparison <u>of disease-free survival</u> between patients with locally advanced rectal cancer (ARC) and locally recurrent rectal cancer (RRC) undergoing pelvic exenteration.

International Journal of Surgery 48 (2017) 69–73



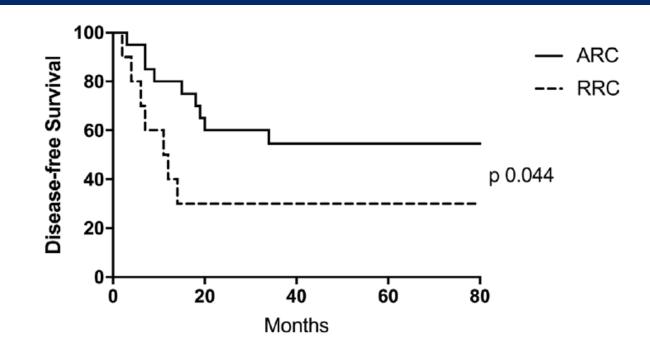


Fig. 2. Comparison of disease-free survival between patients with locally advanced rectal cancer (ARC) and locally recurrent rectal cancer (RRC) undergoing pelvic exenteration including only R0 resections.

International Journal of Surgery 48 (2017) 69–73



Rectal Cancer - Pelvic Exenteration Outcomes – Quality of Life

A systematic review examining quality of life following pelvic exenteration for locally advanced and recurrent rectal cancer

E. Rausa*†, M. E. Kelly*, L. Bonavina†, P. R. O'Connell*‡ and D. C. Winter*‡

*Department of Colorectal Surgery, St Vincent's University Hospital, Elm Park, Dublin, Ireland, †Department of Surgery, IRCCS Policlinico San Donato, University of Milan Medical School, San Donato Milanese (Milano), Italy and ‡Section of Surgery, UCD School of Medicine, Dublin, Ireland



Rectal Cancer - Pelvic Exenteration Outcomes – Quality of Life

- The median compliance (range) in fully answering the QoL questionnaires was 77% (62–100%)
- Median follow up time 12-24 months
- QoL began to return to pre-surgical levels
 - 2–3 months in two studies
 - 6 9 months in two studies.
- 1 study (Choy et al) QOL improved by 9 months, baseline never fully restored in those with LRRC
- Difference in QOL between R0 and R1 resections not consistent



Rectal Cancer - Pelvic Exenteration Outcomes – Quality of Life

- Comparing QOL between APR Vs TPE
- Austin et al. observed similar QOL scores at 3 months post-surgery.
- Radwan et al. reported significant difference
 - regarding physical (P = 0.010), role (P = 0.047), emotional (P = 0.010) and social functional level (P = 0.005) over the first 3 months in favour of APR. However, **this difference dissipated by the fourth month after surgery**
- Women reduced QoL after exenteration (P = 0.04)
- Patients with vaginectomy significantly reduced QoL vis-a-vis vaginectomy plus vaginal reconstruction Colorectal disease 2017



June 2013 – Feb 2018 102 Pelvic Exenterations

Histology 97 Adenocarcinoma 2 SCC 1 melanoma 1 Neuroendocrine 1 GIST

83 Primary Rectal Cancer19 Recurrent Rectal Cancer



	Number (n=102)
Age (years)	43 (19-69)
Males	51%
BMI (Kg/m2)	22.43 (14.9 - 33.2)
NACTRT (79 primary, adenoca)	75.9%
Approach Open Laparoscopic Robotic	83 (81.4%) 14 (13.7%) 5 (4.9%)
Procedure Total Pelvic Exenteration Posterior Extention Supralevator exenteration Lateral Pelvic Node dissection	54 (52.9%) 40 (39.2%) 8 (7.8%) 23.5%



102 Pelvic Exenterations

	Number		
ASA 1/2	99%		
Blood loss (ml)	1400 (150 - 4500)		
Hospital Stay (days)	12 (5-71)		
All complications	52%		
Clavien Dindo			
Grade 1/2	33.3%		
Grade 3/4	17.6%		
Grade 5	1%		



102 Pelvic Exenterations

Pathological Outcomes	Number		
R Status			
R0	86.3%		
R1	6.9%		
R2	6.9%		
pT4	41.2%		
Total nodes	14.38(11.5)		
pN+ disease	36.2%		

Median follow up 11.2 months

Rectal Cancer - Pelvic Exenteration



R0 resection is the holy grail of pelvic exenteration

Pelvic extenteration is only worth it if it is R0 !



Stage IV (metastatic) disease – **Peritoneal disease**

Incidence
Primary cancer – 5-10%
Recurrent cancer – 15-30%

•**Recurrent colorectal cancer** – Peritoneum is the only site of recurrence in 15-20%

Conventional treatment – systemic chemotherapy
Median survival – 9 months
Addition of Bevacizumab/cetuximab – 19-20 months



Stage IV (metastatic) disease – Peritoneal disease

Complete Cytoreductive Surgery Plus Intraperitoneal Chemohyperthermia With Oxaliplatin for Peritoneal Carcinomatosis of Colorectal Origin

Dominique Elias, Jérémie H. Lefevre, Julie Chevalier, Antoine Brouquet, Frédéric Marchal, Jean-Marc Classe, Gwenaël Ferron, Jean-Marc Guilloit, Pierre Meeus, Diane Goéré, and Julia Bonastre

	2yr OS	5 yr OS	Median Survival (months)
Systemic chemotherapy ± Palliative surgery	65%	13%	23.9
CRS + HIPEC + Systemic chemotherapy	81%	51%	62.7 (p<0.05)

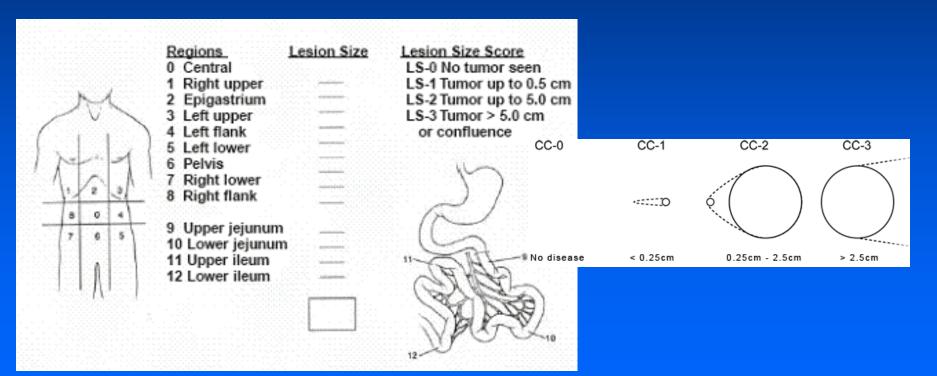
Highly selected patients

J Clin Oncol 2009 Feb 10;27(5):681-5.



Stage IV (metastatic) disease – Peritoneal disease CRS + HIPEC – Patient selection

• Morbidity (23%-45%); Mortality (0-12%)



•Peritoneal carcinomatosis index (PCI)



Stage IV (metastatic) disease – Peritoneal disease CRS + HIPEC – Patient selection

•PCI

- •PCI <12 most favourable results
- •PCI >17-20 no benefit Vs Systemic chemotherap •ECOG 0-1
- •No evidence of extra-abdominal disease
- •Upto 3 small resectable liver metastasis
- •No evidence of biliary obstruction
- No evidence of ureteral obstruction
- •No evidence of intestinal obstruction at > 1 site
- •Small bowel no gross disease in mesentery / multi level partial obstruction
- •Small volume disease in the gastrohepatic omentum.



Colorectal Peritoneal Metastasis Stage IV (metastatic) disease – Peritoneal disease CRS + HIPEC – Unresolved issues

• Complete CRS offers best results (possible in low PCI) - ?Role of HIPEC itself (PRODIGE 7)

• HIPEC methodology

- Drugs / combination
- Doses
- Temperature
- Contact time
- Volume and composition of perfusion solution.
- Open / Closed technique
- Bidirectional chemotherapy
- Role of EPIC
- Preventing peritioneal metastasis in high risk (pT4, PCI 0)



Rectal Cancer – Surgical Options Summary

- Total Mesorectal excision (TME) standard approach for ALL radical rectal surgery
- Tumour specific TME Upper rectal tumours
- Sphincter preservation wherever possible
- Intersphincteric resection better than APR
- Extralevator APR if levator involved
- Beyond TME pelvic side wall, sacrectomy
- Sacrectomy for recurrent disease poor outcomes
- Pelvic Exenteration worth it if R0
- CRS+HIPEC good outcomes in selected cases



Thank you !



Division of Gastrointestinal and HPB Oncology Tata Memorial Centre, Mumbai