

Gateways for management of advanced colorectal cancer

Ahmed Mostafa, MD Professor of surgical oncology NCI, Cairo University.

#### **Advanced CRC**

- T4 a/b CRC Tumor adherent or direct invading adjacent organ or structure.
- 10% of all resected colon cancer
- Incidence higher in emergent cases than elective cases (70% vs 25%)
- Higher positive margin rate
- Biologic more aggressive MSI-H/ mucinous



#### Radiological

#### evaluation of cT4

| [       | Accuracy | Sensitivity | Specificity |
|---------|----------|-------------|-------------|
| CT scan | 70-77%   | 17-25%      | 82-93%      |
| PET/CT  | 80-82%   | 50-58%      | 86-91%      |



#### <u>assessed T4</u> were confirmed pT4 histological, tendency of over diagnosis



## Confusing/misleading PET



#### Principles Surgical Management

- En block resection of contiguous structure either attachment, adhesions or infiltration,
- Don't disrupt the plane of adhesions as 34%-84% are malignant.

(Gezan et al. 2012, Eveno et al. 2014)



#### Principles Surgical Management

- **Stenting** distal lesions as bridge to definitive resection.
- MIS approach is still an option with smart approach and higher conversion rate.
- Post operative radiation for resected tumors with positive margin is considered after placing surgical clips at resected tumor bed.





#### Principles Surgical Management

## Journal of Clinical Oncology®

An American Society of Clinical Oncology Journal

Meeting Abstract | 2019 ASCO Annual Meeting I

GASTROINTESTINAL (COLORECTAL) CANCER

**Conclusions:** Six weeks of NAC for operable CC can be delivered safely, with marked histopathological down-staging, and may result in better disease control at 2 years in pMMR disease. **28% lower event rate** 

 Consider neoadjuvant chemotherapy in clinical T4b

#### Microsatellite status











#### Gateways for locally advanced tumors



#### Lateral Pelvic Pathways

1- Pararectal space divided by the ureter
2-Paravesical space divided by the obliterated umbilical artery

**3-Triangle of Marcille** 

# Denonvilliers

## Waldeyer



Urogenital hypogastric fascia





#### Pararectal space: divided by the ureter

The medial pararectal space Okabayashi space contains the superior hypogastric nerve .

The lateral pararectal space <u>Latzko</u> space is the best space for dissection of the uterine artery





#### The para vesical space divided by OUA

Paravesical space is divided by umbilical artery into lateral (obturator space) and medial spaces and separated from the pararectal space by the uterine artery



#### Triangle of Marcille

#### Vessel s:

1.common iliac artery 1- common iliac artery ex il a - external iliac artery in il a- internal iliac artery 2/ex il v- external iliac vein ill u a- ililumbar artery I s a- lateral sacral artery

#### Nerves:

L4 L5 n- Lumbosacral trunk S1- First sacral nerve meets Lumbosacral trunk (inferior to the triangle) Ob n- obturator nerve gf n- genitofemoral sym t- sympathetic nerve from the trunk to the superior hypogastric plexus

Other: Ur Ureter L5- 5<sup>th</sup> lumbar vertebra



#### (Triangle of Marcille)

## Anterior Pelvic Pathways

1- <u>Vesico</u>uterine

2- <u>Recto</u>uterine





Rectovaginal space Rectovesical pouch Posterior pelvic Pathway











## Laparoscopic approach

## The laparoscopic approach

**Below the IC pedicle** 

**Below the IMV** 





Beyond TME Don't force dissection plane

 The situation where the tumor extends beyond the TME plane, so TME is not enough, enblock resection with urinary bladder, ureter, distal sacrum or lateral pelvic wall, otherwise you will declare the operation as non-curative, as you come across tumor that you was not expecting to see.



## Exentrations

#### T4 risk of PC

| Independent predictors<br>for metachronous PC | HR   | 95% CI     | <i>P</i> = |
|---|------|------------|------------|
| T4  | 9.98 | 3.10-32.11 | <0.001     |
| N2 with <12 nodes exam                        | 7.41 | 4.78-11.51 | <0.001     |
| R2 resection                                  | 2.75 | 2.10-3.61  | <0.001     |
| Right side                                    | 1.77 | 1.31-2.39  | 0.002      |
| <b>Emergency surgery</b>                      | 2.11 | 1.66-2.69  | <0.001     |
| Age >70                                       | 0.69 | 0.55-0.87  | 0.003      |

Segelman et al. Br J Surg 2012

Magnitude of the problem **<u>10</u>%** of cases present with synchronous carcinomatosis.

20 % of patients will develop metachronous disease at follow up.

<u>5 %</u> PC is the sole pattern of recurrence.

## Solution of the problem

Median survival of <u>6 months</u> in untreated cases

Modern chemotherapy and targeted agents the median overall survival has dramatically improved up to <u>24 months</u>.

**CRS** and HIPEC improved median survival up to <u>40-60</u> months.

## Evolution of median survival PC of CRC

|                |                            | Median survival                                   |
|----------------|----------------------------|---|
| Before<br>1990 | Systemic<br>chemotherapy   | 6 months<br>16 months with modern<br>chemotherapy |
| 1990-<br>2000  | Verwal 2003<br>Glehan 2004 | 20 months   |
| 2000-<br>2010  | Elias 2010                 | 30 months   |
| 2010-<br>2020  | Quenet ASCO 2018           | 40 months   |

## (CRS <u>&</u> HIPEC) vs Systemic

A randomized study



Surgery + HIPEC > Systemic chemotherapy

Verwall et al. J Clin Oncol 2003, Ann Surg 2008

#### (CRS <u>&</u> HIPEC) vs Systemic chemptherapy



#### (CRS <u>&</u> HIPEC) vs Systemic chemptherapy

1013 patients

CT-PCI PC1 < 10.0.496)

PCT 15-35 (Medium)

PCE > 30 (High)

and/or Lot milds Yo

No systems

MOLD CYCRIPTICATE

on symptom

#### Peritoneal Surface Disease Severity Score (PSDSS)

American Society of Peritoneal Surface Malignancies 1 013 patients Median Survival (months)

| PSDSS   | Chemotherapy<br>alone | Cytoreductive<br>surgery and HIPEC |
|---------|-----------------------|------------------------------------|
| PSDSS 1 | 45                    | 86                                 |
| PSDSS 2 | 19                    | 43                                 |
| PSDSS 3 | 8                     | 29                                 |
| PSDSS 4 | 6                     | 28                                 |

Ann Surg Oncol 2014





#### Cure 16%

#### Peritoneal metastasis from colorectal cancer

- Survival at 5 years withour recurrence : 16%
  - We can cure PM from colorectal cancer

Goere et al Ann Surg 2012

- Median survival > 60 months
  - Strict selection of patients
  - Systemic chemotherapy

Passot et al. Ann Surg 2012

Selection of patients with (localized disease, well differentiated, no clinical symptoms) CRS and HIPEC do better





#### Is it CRS or HIPEC



#### **Positive Lessons from Negative trial**

#### **Overall survival (ITT)**



Median Follow Up: 64 months [95% CI:58.9-69.8]

|   | HIPEC                   | Non-HIPEC                  | P-value |
|---|-------------------------|----------------------------|---------|
| Median Survival<br>(months)<br>[95% CI] | <b>41.7</b> [36.2-52.8] | <b>41.2</b><br>[35.1-49.7] | 0.995   |
| 1-year Survival                         | 86.9%                   | 88.3%                      |         |
| 5-year Survival                         | 39.4%                   | 36.7%                      |         |

HR=1.00: 95%CI [0.73 - 1.37] p=0.995

# Conclusion

- Pelvic spaces are gate ways to navigate safely during advanced oncologic procedures.
- Familiarity and surgical competence in dealing with cytoreductive procedures are prerequisite before dealing with advanced CRC.
- Neoadjuvant treatment or postoperative irradiation may improve outcome in selected cases.





#### duodenum



#### Pancreas







#### Umbilicovesical fascia (UVF)



#### Urogenital-hypogastric fascia (UGHF)



#### Sacro-recto-genito-pubic ligament SPL



#### the superfcial layer of the vesicouterine ligament (sIVUL) between the vagina and bladder



• Schematic illustration of the four pelvic retroperitoneal compartments. The parietal compartment is indicated in blue; the vascular compartment in red; the neural compartment in yellow; the visceral compartment in green. Ela external iliac artery, Ila internal iliac artery, MRa middle rectal artery, OUA obliterated umbilical artery, SPL sacropubic ligament, SUL sacrouterine ligament, SVa superior vesical artery, Ua Uterine artery, UGHF urogenitalhypogastric fascia, Ur ureter, UVF umbilicovesical fascia, VUL vesicouterine ligamen





#### The visceral compartment

- The *visceral* compartment is so-called as it contains the pelvic organs (Fig. <u>5</u>). It is linear in shape and lies in the center of the pelvis between the two sacropubic ligaments (Fig. <u>6</u>).
- The visceral compartment includes the rectovaginal and vesicovaginal spaces from classical surgical anatomy

#### The neural compartment

 The neural compartment, so-called because of the presence of the organ-specific afferent and efferent vegetative bundles, extends from the UGHF to the bladder and is crescent shaped with its concavity tilted towards the rectum (Fig. 6). It is bordered:
 Dorsolaterally, by the UGHF extending along the mesoureter;

Medially, by the portion of the SPL stretched between the rectum and the bladder Ventrally, by the bladder

 In a dorso-ventral direction, the neural compartment includes the following spaces from classical surgical anatomy:

Heald's retrorectal space (Heald 1988), Okabayashi's pararectal space (Okabayashi 1921) Yabuki's fourth space (Yabuki et al. 2000)



#### The vascular compartment

- The *vascular* compartment, so-called because of the presence of the internal iliac vessels and their collaterals to the organs, extends from the sacrum to the UVF is crescent shaped like the parietal compartment but with a ventral concavity.
- It is bordered: anterolaterally, by the UVF and by its dorsal extension along the OUA up to the internal iliac artery; medially, by the UGHF and its ventral development along the mesoureter; and dorsally, by the sacrum.
- In a dorso-ventral direction, the vascular compartment includes the following spaces from classical surgical anatomy:
- The presacral,
- The Latzko's pararectal
- Medial paravesical spaces (Fig. <u>6</u>).

#### The parietal compartment

- The parietal compartment is crescent shaped with a dorsal concavity and includes a single uninterrupted space, extending from the lateral portions of the sacral wings to the retropubic area.
- It is bordered:

laterally, in a dorso-ventral direction, by the piriformis and internal obturator muscles and the pubic insertion of the levator ani muscles; medially, by the internal iliac artery and the Umbilicovesical fascia UVF extending along the OUA; and dorsally, by the sacral wings



Techniques in Coloproctology (2018) 22:835–845 https://doi.org/10.1007/s10151-018-1883-1

**REVIEW ARTICLE** 

CrossMark

Systematic review of outcomes following pelvic exenteration for the treatment of primary and recurrent locally advanced rectal cancer

E. Platt<sup>1</sup> · G. Dovell<sup>1</sup> · S. Smolarek<sup>1</sup>

Review > Dis Colon Rectum. 2013 Apr;56(4):519-31. doi: 10.1097/DCR.0b013e31827a7868.

#### Pelvic exenteration for rectal cancer: a systematic review

Timothy X Yang <sup>1</sup>, David L Morris, Terence C Chua

Affiliations + expand PMID: 23478621 DOI: 10.1097/DCR.0b013e31827a7868



Articles & Issues 🗸 Videos Infographics For Authors 🖌 Journal Info 🗸

**ORIGINAL ARTICLES** 

#### What Constitutes a Clear Margin in Patients With Locally Recurrent Rectal Cancer Undergoing Pelvic Exenteration?

Koh, Cherry E. FRACS<sup>\*,†,‡,5</sup>; Brown, Kilian G. M. MBBS<sup>\*,†,‡</sup>; Steffens, Daniel PhD<sup>\*</sup>; Young, Jane PhD<sup>\*,†,5</sup>; Salkeld, Glenn PhD<sup>\*,†</sup>; Solomon, Michael J. DMedSc, FRACS<sup>\*,†,±,5</sup>

