

# Contents

1	Welcome	7		
2	Overview	9		
3	CMC Inpatient	11		
4	Pineville Inpatient	15		
5	Rounds	17		
6	Progress Notes	19		
7	Discharges	21		
8	Education	23		
9	Clinic	27		
10	Postop Care after Esophagectomy	29		
Postoperative Care 33				
11	Colectomy	33		
<b>12</b>	LAR + Ileostomy	37		
13	Abdominoperineal Resection	41		
14	Esophagectomy Postop	45		
15	Esophagectomy Ward	49		
16	Jejunostomy Feedings	55		

4		CONTENTS

17 Gastrostomy Feeds	59
Hill OR	63
18 Colorectal Cases - Hill/Squires	63
Salo OR	67
19 CV Port (IJ)	67
20 Lap Jejunostomy	71
21 Lap Gastrostomy	75
22 Esophagectomy 1 Stage	79
23 Lymph Node Bioppsy	89
Esophageal Cancer	93
24 EsoCa SCORE - JR	93
25 EsoCa Objectives - Chief	95
26 Esophageal Overview	97
27 Esophageal Surgery	99
28 Staging	101
29 Superficial EsoCa	103
30 Localized EsoCa	105
31 Locally Advanced EsoCa	107
32 Chemoradiation	111
33 Radiation	113
34 Esophagectomy	115
35 Metastatic EsoCa	119
36 Survivorship	121

Gastric Cancer	125
37 Gastric Ca SCORE	125
38 Superficial Gastric	127
39 Locally-Advanced Gastric	129
40 Hereditary Diffuse Gastric Cancer	131
41 Gastrectomy	133
42 Gastric GIST	135
Colon Cancer	139
43 ColonCa SCORE	139
44 ColonCa Genetics	141
45 Partial Colectomy SCORE	143
46 Colostomy SCORE	145
47 Total Colectomy SCORE	147
48 Stage I Colon Cancer	149
49 Stage IV Colon Cancer	151
50 Colectomy	153
51 Chemotherapy	155
52 Appendiceal	157
Rectal Cancer	161
53 RectalCa SCORE	161
54 Objectives - APR/Exent	163
55 Rectal Cancer Staging	165
56 Rectal Ca Surgery	167
57 Rectal Adjuvant Therapy	169

6	CONTENTS
58 Neoadjuvant Chemotherapy	173
59 Non-operative management of Rectal Cancer	175
60 Anal Squamous Cell Carcinoma	177
Sarcoma	181
61 Soft Tissue Sarcomas	181

# Welcome

Welcome to the GI Surgical Oncology service. This orientation manual is designed to provide an introduction to the service, its workflow, and an introduction to relevant literature. Please let us know about additional information which could be included may be helpful to others.

## Overview

### 2.1 Absences

Please notify Dr Hill, Dr Salo and Dr Squires before the beginning of the rotation if you will be away during the month. This includes vacations, meetings, interview trips, and other absences.

### 2.2 Communication

Please use Haiku for messaging service attendings rather than text. Please check our status prior to messaging nights and weekends. If we are listed as unavailable, please contact *GI Surg Onc Attending LCI CMC*.

### 2.3 Medical Records

Completing medical records in a timely fashion is critical for patient safety, billing, and compliance. Timeliness also demonstrates an understanding of how the world of surgery for which residents are being prepared functions.

## 2.4 Operative Logs

Completion of operative logs is critical for board certification of the individual resident but also has implications for the appropriate assignment of residents to surgical rotations AND impacts the ability of the residency to maintain accreditation and recruit resident candidates. Residents who find it difficult to find time to maintain operative logs may find themselves excused from the operating room to complete them. Residents are expected to complete operative logs within two weeks of the end of the rotation.

### 2.5 Case Assignment

The senior resident will be expected to make case assignments for junior residents and students. It in not necessary split the month by attending - splitting by case is acceptable as well. We also expect that both residents know all the patients rather than just for one attending. This helps with nursing questions, etc.

### 2.6 Clinic

Clinic is an important part of a surgeon's education, where decisions are made regarding diagnostic workup, patient evaluation, and treatment planning. The expectation is that all residents on the service attend clinic once per week.

### 2.7 Work Hours

If the service workload jeopardizes your ability to abide by the work hour restrictions, you must notify an attending so that arrangements can be made. The service attendings are committed to abiding by work hour restrictions. Service residents are *not* expected to cover late cases (after 6pm)

# **CMC** Inpatient

Colorectal Surgery (Davis/Kasten) and GI Surgical Oncology (Hill/Salo/Squires) will cover Pineville and CMC. For efficiency, the services at each hospital will merge for patient care. Each patient will continue to have an attending surgeon, but rounding and inpatient care will be provided by the service.

### 3.1 Admissions

Provider group: "GI Surg Onc Attending LCI CMC"

List Attending Surgeon in addition

Patient List is CMC GI Surgical Oncology

### 3.2 Rounds

Work rounds for both services (CR and SurgOnc) start at 6am in STICU or 11T. Service attendings will be updated after rounds.

### 3.3 Resident Epic teams:

- GI Surgical Oncology LCI CMC
- Colorectal CMC
- Colorectal Surgery Pineville

It is critical that you notify service attendings before the start of the month to adjust the resident call schedule. Each "shift" is 5:50am to 6pm. At 6pm the resident Epic Teams will be forwarded to the night team.

Please append a text block to the bottom of each progress note specifying the Epic Team *GI Surgical Oncology Colorectal LCI CMC* for that patient to facilitate communication from nursing.

### 3.4 Consults

Established patients and directed should be discussed with the attending surgeon.

Unassigned Colorectal: Contact Attending directly

Unassigned Surgical Oncology: "GI Surgical Oncology LCI CMC"

In general, benign colorectal consults are staffed by Dr Davis. Colorectal malignancies are staffed as below. Esophageal and GE junction staffed by Dr Salo. Adenocarcinoma of distal stomach: Drs Salo/Squires. Gastric GIST: Drs Hill/Salo/Squires. Squires/Hill alternate week

	Mon	Tues	Weds	Thu	Fri
CR Malig GI Surg Onc			MHSJSH MHSJSH		

### 3.5 Postop Clinic Appt

Postoperative patients are generally seen for a Transition of Care visit within the first week

Discharge appointments are made by sending a message in Canopy the evening prior (preferred) OR the morning of discharge before 8am to:

- LCI CMC GI, Clerical
- Marsha Sukhdeo (Salo and Squires)
- Carmen Davis (Hill)
- Mychal Lacombe (Salo and Squires)
- Rebecca Wicks (Hill)
- Brandon Galloway

Please include the following information in the Epic Message:

- Name of attending
- Ward from which the patient is being discharged
- Desired date for appt @ same time
- Need for bloodwork at first visit
- Other studies to be done after discharge
  - Upper GI
  - Chest X-ray
  - Modified Barium Swallow

If messages cannot be sent in time, clinic schedulers can be reached at: Marsha Sukhdeo (Salo and Squires): 980-442-6110 or Carmen Davis (Hill): 980-442-6183.

If schedulers are not available, clinic RNs can be reached at: (Hill) 980-442-6146 or (Salo and Squires) 980-442-6143.

For patients likely to go home over the weekend or holidays, please plan to send a canopy message before 3pm on Friday or the day prior.

The scheduler will respond with a message to the discharging resident AND to the ward CNL with the appointment time, which can be included within the discharge summary. Copies of the message will also be sent to clinical nurse leaders:

11Tower: Sharon Hood6Tower: Amy Peterson

### 3.6 Conferences

- GI Tumor Planning Conf Mon 7-8am (via Teams and LCI I 3rd floor Conf Rm)
- Resident Teaching Conf Tues 7-8am 5th floor LCI II. Please review the upcoming clinic schedule and choose a case to present.
- Bone and Soft Tissue Conf Fri 7-8am (via Teams)

### 3.7 Medicine Consults

For medicine consults, please use "CHG Service Hospitalists CMC" for all new consult requests as of 11/2023 due to merging of CHG and Staff Medicine services.

# Pineville Inpatient

### 4.1 Rounds

Starting time for rounds is variable from day to day. Maddie Georgino will help organize work and timing of rounds, etc.

### 4.2 Resident Epic teams:

• Colorectal Surgery Pineville

Residents will be assigned to Epic teams by schedule. It is critical that you notify service attendings before the start of the month to adjust the resident Epic schedule. Each "shift" is 5:50am to 6pm. At 6pm the resident Epic Teams will be forwarded to the night team.

Please append a text block to the bottom of each progress note specifying the Epic Team for that patient to facilitate communication from nursing.

### 4.3 Consults

Established patients and directed should be discussed with the attending surgeon.

## 4.4 Postop Clinic Appts

Postoperative patients are generally seen for a Transition of Care visit at about two weeks.

Discharge appointments are made by sending a message in Canopy the evening prior (preferred) OR the morning of discharge before 8am to:

- Hale Mock
- Kamisha Wilson
- Madeline Georgino

Please include the following information in the Canopy Message:

- Name of attending
- Ward from which the patient is being discharged
- Desired date for appointment
- Need for Wound Ostomy RN appointment at same time (essential for new stomas)
- Need for bloodwork at first visit
- Other studies to be done after discharge
  - Upper GI
  - Chest X-ray
  - Modified Barium Swallow

For patients likely to go home over the weekend or holidays, please plan to send a canopy message before 3pm on Friday or the day prior.

### 4.5 Conferences

- GI Tumor Planning Conf Monday 7-8am (Teams)
- Resident Teaching Conf 7-8am in Conference Room. Please review the upcoming clinic schedule and choose a case to present.
- Bone and Soft Tissue Conference Friday 7-8am (Teams)

## Rounds

The following format will help speed communication of data on rounds.

ID: One line description: "Mr Glenn: PostOp day 3 after low anterior resection"

24 hour events: Summary of important events in prior 24 hrs

### Data Communication (organized by system)

Neuro: Pain control, level of alertness, psychotropic meds, sedatives, and pain meds.

CardioVascular: Vital signs (normal OR cite the range of systolic blood pressures and range of heart rate). Heart rhythm. Cardiac meds. Most recent recommendations of cardiology consult.

Respiratory: Pulmonary exam, oxygen saturation, supplied oxygen, ventilator setting. Results of CXR.

GI: Diet, bowel function, NG output. Drain outputs can often be summarized unless they are unusually high or low (and ready to be removed. New finding of bile in any abdominal drain needs special emphasis. GI meds (eg protonix, Entereg). Tube feed formula, rate and duration (continuous or nocturnal). Status of C Diff tests. Results of JP drain amylase levels (gastroesophageal patients). Results of JP triglycerides or creatinine, if sent,

Renal: Urine output in 24 hours AND in most recent 8 hour shift. Presence (or absence) of Foley catheter and plans for removal, if present. Most recent creatinine. If diuretics administered, dosage and amount of urine output during the shift when it was administered. Most recent potassium in any patient receiving (or about to receive) furosemide (Lasix). Results of Mg and Phos if abnormal.

Heme: Hemoglobin, platelets, DVT prophylaxis. PLEASE CHECK THE MAR SUMMARY DAILY to be certain that the ordered DVT prophylaxis has been given.

ID: WBC, Tmax in past 24 hours, culture results.

Endo: Diabetic regimen, blood sugar range, and amount of sliding scale insulin administered in the prior 24 hours.

### Assessment/Plan (organized by Problem List):

Each of the patients problems are addressed with an assessment and plan. Preexisting medical problems and postoperative complications need to be addressed in the plan. An assessment and plan for each organ system is usually not necessary, except for the most complex patients. Patients active medical problems should be documented on the Patient List for rounds. This helps to remind the team about medical problems which the team is managing:

- Chronic anticoagulation
- Diabetes
- Malnutrition

This problem list-oriented approach will also be helpful to writing problemoriented notes.

# Progress Notes

Progress notes need to reflect the problems which are being managed by the team. The current medical problems being managed by the team should be added to the problem list for that hospitalization. This will make it easier to generate notes which are oriented to the patient's problem list.

In addition, the progress notes provide a narrative which can later be used to generate the discharge summary (particularly for complex patients). Each day, the event of the hospitalization is carried from note to note. Each day, an additional line is added to the progress note which summarizes events for that day. This makes it possible to see within each Progress Note the pertinent events for the hospitalization. These events would include extubation, re-intubation, positive cultures, dates lines are inserted or removed, dates of removal of NG tubes and drains, and transfer to ward or re-admission to ICU. This chronology assists in treatment decisions ("how old is the IJ line" or "when did we start antibiotics?" or "when is the planned antibiotic stop date"?) but also makes the discharge summary much easier to prepare.

# Notes should be forwarded to the patient's attending (unless out of town)

#### Esophagectomy Events to be Documented:

- Extubation date/time
- NG Removal date
- Chest tube removal date
- MBS date(s) and results (aspiration | penetration)
- ICU DC orders written
- ICU discharge (transfer to ward)

### Esophagectomy Complications to be Documented:

N	Delirium
	Stroke
CV	New arrhythmia req Rx
	MI
$\mathbf{R}$	Pneumonia (3 of fever   WBC   infiltrate   abx   sputum cx)
	Effusion req drainage
	Reintubation
	Atelectasis req bronchoscopy
	ARDS
	PE
	Ventilation >48 hours after leaving OR
GI	Anastomotic leak (medical rx   stent   surgery)
	Delayed gastric emptying req botox or NG >7d
	C Diff
$\operatorname{GU}$	Urinary Retention
	Discharge with foley catheter
Η	DVT req treatment
	Return to OR
	Return to ICU

#### Communication

Please add an addendum at the BOTTOM of each progress note which includes a means for contacting the team:

Please message "GI Surgical Oncology LCI CMC" via Haiku 24/7. Messages are automatically forwarded to the General Surgery Resident on Call evenings and weekends."

#Signout

### **Evening Signout**

The Handoff Tool should be completed for all inpatients, and the responsible attending designated. This tool is critical for the safe care of patients by the nigh team. If there are studies which are pending at the time of signout (CT scan, follow-up Hb), it is critical that a plan be in place for whom to notify with an abnormal or critical study. In general, Drs Hill and Salo are always available until 10pm. Attending notification plans (service attending vs covering attending) for unstable patients should be negotiated before nightfall.

#### Weekend Signout

The senior resident is responsible for making certain that the weekend rounding resident is familiar with the patients, their problems, and the plan of care. A signout email should be prepared Friday afternoon and forwarded to the service attendings by 6pm for their review. This signout can then be edited with the attendings' notes and forwarded to the weekend rounding attending.

# Discharges

### Discharge Prescriptions

Prescriptions should be ideally be prepared the day prior to anticipated discharge and sent to the patient's pharmacy. According to North Carolina STOP guidelines, opioid prescriptions for postoperative patients should be for no more than a 7 day supply. At the same time, patients taking narcotics in the hospital should have the same dosages for their outpatient prescription, to avoid patients running out of narcotics between the time of discharge and their first clinic visit.

#### **Additional Appointments**

If followup appointments in additional to surgical followup are needed, these should be designated on the discharge orders. Particularly:

- Primary Care Physician
- Cardiologist (if new cardiac medicines)
- Co-surgeons (Urology, Thoracic Surgery, GYN)

### Discharge Summary

The discharge summary documents important events and complications in the postoperative course and serves to inform the referring physician and primary physician about these events, but also serves as a blueprint for post-discharge treatment planning. Please recognize that the first post-operative visit may be with a resident who may be meeting the patient for the first time. Key items to include:

- N Followup plan for chronic pain management Stroke
- CV Postop Arrhythmia? | MI? | CHF?

If new cardiac meds: Who is managing medications If afib: CHADS score and anticoagulation plan

R	Pneumonia?   ARDS?   TRACH?
	Need for home oxygen?
	CXR needed at first postop visit?
GI	Delayed gastric emptying?   leak?   ileus?
	Tube feed regimen
	Diet at discharge (Low residue   Full liquds   Meds with thickened
	water  NPO)
	New stoma (ileostomy   colostomy)
	Wound care needs (VAc   Prevena)
GU	Urinary Retention
	Discharge with foley catheter
Η	Complications: DVT   PE
	Anticoagulation Plan
Endo	Insulin regimen at DC (dose will be in med rec)
ID	Antibiotics at DC
	Return to ICU

### Communication

It is essential that discharge summaries be sent to the patient's primary MD and referring physician. Please review the initial consultation note for the names of providers involved in a patient's care.

# Education

The service will host Third- and Fourth-year medical students from Wake Forest University as well as externs.

Student notes should be forwarded to the patient's attending for attestation and signature.

Third year students will have a 'green card' of diagnoses and procedures which need to be checked off (and signed) during the rotation. Students: Please remind the chief resident and attendings about items which remain to be completed.

## 8.1 Medical Student Duty Hours

### Hours:

- Students will not work longer hours than residents on the same service
- Students will not work more than 80 hours/week averaged over 4 weeks.

#### Breaks: Students will

- have 4 24-hour periods free from assigned activities over a 4 week period
- not work longer than 16 continuous hoours
- have a 8 hour break from clinical/academic hours following a 16-hour shift
- can only work a maximum of 5 sequential overnight shifts

#### Exams:

 Must be excused no later than midnight prior to the day of the shift or final exam

#### Holidays:

• Must be excused from responsibilities from 5pm on the day prior to the holiday on the academic calendar through the holiday<sup>1</sup>

### 8.2 Procedures/Diseases

- Wound Care (VAC/dressing change, identify infection)
- Suture/Staple removal
- Suture Skin
- Foley catheter insertion (adult)
- Insert nasogastric tube (or OG in OR)
- Make an incision, any site
- Participate in intubation, bag mask ventilation in OR
- Xray 3-way of abdomen (interpret)
- Xray chest (interpret)
- Assist with insertion of chest tube or pigtail

### 8.3 Ask a Resident (5min discussion)

- Abdominal Pain (RUQ)
- Acute Limb Ischemia (vascular disease)
- Diverticulitis
- Neoplastic process (Breast Mass, GI Mass, soft tissue)
- Abdominal wall mass or hernia
- ABC's of trauma, Primary/Secondary survey
- Ileus, small and large bowel obstruction
- Evaluate acute surgical abdomen (participate)
- Post-op fever in surgical inpatient (discuss and participate)
- Post-op pain management (discuss and participate)

### 8.4 Medical Student Resources

Subcuticular suturing video

## 8.5 Recommended Resources:<sup>2</sup>

- Essentials of general surgery, 6th edition, [edited by] Peter F. Lawrence
- Surgery: A Case Based Clinical Review (Christian de Virgilio)
- Kaplan Surgery Notes
- NMS Surgery CaseBook
- Surgical Recall (Lorne H. Blackbourne)
- UWorld QBank

<sup>&</sup>lt;sup>1</sup>From Faculty as Teacher 2023

<sup>&</sup>lt;sup>2</sup>from Spring 2023 Surgery Syllabus

- Aquifer / Wise-MD
- OnlineMedEd

### 8.6 SHELF exam

SHELF exam topic areas

### 8.7 Entrustable Professional Activities

- 1. Gather a history and perform a physical examination
- 2. Prioritize a differential diagnosis following a clinical encounter
- 3. Recommend and interpret common diagnostic and screening tests
- 4. Enter and discuss orders and prescriptions
- 5. Document a clinical encounter in the patient record
- 6. Provide an oral presentation of a clinical encounter
- 7. Form clinical questions and retrieve evidence to advance patient care
- 8. Give or receive a patient handover to transition care responsibility
- 9. Collaborate as a member of an interprofessional team
- 10. Recognize a patient requiring urgent or emergent care and initiate evaluation and management
- 11. Obtain informed consent for tests and/or procedures
- 12. Perform general procedures of a physician
- 13. Identify system failures and contribute to a culture of safety and improvement

## Clinic

Clinic is an essential part of the educational experience as this is where preoperative evaluation and surgical planning is done.

### 9.1 Salo Clinic

Notes are generated using the .LCICONSULT SmartPhrase template. The will automatically pull in the names of the cancer care team (medical oncologist, gastroenterologist). History from prior notes can be pulled in and placed in the Subjective section. The problems pertinent to the visit are designated.

As always, notes which are copied forward need to be carefully proofread to avoid errors. For instance, when a patient returns for a postop visit, it is important not to carry forward text from the preoperative note that states 'surgery next week.

It is important that authorship be clearly communicated, particularly with regard to assessment and plan. In many cases, the assessment and plan area straightforward and can be copied forward from the prior Assessment/Plan. In some cases, clinical decision-making involves a judgement call. This is indicated in Dr Salo's notes as 'I would recommend…' In these cases, please do no copy the assessment notes verbatim to avoid confusion about whose opinions are being quoted.

# Postop Care after Esophagectomy

### 10.1 Weaning Tube Feeds in Diabetics

As outpatients begin eating more orally, their tube feeds are reduced.

Weaning from 5 cans to 4 cans: Easiest method is to maintain the same schedule (16 hours) and reduce insulin dosage by 20%. For instance, the above patient who is on 5 cans at  $75 \text{mL/hour} \times 16$  hours is receiving 16 N + 8 R at start of tube feeds and 6 hours later. This patient could be weaned by reducing rate form  $75 \text{mL/hour} \times 16$  hours to  $60 \text{mL/hour} \times 16$  hours and reducing insulin to 12 N + 6 R at the start of tube feeds AND another dose of 12 N + 6 R after 6 hours.

Weaning from 4 cans to 3 cans: One option is to decrease the duration of tube feeds from 16 hours to 12 hours, while maintaining rate of 60mL/hour. In this case, the insulin dosage could be kept the same at the start of tube feeds, BUT the dose 6 hours after the start of tube feeds could be omitted.

Once patients are on 3 cans per night, further weaning can be accomplished by skipping tube feeds (and insulin) every other night in a "tube feed holiday". This allows an evening of interrupted sleep and can tend to increase the appetite the morning after tube feeds are held.

# Postoperative Care

# Colectomy

#### Clinic

- Opiod Cessation
- Smoking Cessation
- EtOH/Drugs of Abuse Social Work
- Nutritonal Evaluation
  - All Patients Ensure for 3d preop
  - Poor nutrition ?Delay surgery
- Preop Anesthesia/ERAS Class
- Expectations of Surgery:
  - Length of Stay 1-3 days
  - Diet (self-limiting)
  - Pain control (low-opoid)
  - Activity (OOB at 6am, OOB 3x/day)
- Bowel Prep Abx and mechanical

### **Preop Holding**

- Colon PowerPlan (Hill)
- Antibiotic PowerPlan
- No PCN only for severe allergy
- Entereg (if no preop opiods)
- VTE prophylaxis
- Carbohydrate load (2hr preop)

#### $\mathbf{OR}$

- Goal-directed fluid administration
- 2L total in OR
- 3L total/first 24hrs
- Open procedures: No epidural

### Postop Day 0

- Goal-directed fluid administration
- OOB (Dangling not compliant)
- Diet
  - Low Residue diet
  - Ensure Supplements
- Teaching
- Gum, Mag & Entereg
- Pain Management
  - PCA
  - Tylenol 1gm q6 ATC
  - Gabapentin 300mg TID
  - Tramadol PRN
  - Resume all baseline pain meds
- Home Medications
  - Resume all home medications
  - No therapeutic anti-coagiation
  - Diabetes medicines
    - \* Prefer to resume all oral diabetes meds
    - \* Sliding-scale insulin ordered for all diabetics

### Postop Day 1

- Labs: K+ and CBC only
- Heparin lock IV
- Remove Foley
- d/c PCA (unless open incision)
- Out of bed > 6 hrs
- Diet: low residue
- Pain management
  - 1) d/c PCA
  - 2) Tylenol
  - 3) Gabapentin 300 tid
  - 4) Tramadol
  - 5) Home meds
  - 6) Oxy if lots of pain
- Afternoon rounds
- 1) Check patient 2-4PM
- 2) Ambulation 2x's by PM
- 3) Patient education

#### Postop Day 2

- No IV fluids unless indicated
- No labs unless indicated
- OOB >6 hrs
- No PT unless going to rehab or SNF (ask Hill first)

- Pain management
  - Same as POD#1
- Discharge planning
  - Consider early D/C (median LOS=2d)
- Otherwise plan
  - Patient
  - Nursing
- Afternoon rounds
- 1) Possible home today!
- 2) Check patient 2-3PM
- 3) Ambulation 2x's by PM
- 4) Check for dehydration
- 5) Patient education
- 6) Give estimated date of discharge

### Postop Day 3-5

- IVF
  - Consider bolus of IV fluids
  - Consider maintenance IV if we feel won't resolve soon
- Labs
  - No labs unless indicated
  - Consider ordering QOD Chem7 if prolonged ileus
  - OOB >6 hrs
  - No PT unless likely to go to rehab or SNIF (ask Hill first)
- Pain management
  - Same as POD#1
- Afternoon rounds
  - 1) Possible home today!
  - 2) Check patient 2-3PM
  - 3) Ambulation 2x's by PM
  - 4) Check for dehydration
  - 5) Patient education
  - 6) Give estimated date of discharge

# LAR + Ileostomy

#### Clinic

- Opiod Cessation
- Smoking Cessation
- EtOH/Drugs of Abuse Social Work
- Nutritional Evaluation
  - All Patients Ensure for 3d preop
  - Poor nutrition ?Delay surgery
- Arrange Wound Ostomy Nursing
- Pre-approval for Home Health
- Preop Anesthesia/ERAS Class
- Expectations of Surgery:
  - Length of Stay 1-3 days
  - Diet (self-limiting)
  - Pain control (low-opoid)
  - Activity (OOB at 6am, OOB 3x/day)
- Bowel Prep Abx and mechanical

### **Preop Holding**

- Colon PowerPlan (Hill)
- CCM order
- WOCN order
- Antibiotic PowerPlan
- No PCN only for severe allergy
- Entereg (if no preop opiods)
- VTE prophylaxis
- Carbohydrate load (2hr preop)
- Confirm stoma marking
- Confirm CCM aware of ostomy

### $\mathbf{OR}$

- Goal-directed fluid administration
  - 2L total in OR
  - 3L total/first 24hrs
- Open procedures: No epidural

### Postop Day 0

- Goal-directed fluid administration
- OOB (Dangling not compliant)
- Diet
  - Low Residue diet
  - Ensure Supplements
- Teaching
- Gum, Mag & Entereg
- Pain Management
  - PCA
  - Tylenol 1gm q6 ATC
  - Gabapentin 300mg TID
  - Oxycodone once at night
  - Tramadol PRN
  - Resume all baseline pain meds
- Home Medications
  - Resume all home medications
  - No therapeutic anti-coagiation
  - Diabetes medicines
    - \* Prefer to resume all
    - \* Sliding-scale insulin if needed
- Wound Ostomy teaching
- CCM for Home Health for stoma care

### Postop Day 1

- Labs: K+ and CBC only
- d/c "pre" plan & colon visit
- Heparin lock IV
- Remove Foley
- d/c PCA (unless laparotomy)
- Out of bed > 6 hrs
- Diet: low residue
- Pain management
  - 1) d/c PCA
  - 2) Tylenol
  - 3) Gabapentin 300 tid
  - 4) Tramadol
  - 5) Home meds
  - 6) Oxy if lots of pain

- Discharge planning
- 1) Possible home today!(25% will go home POD1)
- 2) Check patient 2-3PM
- Afternoon rounds
- 1) Possible home today!
- 2) Check patient 2-4PM
- 3) Ambulation 2x's by PM
- 4) Patient education

### Postop Day 2

- No IV fluids unless indicated
- No labs unless indicated
- OOB >6 hrs
- No PT unless going to rehab or SNF (ask Hill first)
- Pain management
  - Same as POD#1
- Wound Ostomy Teaching
- Discharge planning
  - Consider early D/C (median LOS=2d)
- Otherwise plan
  - Patient
  - Nursing
  - CCM for home health for stoma care
- Afternoon rounds
- 1) Possible home today!
- 2) Check patient 2-3PM
- 3) Ambulation 2x's by PM
- 4) Check for dehydration
- 5) Patient education
- 6) Give estimated date of discharge

Postop Day 3-5 - IVF - Consider bolus of IV fluids - Consider maintenance IV if we feel won't resolve soon - Labs - No labs unless indicated - Consider ordering QOD Chem7 if prolonged ileus - OOB >6 hrs - No PT unless likely to go to rehab or SNIF (ask Hill first) - Pain management - Same as POD#1 - Afternoon rounds 1) Possible home today! 2) Check patient 2-3PM 3) Ambulation 2x's by PM 4) Check for dehydration 5) Patient education 6) Give estimated date of discharge

# Abdominoperineal Resection

### Clinic

- Opiod Cessation
- Smoking Cessation
- EtOH/Drugs of Abuse Social Work
- Nutritional Evaluation
  - All Patients Ensure for 3d preop
  - Poor nutrition ?Delay surgery
- Arrange Wound Ostomy Nursing
- Pre-approval for Home Health
- Preop Anesthesia/ERAS Class
- Expectations of Surgery:
  - Length of Stay 1-3 days
  - Diet (self-limiting)
  - Pain control (low-opoid)
  - Activity (OOB at 6am, OOB 3x/day)
- Bowel Prep Abx and mechanical

### **Preop Holding**

- Colon PowerPlan (Hill)
- CCM order
- WOCN order
- Antibiotic PowerPlan
- No PCN only for severe allergy
- Entereg (if no preop opiods)
- VTE prophylaxis
- Carbohydrate load (2hr preop)

- Confirm stoma marking
- Confirm CCM aware of ostomy

### $\mathbf{OR}$

- Goal-directed fluid administration
  - 2L total in OR
  - 3L total/first 24hrs
- Open procedures: No epidural

### Postop Day 0

- No sitting
  - Order Sign over bed "No sitting"
- Goal-directed fluid administration
- OOB (Dangling not compliant)
- Diet
  - Low Residue diet
  - Ensure Supplements
- Teaching
- Gum, Mag & Entereg
- Pain Management
  - PCA
  - Tylenol 1gm q6 ATC
  - Gabapentin 300mg TID
  - Oxycodone once at night
  - Tramadol PRN
  - Resume all baseline pain meds
- Home Medications
  - Resume all home medications
  - No therapeutic anti-coagiation
  - Diabetes medicines
    - \* Prefer to resume all
    - \* Sliding-scale insulin if needed
- Wound Ostomy teaching
- CCM for Home Health for stoma care

### Postop Day 1

- $\bullet$  Labs: K+ and CBC only
- d/c "pre" plan & colon visit
- Heparin lock IV
- Out of bed > 6 hrs
- Diet: low residue
- Pain management
  - 1) d/c PCA
  - 2) Tylenol
  - 3) Gabapentin 300 tid
  - 4) Tramadol

- 5) Home meds
- 6) Oxy if lots of pain

### Postop Day 2

- No IV fluids unless indicated
- No labs unless indicated
- OOB >6 hrs
- No PT unless going to rehab or SNF (ask Hill first)
- Pain management
  - d/c PCA (unless laparotomy)
- Wound Ostomy Teaching
- Foley voiding challenge
  - 250mL saline into foley -> pull
  - Must void within 2 hours
- Discharge planning
  - Consider early D/C
- Otherwise plan
  - Patient
  - Nursing
  - CCM for home health for stoma care
- Afternoon rounds
- 1) Possible home today!
- 2) Check patient 2-3PM
- 3) Ambulation 2x's by PM
- 4) Check for dehydration
- 5) Patient education
- 6) Give estimated date of discharge

Postop Day 3-5 - IVF - Consider bolus of IV fluids - Consider maintenance IV if we feel won't resolve soon - Labs - No labs unless indicated - Consider ordering QOD Chem7 if prolonged ileus - OOB >6 hrs - No PT unless likely to go to rehab or SNIF (ask Hill first) - Pain management - Same as POD#1 - Afternoon rounds 1) Possible home today! 2) Check patient 2-3PM 3) Ambulation 2x's by PM 4) Check for dehydration 5) Patient education 6) Give estimated date of discharge

## Esophagectomy Postop

#### **ICU Care**

All patients admitted to STICU with Surgical Critical Care Consultation. Surgical critical care will need a phone call immediately after surgery at 6-0366. Listed: Dr Salo, CMC-GI Surgical Oncology, CMC-Surgical Critical Care. Diabetic patients (on insulin preop): Consult Kelli Dunn.

#### Ward Care

Once stable, patients are transferred to 6T. If a 6T bed is not available, please notify Dr Salo. Historically, over 95% of patients are transferred to 6T after leaving the ICU.

#### Neuro

Multimodal pain control:

- gabapentin (300mg tid liquid via Jejunostomy)
  - Tylenol (1000mg q6hrs as pediatric liquid via Jejunostomy).
  - PCA with subsequent conversion to oxycodone elixir via Jejunostomy.
  - No ketorolac (Toradol) given risk of anastomotic failure1
  - Home anxiolytics are generally administered at half the home dose

#### Cardiovascular

Postoperative atrial fibrillation is a common occurrence (20%) after esophagectomy, with the risk increasing in older patients. For patients over age 70, half of patients will develop atrial fibrillation in the postoperative period.

For prevention of atrial fibrillation, beta blockade is used. For patients receiving beta blockers prior to surgery, continuation of beta blockade is recommended. For others, patients are given metoprolol 2.5 mg IV q6hrs which can be titrated up to 10mg IV q6hrs as needed. See STS Guidelines.2

Home anti-hypertensives are usually held in order to allow beta-blockade. Patients who have elevated blood pressures once they are adequately beta-blocked (eg HR 60-70) are usually restarted on their home anti-hypertensives. Home anti-hypertensives are not routinely restarted postoperatively

### Respiratory

Chest tubes generally consist of a 28Fr Blake drain placed into the right chest. This is placed to water seal when output is less than  $200 \mathrm{mL/day}$  and there is no leak visible in the Pleurevac container. Chest tubes are usually removed once output is less than  $150 \mathrm{mL/day}$  and drainage is clear without evidence of chyle (milky appearance). A chest x-ray is obtained after removing a chest tube to look for a pneumothorax.

 ${\bf Gastrointestinal}$  All patients receive pantoprazole 40mg IV daily and meto-clopramide 5-10mg IV q 6hrs

### Nutrition

All esophagectomy patients receive a feeding jejunostomy at the time of operation. In the immediate post-operative period, patients receive Osmolite 1.5 starting the day of surgery once they are off pressors. Tube feeds are started at 20mL/hour until flatus and then advanced at 10mL per hour every 8 hours, up to a goal of 60mL per hour (x24 hours). Patients who are on tube feeds prior to surgery are generally restarted on their home tube feed formula.

Patients who do not tolerate Osmolite are switched to Vital 1.5, which is prehydrolyzed. In order to allow enough time for switching of tube feedings, patients are generally switched from Vital to their home tube feeding formula 3-4 days prior to discharge. This is typically done when they are transferred out of the ICU.

Obese patients (BMI>30) are started on Promote at 20mL/hour and increased to a goal of 60mL/hour. Promote contains a more protein relative to carbohydrdates. An alternative is Vital High Protein, which is similar to Promote but using hydrolyzed proteins.

Diarrhea in patients on tube feeds (especially nocturnal diarrhea) needs to be addressed. Despite STICU guidelines for nutrition in trauma patients, diarrhea (especially night-time diarrhea) is justification for alteration in tube feeds. See Diarrhea and Jejunstomy Feeding

Diabetic Patients

Patients who require EndoTool in the ICU will need an endocrinology consult. See also Jejunostomy Feedings with Diabetes

'Free Water'

In addition to tube feeds, most patients will receive 'free' water flushes through the jejunostomy. This is typically done as 240mL four times per day (for a total of 32oz)

### Nasogastric Tubes

A silicone nasogastric tube (Covidien Salem Sump) is placed in all patients during surgery and the position confirmed by ultrasound intraoperatively. Tubes are positioned so that all 4 dots are outside. Gastric emptying is evaluated with upper GI prior to NG tube removal. Once extubated, upper GI is typically performed on the 2nd through 4th postoperative day. Conversely, patients who are intubated will keep their nasogastric tube until extubated. Radiology ordered as "upper GI Series". In the comments section please add "IsoVue through NG tube. Contact Dr Salo for study". See Evaluation of GI Function

#### Drains

- JP1: 19Fr Blake drain in left pleura. The exit site the most lateral drain and is secured with a blue suture
- JP2: 19Fr Blake drain in right pleura. The exit site is medial to JP1 and is secured with a black suture
- JP3: 19 Fr Blake drain in abdomen. If used, the exit site is most medial
  and is secured with a blue suture
- JP4: 15Fr Blake drain in neck (for cervial incision)
- JP5: 15Fr Blake drain in subcutaneous tissue of incision

### Evaluation for Anastomotic leak

Drain amylase is an inexpensive, specific, and relatively sensitive test for anastomotic leak. Fluid from JP2 (right chest) is sent for "Body Fluid Amylase" starting on postoperative day #4 and continued until postoperative day 9. Drain amylase over 400IU/ml is considered positive and prompts a CT esophagram for confirmation. See also Evaluation for Anastomotic Leak

**Renal** Total fluids (IV + tube feeds) are generally run at 75mL/hour. Foley catheter is removed on the first or second day after surgery. Some patients will need diuresis on the 3rd or 4th postoperative day

Heme All patients require VTE prophylaxis with Lovenox or heparin SQ.

Preoperative anti-platelet agents are started on the first (aspirin) or second (Plavix) postoperative day if there is no excessive bleeding from the chest tube or JP drains. Patients on preoperative anticoagulation are transitioned to therapeutic Lovenox on the second postoperative day if no signs of bleeding.

ID Prophylactic Cefazolin and Flagyl are administered for 24 hours and stopped

# **Esophagectomy Ward**

### Ward Care

Once stable, patients are transferred to 6T. If a 6T bed is not available, please notify Dr Salo. Historically, over 95% of patients are transferred to 6T after leaving the ICU.

### 15.1 Anti-Hypertensives

Patients are transitioned to enteral metoprolol once they are transferred to the ward. Patients on IV metoprolol at 2.5mg IV q6 hours are started on 25mg enteral bid, while patients receiving 5mg IV q6 hours are started on 50mg bid. For patients who are not taking medicines by mouth, liquid metoprolol can be ordered while an inpatient. (Liquid metoprolol is not available as a home medicine.)

Home anti-hypertensives are usually held in order to allow beta-blockade. Patients who have elevated blood pressures once they are adequately beta-blocked (eg HR 60-70) are usually restarted on their home anti-hypertensives. *Home anti-hypertensives are not routinely restarted postoperatively*. Home antihypertensives are restarted on a selective basis

### 15.2 Chest tubes

Chest tubes generally consist of a 28Fr Blake drain placed into the right chest. This is placed to water seal when output is less than 200mL/day and there is no leak visible in the Pleurevac container. Chest tubes are usually removed once output is less than 150mL/day and drainage is clear without evidence of chyle (milky appearance). A chest x-ray is obtained after removing a chest tube to look for a pneumothorax.

### 15.3 GI Medicines

All patients receive pantoprazole 40mg IV daily and metoclopramide 5-10mg IV q 6hrs. This is later switched to enteric PPI and reglan. For patients <age 75, remeron is added as 15mg enteral qhs.

### 15.4 Evaluation for leak

Experience suggests that the median time to the diagnosis of leak is 7 day after surgery. Currently the risk of anastomotic leak at CMC after transthoracic esophagectomy is 2%. The most sensitive test for the diagnosis of leak is CT esophagram (see below). Based upon institutional experience, patients are divided into low risk of leak vs high risk for leak based upon drain amylase level and white blood cell count. Patients with a normal drain amylase (400IU/ml) between postoperative day 4 and day 7 AND white blood cell count less than 12 are considered low risk. Patients with either an elevated drain amylase or WBC greater than 12 are evaluated with CT esophagram. In a group of 100 patients, several were found to have elevated drain amylase in the first four days (up to 2000Iu/ml) which subsequently declined, and no evidence of leak was found

### 15.4.1 Drain Amylase

JP2 is placed into the right pleura, passes through the hiatus, and is brought out through a trocar site in the medial left upper quadrant. Drain amylase from JP2 is tested beginning on postoperative day #4 until discharge or postoperative day #9. JP2 is generally removed prior to patient discharge. JP1 is placed in the left pleura and is generally not tested for amylase.

### 15.4.2 CT esophagram

This is the most sensitive study for the detection of anastomotic leak. In order to obtain sufficient sensitivity, it requires a pre-contrast scan and the administration of contrast into the esophagus. The need for a pre-contrast scan means that the presence of remnant barium in the esophagus (from a Modified Barium Swallow) makes it more difficult to interpret the scan and should be avoided. Awake patients can drink the contrast. Patients with an NG tube present at the time of the study generally will have contrast administered through their NG tube. Almost all esophagectomy patients will have a Covidien silicone Salem Sump 18Fr tube in place. This has four marks on the tube at 45, 55, 65, and 75cm. The tube is typically positioned with the 4th mark at the nares, which means that the tip is 45cm from the nares and is usually within the gastric conduit AND below the anastomosis. The NG tube also contains side holes which extend 8.5cm above the tip of the tube. The optimal study is done with the NG tube withdrawn so that there are two side-holes above the anastomosis, which means that the tip is approximately 6cm below the anastomosis. A scout CT

will be performed, and the radiologist will determine how far back the NG tube needs to be withdrawn. This is communicated to the CT technician, who asks the nurse (or physician) to withdraw the tube the calculated amount (making a note of the starting position of the tube relative to the four marks). This should keep the tip below the level of the anastomosis, so that after the CT scan, the NG tube can be (blindly) advanced back to its original position (at approximately 45cm from the nares).

Patients who clinically deteriorate prior to post-operative day 7 in whom there is high suspicion for a leak (fevers, pleural effusion on chest X-ray, elevated JP amylase, respiratory failure) will undergo a CT esophagram earlier than post-operative day 7.

### 15.5 Anastomotic Leak Treatment

If a patient is demonstrated to have a leak, they are made NPO AND will have any pleural effusion treated with a pigtail catheter. Conservative management will generally be successful for most patients with leaks provided 1) there is no evidence of conduit necrosis 2) nutrition is optimized and 3) empyema is treated. Patients with leaks may even need a decortication, which emphasizes the importance of CT scan in the sick post-operative esophagectomy patient, so that an empyema can be diagnosed and treated. Patients with leaks who show signs of systemic illness may need to be considered for an intraluminal stent. Patients who are profoundly ill need to be evaluated for gastric necrosis with upper endoscopy.

### Nutrition

All esophagectomy patients receive a feeding jejunostomy at the time of operation. In the immediate post-operative period, patients receive Osmolite 1.5 starting the day of surgery once they are off pressors. Tube feeds are started at 20mL/hour until flatus and then advanced at 10mL per hour every 8 hours, up to a goal of 60mL per hour (x24 hours). Patients who are on tube feeds prior to surgery are generally restarted on their home tube feed formula.

Patients who do not tolerate Osmolite are switched to Vital 1.5, which is prehydrolyzed. In order to allow enough time for switching of tube feedings, patients are generally switched from Vital to their home tube feeding formula 3-4 days prior to discharge. This is typically done when they are transferred out of the ICU.

Obese patients (BMI>30) are started on Promote at 20mL/hour and increased to a goal of 60mL/hour. Promote contains a more protein relative to carbohydrdates. An alternative is Vital High Protein, which is similar to Promote but using hydrolyzed proteins.

Diarrhea in patients on tube feeds (especially nocturnal diarrhea) needs to be addressed. Despite STICU guidelines for nutrition in trauma patients, diarrhea

(especially night-time diarrhea) is justification for alteration in tube feeds. See Diarrhea and Jejunstomy Feeding

### Diabetic Patients

Patients on tube feeds are typically started on continuous (around-the-clock) tube feedings, and subsequently changed to a nocturnal regimen (typically 6pm to 10am). See Jejunostomy Feeds in Diabetic Patients

'Free Water'

In addition to tube feeds, most patients will receive 'free' water flushes through the jejunostomy. This is typically done as 240mL four times per day (for a total of 32oz)

### Nasogastric Tubes

A silicone nasogastric tube (Covidien Salem Sump) is placed in all patients during surgery and the position confirmed by ultrasound intraoperatively. Tubes are positioned so that all 4 dots are outside. Gastric emptying is evaluated with upper GI prior to NG tube removal. Once extubated, upper GI is typically performed on the 2nd through 4th postoperative day. Conversely, patients who are intubated will keep their nasogastric tube until extubated.

Upper GI is ordered as FL Upper GI Track Single Contrast

#### Drains

- JP1: 19Fr Blake drain in left pleura. The end of the tube is cut at an angle. The exit site the *usually* the most lateral drain.
- JP2: 19Fr Blake drain in right pleura. The exit site is usually medial to JP1
- JP3: 19 Fr Blake drain in abdomen. If used, the exit site is usually most medial
- JP4: 15Fr Blake drain in neck (for cervial incision)
- JP5: 15Fr Blake drain in subcutaneous tissue of incision

### Evaluation for Anastomotic leak

Drain amylase is an inexpensive, specific, and relatively sensitive test for an astomotic leak. Fluid from JP2 (right chest) is sent for "Body Fluid Amylase" starting on postoperative day #4 and continued until postoperative day 9. Drain amylase over  $400 \mathrm{IU/ml}$  is considered positive and prompts a CT esophagram for confirmation. See also Evaluation for An astomotic Leak

Renal Total fluids (IV + tube feeds) are generally run at 75mL/hour. Foley catheter is removed on the first or second day after surgery. Some patients will need diuresis on the 3rd or 4th postoperative day

**Heme** All patients require VTE prophylaxis with Lovenox or heparin SQ.

Preoperative anti-platelet agents are started on the first (aspirin) or second (Plavix) postoperative day if there is no excessive bleeding from the chest tube

or JP drains. Patients on preoperative anticoagulation are transitioned to therapeutic Lovenox on the second postoperative day if no signs of bleeding.

ID Prophylactic Cefazolin and Flagyl are administered for 24 hours and stopped

### 15.5.1 Labs

Once on the ward, BMP and CBC are checked every other day. Patients with leukocytosis (>12,000) are monitored with daily CBC until resolved.

### 15.5.2 Discharge Medicines

Patients after esophagectomy typically go home with the following medicines:

Proton pump inhibitors (will continue for 2 years) Oxycodone elixir via feeding tube. Current STOP guidelines dictate that patients receive no more than a 7 day supply of opiods at discharge Reglan 10mg po qid (will stop at 6 weeks postop) Remeron 15mg qhs (will continue for 3 months post-op) Tylenol 1000mg q6 hours as elixir (pediatric form) Gabapentin 300mg as liquid tid x 14 days Metoprolol if started postoperatively (most patients). If not on beta blockers preoperatively, this will be cut in half at the first visit and stopped at the second postoperative visit. It is important to limit the use of medicines via jejunostomy in order to lower the risk of clogging the feeding tube. While liquid metoprolol is available for inpatients, it is not available from outpatient pharmacies.

### 15.5.3 Diet at discharge

- 70% will pass their MBS for thin liquids are and discharged on protein shakes
- 15% will pass for nectar thick but fail for this liquids. They are discharged taking medicines with a sip of thickened water but otherwise NPO
- 15% will fail for nectar thick and are discharged NPO with medicines via jejunostomy

# Jejunostomy Feedings

Due to the osmotic load, jejunostomy feedings are given via enteral (Kangaroo) pump rather than bolus feeding. Feedings are generally begun as continuous (around-the-clock) and are then transitioned to nocturnal (generally 6pm to 10am) prior to discharge.

Jejunostomy feedings carry a small but significant risk ( $\sim$ 1%) of small bowel necrosis, as evidenced by the findings of pneumatosis on CT scan and in some cases small bowel necrosis and perforation. The existing literature would suggest that the early symptoms associated with small bowel necrosis are abdominal distension. As a result, patients on jejunostomy tube feeding need to be carefully monitored for distension, and tube feeds held if distension develops.(Taylor et al., 2014) Tube feeds are generally started at 30mL/hour in the immediate postoperative patients. In patients who are awake and in whom it is possible to determine whether or not there are issues of tube feed intolerance, the rate of tube feedings is increased 10mL/hour every 12 hours to a goal of 60mL/hour for women and 75mL/hour for men. In patients who are intubated/sedated, the advancement of tube feeds is individualized, and decisions are made on a daily basis on rounds.

For patients who receive tube feedings preoperatively, the same formula is generally used after surgery.

For patients with BMI less than 35, Osmolite 1.5 is used as a starting formula. If Osmolite is not tolerated due to diarrhea, Vital 1.5 can be used. For patients with BMI greater than 35, Promote is used as a starting formula, as it contains a lower amount of carbohydrates then Osmolite 1.5. Patients who are intolerant to Promote can be switched to Vital High Protein.

Patients less than 150 pounds receive 4 cans of tube feedings, administered at 60mL/hour x 16 hours (6pm to 10am). Patients greater than 150 pounds receive 5 cans of tube feedings, administered at 75mL/hour x 16 hours (6pm to 10am).

Patients with BMI >35 receive 5 cans of Promote (or Vital High Protein).

Glucerna is formulated with carbohydrates with low glycemic index, which is particularly helpful for bolus feeding (via gastrostomy) or patients who are eating. Glycemic index is less important in patients on continuous tube feeds (eg via jejunostomy). In addition, the high fiber context of Glucerna may make this formula more prone to causing clogging of jejunostomy tubes. Free Water Flushes

Once patients are transferred to the ward, free water via jejunostomy should be ordered as 240mL via jejunostomy qid. This is entered under Flush Enteral Tube for Free Water Requirements. Diarrhea with Jejunostomy Feedings

Patients who experience diarrhea during jejunostomy administration (especially diarrhea at night) will need this addressed. Several steps

- Send stool for C Diff
- Consider changing tube feedings to a more easily digestible formula. Patients on Osmolite 1.5 can be changed to Vital 1.5. Patients on Vital 1.5 can be changed to Vivonex. Patients on Promote can be changed to Vital High Protein
- Stop tube feedings for 2-4 hours to allow diarrhea to resolve
- restart tube feeds at a lower rate (eg 20mL/hour lower than the prior rate).
- Bannatrol can be given to patients taking an oral diet. Bannatrol is NOT given via jejunostomy tube to avoid clogging.
- Lomotil is generally used as a last resort.

### 16.1 Med Administration via J-Tubes

Patients who have difficulty with dysphagia or complete esophageal obstruction will need to have their medicines administered via jejunostomy tube. The process of designing a medicine regimen which can safely be administered via enteral tube can be a challenge and may require a consultation with the hospital pharmacist. One the other hand, the financial consequences of a clogged feeding tube are substantial. Flomax is never given via jejunostomy tubes due to risk of clogging.

Several common medicines are available in liquid form:

- Acetaminophen (pediatric formulation)
- Gabapentin
- Oxycodone
- Hydrocodone + acetaminophen
- Reglan

Apart from delayed-release medicines and Flomax (tamsulosin), most medicines can be crushed and resuspended and administered via gastrostomy tube if

needed.

Nexium can be administered by opening the capsule and resuspending the beads in 50mL of water. The beads are resuspended and administered. In this case, the beads don't dissolve in the syringe, but can be administered without risk of clogging.

### 16.2 Occluded Jejunostomy Tubes

Jejunostomy tubes which are refractory to the usual non-invasive means (warm water, Coca-Cola) will need to be changed over a wire in Interventional Radiology.

### 16.3 Jejunostomy feedings on preop Patients

Scope Anesthesia policy states the jejunostomy feeds are considered equivalent to oral solid food in terms of NPO interval. In general, jejunostomy tube feeds should be held at midnight the night before surgery.

### 16.4 Jejunostomy + Diabetes

### Inpt – Jejunostomy ~ Diabetes

Patients on tube feeds are typically started on continuous (around-the-clock) tube feedings, and subsequently changed to a nocturnal regimen (typically 6pm to 10am). The diabetic management for these patients differs depending upon their tube feeding regimen:

### Diabetics on Continuous Tube Feeds.

Non-insulin diabetic patients are generally initially given Osmolite 1.5 as it provides higher caloric density and does not contain fiber, which tends to clog the feeding tubes. Diabetic patients requiring insulin can be initially trialed on Osmolite 1.5 but are changed to Promote or Glucerna 1.5 if their blood sugars prove difficult to control (as evidence by either the need for EndoTool or requiring a q6 hour regimen of insulin N + insulin R).

Diabetic patients who need insulin while receiving tube feedings are typically treated initially with continuous tube feedings and around-the-clock insulin. Patients with large insulin requirements may need hourly intravenous insulin (with dosages calculated via EndoTool). Once their insulin requirements are stabilized, they are transitioned to a q6 hour regimen consisting of N and R insulin, typically twice as many units of N insulin as R (for instance, 6Units of N + 3 Units of R insulin every six hours). Patients who require EndoTool or a q6hr regimen need an endocrinology consultation with Dr Kelli Dunn to assist in diabetic management.

#### Diabetics on Nocturnal Tube Feeds

Most patients are transitioned from continuous tube feeds to nocturnal prior to discharge. Diabetic patients on nocturnal tube feedings typically receive tube feeding from 6pm to 10am and receive insulin at initiation of tube feeds (6pm) and again at 6 hours later (midnight). A typical regimen might be 18U of 70/30 at 1800 and 18U of 70/30 at MN. An alternative might be 12U NPH + 6U Regular at 1800 and 16U NPH and 8U Regular insulin at MN. Because it can take several days to determine the correct insulin regimen, diabetic patients receiving jejunostomy feedings are cycled as early in their hospital course as possible to avoid delaying discharge for blood sugar management.

Diabetic patients receiving insulin will need careful coordination of tube feeding and insulin administration when they are being transitioned from continuous to nocturnal tube feeds. Patients on continuous tube feeds may receive insulin on a q6 hour schedule, while those receiving nocturnal tube feeds receive insulin at 1800 and MN. When patients on continuous tube feeds are transitioned, the tube feeds are stopped at 10am, to be restarted at 6pm that evening. It is critical that as soon as the tube feeds are stopped at 10am, that the q6 hour insulin as stopped as well.

In either case it is critical that if tube feedings are stopped, standing insulin administration (either q6 hour OR 1800 and MN) be stopped as well. In these cases, sliding scale insulin is generally continued.

Day	Time	Tube Feeds	Insulin
SUN	MN	60mL/hr	8N+4R
Mon	6am	$60 \mathrm{mL/hr}$	8N+R
Mon	Noon	Stop	None
Mon	$6 \mathrm{pm}$	$75 \mathrm{mL/hr}$	16N + 8R
Mon	MN	$75 \mathrm{mL/hr}$	16N + 8R
Tues	10am	Stop	None
Tues	$6 \mathrm{pm}$	$75 \mathrm{mL/hr}$	16N + 8R
Tues	MN	$75 \mathrm{mL/hr}$	16N + 8R
Weds	10am	Stop	None

In this example, the patient was receiving 8N + 4R every 6 hours, so total insulin units per day is  $(8+4) \times 4 = 48$ units. Because the carbohydrate load of  $60\text{mL/hour} \times 24$  is roughly equivalent to  $75\text{mL/hour} \times 16$  hours, the total insulin administered is roughly the same. When converted to nocturnal dosing, the patient now received 16+8=24 units twice (6pm and MN) = 48U. In practice, it may be wiser to begin by adjusting the dose down a little, to perhaps 14U N and 7U R for the first night.

# Gastrostomy Feeds

Gastrostomy feeds are generally given via bolus feeds 3-4 times per day. Feeds can be given 1-2 cartons at a time.

Patients with severe reflux may require gastrostomy feeds via infusion pump, but this is much less common.

Osmolite 1.5 is generally used as an initial formula. Glucerna is formulated with carbohydrates with low glycemic index, which is particularly helpful for bolus feeding (via gastrostomy). However, the high fiber context of Glucerna may make this formula more prone to causing clogging of gastrostomy tubes.

### 17.1 Free Water Flushes

Most patients require free water in addition to tube feeds. Most patients require 240mL 4 times per day via gastrostomy

### 17.2 Med Administration via G-Tubes

Patients who have difficulty with dysphagia or complete esophageal obstruction will need to have their medicines administered via jejunostomy tube. The process of designing a medicine regimen which can safely be administered via enteral tube can be a challenge and may require a consultation with the hospital pharmacist. One the other hand, the financial consequences of a clogged feeding tube are substantial. Flomax is never given via enteral feeding tubes due to risk of clogging.

Several common medicines are available in liquid form:

- Acetaminophen (pediatric formulation)
- Gabapentin

- Oxycodone
- $\bullet \ \ Hydrocodone + acetaminophen$
- Reglan

Apart from delayed-release medicines and Flomax (tamsulosin), most medicines can be crushed and resuspended and administered via gastrostomy tube if needed.

Nexium can be administered by opening the capsule and resuspending the beads in 50mL of water. The beads are resuspended and administered. In this case, the beads don't dissolve in the syringe, but can be administered without risk of clogging.

# Hill OR

# Colorectal Cases - Hill/Squires

### Pre-op holding

ADULT SURG Colorectal ERAS MPP Hill This is what has all of the main ERAS components. Tylenol, gabapentin, Decadron, Alvimopan and heparin are all given in pre-op holding

ADULT STANDING Antimicrobial colorectal In general, I will give Ancef to patients with almost all patients with allergies to PCN. They have to remember the "severe" reaction. If it is a severe reaction, please use the second line antibiotics listed in the power plan.

Type and Screen are not typically needed for colectomy. They will have an antibody screen from office. If antibodies present, then d/w attending

### Intra-op

Positioning: I typically like to position myself.

Right sided=supine; Left sided=lithotomy; All laparoscopic colon cases will have their arms tucked with a chest tape strap.

NG/OG tubes Not needed. I will have an esthesia place if we have gastric distension. We give a multiple PO meds prior.

Review anesthesia fluid management during time out. 2L max volume, urine output not an accurate indicator.

# Salo OR

# CV Port (IJ)

### Room Prep

- Slider bed (Skytron 3600B) with head section
- C-Arm
  - Radiology technician alerted to need for C-Arm
  - Will need lead and thyroid shields for everyone in room
- Ultrasound with hockey-stick probe near patient's RIGHT SHOULDER

### Instruments - Minor instrument pan

### Disposables/Meds

- Confirm choice of port with surgeon. Usual options
  - Bard PowerPort VUE with 8Fr attachable catheter (1708062)
  - Bard PowerPort slim Implanted port (for patients with low BMI) Heparin 5mL of 1000U/ml labeled as "1000 U/ml" -Heparin 5mL of 1000U/ml + 45mL saline labeled as "100 U/ml"
  - Local
    - \* If general anesthesia: Marcaine 0.5% with epinephrine
    - \* If MAC: Xylocains 1% with epinephrine
- 1000 drape x3 AND blue paper drapes 4 packs of 2 each = 8 total
- Suture
  - 3-0 Prolene RB-1 double-arm
  - 3-0 Vicryl SH
  - 4-0 Monocryl PS2

### Position

- Supine with left arm tucked, right arm on armboard at side.
  - Right arm on armboard in case needed by anesthetist
  - NO shoulder roll
- Foley catheter: usually NOT required check with surgeon

• Lower body Bair Hugger from abdomen to feet with ONE layer of blankets on top of Bair Hugger. Velcro strap on thighs.

### Prep

Chloroprep: RIGHT chest, neck to chin and earlobe, shoulder to include deltopectoral groove.

### Drape

1000 plastic drapes outline the sterile field for the port. The skin is stretched to avoid a gap between drape and skin. Allow access to the right sternocleidomastoid, right deltopectoral groove, and sternal notch. Blue paper drapes on top of 1000 drapes Transverse drape reversed head-to-foot. Ioban around edges of port field. Skin over SCM is left without Ioban to facilitate ultrasound

**Preop evaluation** Allergies Blood thinners or anti-platelet agents History of prior central venous lines or ports History of neck surgery

### Operation

Reverse Trendelenburg

Port pocket is constructed 1cm below and parallel to clavicle 3cm long. It is essential that there is no bleeding in the pocket (to avoid a port pocket hematoma).

Trendelenburg

Right internal jugular vein is identified and its course cephalad-caudad marked on the skin.

Finder needle (22Ga) OR micropuncture kit passed into IJ. The needle should enter the vein directly beneath the ultrasound probe.

Skin anesthetized and transverse 8mm counter-incision made at needle entry site

Respiration held by anesthesia

16Ga needle passed into IJ under sono. It is essential that the vein is scanned up and down by 'rocking' the probe to visualize the tip of the needle as it passes inferior.

J Wire passed through 16Ga needle and needle withdrawn

Anesthesia resumes respirations

Ultrasound used to confirm presence of wire within the vein by scanning up and down.

Level bed

Fluoroscopy used to confirm position of wire. Dilator and sheath inserted under fluoroscopic visualization ('live'). C-arm backed away off field.

Tunneler connected to tubing on the end with small numbers. Confirm that the collar is in place and place hemostat on the end of the tubing with large numbers (to avoid allowing the collar to fall off). Tunneler bent into a curve to avoid injury to the carotid artery.

Catheter tunneled from port pocket to counter-incision over SCM. The tunneler path describes a gentle arc to avoid kinking the catheter. 1 cm of catheter near tunneler trimmed.

Catheter placed through dilator into central circulation. Most of the catheter is inserted. The catheter will generally not cause arrhythmias.

C-arm brought back onto field.

Peel-away sheath split and removed.

Traction on catheter from the port pocket is used to position catheter approximately 3cm below carina. It may be necessary to 'orbit' the C-arm if the catheter overlies the spine.

Port pocket is measured and catheter trimmed (after sliding collar superior) and attached to port. It is essential that the catheter come to rest within 1mm of the port before locking the collar in place. In order to avoid pulling on the catheter (and changing the position of the catheter tip) the port is rotated (not the catheter). Collar is locked in place.

Port accessed with straight Huber needle with  $100\mathrm{U/ml}$  heparinized saline. Blood is withdrawn into port. Needle is left in the port and the syringe detached. Syringe with concentrated flush ( $1000\mathrm{U/ml}$ ) is attached to the needle and the port flushed (without aspiration of blood). Syringe and needle are removed.

Port sutured to the underlying pectoralis fascia with 2 sutures of 3-0 Prolene, one forehand and one backhand. Sutures are tied and cut.

The port pocket irrigated and the incision closed with subcutaneous 3-0 Vicryl followed by subcuticular 4-0 Monocryl. The skin is dressed with Dermabond.

### Postop Orders

CXR in recovery to confirm central line placement

# Lap Jejunostomy

### Room Prep

- EGD cart near patient's LEFT SHOULDER (with ADULT EGD scope)
- If central venous port is placed at the same time:
- Slider bed (Skytron 3600B) with head section
- Radiology technician alerted to need for C-Arm
- $\bullet$  BK Ultrasound with hockey-stick probe near patient's RIGHT SHOULDER

### Instruments

- 5mm 30 degree scope AND 5mm 0 degree scope
- SRI laparoscopic Pan
- Salo laparoscopic instruments

### Disposables/Meds

- Veress needle (with 10mL syringe and saline)
- 5mm Z-thread optical port (3 on table, 2 more in room)
- Transverse drape AND laparoscopy drape
- Confirm choice of port with surgeon. Usual options
  - Bard PowerPort 8Fr xx8062
  - Bard PowerPort 8Fr xx8000 (low profile)
- Heparinized saline: 100U/ml (dilute) and 1000U/ml (concentrated)
- 1000 drape x3 AND blue paper drapes 4 packs of 2 each = 8 total
- Micropuncture kit available/not open (from Anesthesia)
- Jejunostomy tube: MIC 0301-14
- Silk 2-0 on RB-1 needle (on Surgical Oncology suture cart)

### Position

- Supine with left arm tucked, right arm on armboard at side.
- Foley catheter: Usually required check with surgeon

• Lower body Bair Hugger on thighs. ONE layer of blankets on top of Bair Hugger. Velcro strap on thighs. NO PILLOW UNDERNEATH LEGS.

### Prep

Chloroprep (two sticks) of abdomen (need to keep pubis in field, as well as right anterior superior iliac spine), both costal margins.

If port: RIGHT chest, neck to chin and earlobe, shoulder to include deltopectoral groove

### Drape

If central venous port: Perimeter of field draped with 1000 (clear adhesive) drapes. Four 1000 drapes around port site:

- Medial border: From Angle of Louis superiorly along midline to chin.
- Superior border: Inferior to jaw (to allow access to right internal jugular vein and SCM)
- Laterally: From inferior to ear down to right shoulder
- Inferior: From lateral shoulder medially to Angle of Louis

Abdomen: Two 1000 drapes used inferiorly keeping pubis and right anterior inferior iliac spine in field. This is critical as the far inferior/lateral RLQ needs to be in the field for optimal port placement.

Six Blue Paper Drapes around perimeter of field (on top of 1000 drapes)

If central venous port: Transverse sheet TURNED HEAD-TO-FOOT turned at an angle to keep deltopectoral groove and SCM within the field

Laparoscopy drape skewed to inferior and right to keep pubis and right ASIS in the field.

Turn on Bair Hugger only AFTER drapes in place

### Indications

Laparoscopic jejunostomy is used for enteral nutrition in patients prior to planned (or possible) esophagectomy or gastrectomy or those for whom the stomach is otherwise not available (ie after esophagectomy or gastrectomy). Patients with metastatic esophageal cancer who need enteral access are generally treated with a gastrostomy, which does not require feeding with a pump

Preop (Resident) Preop orderset: search for "Jejunostomy"

Review Clinical Information (Resident) Review staging scans (especially PET scan) to identify suspicious areas on imaging which need to be investigated at the time of laparoscopy Outpatient anticoagulation use (warfarin, Xaralto, aspirin, Plavix) Review dietitian's recommendations (how many cans of feeding per day?) If patient is scheduled for central venous port Confirm that a port has

not already been placed Prior history of central venous lines? Confirm location of port placement with surgeon (left vs right)

#### Operation

If a central venous port is placed, the port is performed first. See IJ Port

Abdominal access is obtained in one of two ways:

Infraumbilical approach using modified Hasson technique. If the peritoneum is not easily entered, a Veress needle is used to insufflate, followed by incision of the fascia with a 15 blade, and a 5mm optical port (Applied Medical Kii Fios First Entry Z-Thread Trocar) Veress needle inserted in LEFT upper quadrant just inferior to costal margin. Abdominal entry with 5mm optical Z-Thread port. 5mm port in right upper quadrant, 5mm port in RLQ just lateral to rectus, 5mm camera port between RUQ and RLQ ports

The transverse colon is now elevated (using the umbilical port, if used) and the ligament of Treitz is identified. The proximal bowel is arranged in a "C" configuration to confirm the proximal and distal ends of bowel.

A site for placement of the jejunostomy is selected on the skin, left lateral and just superior to the umbilicus. A site is selected on the bowel in the most proximal site on the jejunostomy selected which would allow for placement of the jejunostomy without tension, but at least 20cm from ligament of Treitz.

The proximal jejunum is sutured to the anterior abdominal wall with 2-0 silk. This is usually done with a 9 suture which is introduced into the abdomen with a needle driver "Korean Style" or "Paraguayan Style." Two cm distal to this suture, a diamond of sutures is placed around the proposed tube site, and one suture placed distal to avoid torsion. The final arrangement of sutures is one proximal and one distal and 4-6 sutures around the tube. All sutures were marked with hemostatic clips to facilitate replacement of the tube via fluoroscopy should the tube become dislodged.

Using Seldinger technique, a 16Fr Cook catheter introducer kit is placed within the jejunum.

A MIC 14Fr jejunostomy tube (0301-14) with the tabs trimmed with a scalpel, is inserted through the sheath and positioned in the jejunum. The tube was secured with a suture of 0 silk.

If a balloon tube is used, an 18Fr Cook dilater and sheath is used, followed by a MIC 14Fr jejunustomy tube (0200-14) and the balloon inflated with 7mL of sterile WATER.

The tube is secured with 0 silk and dressed with a BioPatch and a Tagederm dressing.

The abdomen is desufflated and the port sites closed with 4-0 Monocryl, followed by dermabond.

After dressings are applied, a Lopez valve is attached with the long Christmastree end placed into the jejunostomy tube.

Endoscopy The scope is set up:

Suction and aspiration valves inserted and working Suction tubing attached Biopsy valve attached and not leaking Cart set up for recording by powering on the Stryker SDC digital capture box A bite block is used and the scope lubricated. A neonatal scope may be necessary in patients with a tight stricture. Important findings to record:

Level in cm from the incisors, of the most proximal area of Barrett's esophagus. Level in cm from the incisors of the GE junction Appearance of the GE junction on retroflexed view. Extent of invasion of the tumor into the cardia or fundus. The scope is withdrawn and the hypopharynx suctioned. The liquid from the 'First Step' disinfectant is suctioned through the scope, followed by water.

## Lap Gastrostomy

#### Room Prep

- EGD cart near patient's LEFT SHOULDER (with NEONATAL EGD scope)
- If central venous port is placed at the same time:
  - Slider bed (Skytron 3600B) with head section
  - Radiology technician alerted to need for C-Arm
  - BK Ultrasound with hockey-stick probe near patient's RIGHT SHOULDER

#### Instruments

- 5mm 30 degree scope AND 5mm 0 degree scope
- SRI laparoscopic Pan (available)

#### Disposables/Meds

- Veress needle (with 10mL syringe and saline)
- 5mm Z-thread optical port (3 more in room)
- Transverse drape AND laparoscopy drape
- Confirm choice of port with surgeon. Usual options
  - Bard PowerPort 8Fr xx8062
  - Bard PowerPort 8Fr xx8000 (low profile)
- Heparinized saline: 100U/ml (dilute) and 1000U/ml (concentrated)
- 1000 drape x4 AND blue plastic adhesive drapes 4 packs of 2 each = 8
- Micropuncture kit available/not open (from Anesthesia)
- 20Fr Laparoscopic gastrostomy kit in room/not open
- 16Fr MIC gastrostomy tube in room/not open
- Gastrostomy 20Fr Pull PEG (in vending machine)
- GI Anchors ("T-fasteners") in room/not open

#### Endoscope Setup - Neonatal EGD scope

- Valves attached and working (suction/aspiration/biopsy cap)
- Suction tubing attached
- Connect water bottle to left-hand port
- Gauze sponges, lubricant, plastic "tray" from gauze filled with water
- Yellow (small) bite block
- "First step" sanitizer

#### Anesthesia

- ET Tube taped to left. Head turned to the left on donut
- No EKG electrodes on anterior right chest

#### Position

- Supine with left arm tucked, right arm on armboard at side.
- Foley catheter: usually NOT required check with surgeon
- Lower body Bair Hugger on thighs. ONE layer of blankets on top of Bair Hugger. Velcro strap on thighs. NO PILLOW UNDERNEATH LEGS.

#### Prep

Chloroprep (two sticks) of abdomen (need to keep pubis in field, as well as right anterior superior iliac spine), both costal margins.

If port: RIGHT chest, neck to chin and earlobe, shoulder to include deltopectoral groove

#### **Indications**

Laparoscopic gastrostomy is used in patients with esophageal obstruction . Gastrostomy feedings are much easier than jejunostomy, as they can be administered via syringe or gravity bag. By contrast, jejunostomy feedings require administration via pump. Gastrostomy is usually done as an outpatient unless there are concerns for refeeding.

#### Review Clinical Information (Resident)

Review staging scans (especially PET scan) to identify suspicious areas on imaging which need to be investigated at the time of laparoscopy Outpatient anticoagulation use (warfarin, Xaralto, aspirin, Plavix) Review dietitian's recommendations (how many cartons per day?)

#### Drape

If central venous port: Perimeter of field draped with 1000 (clear adhesive) drapes. Four 1000 drapes around port site:

- Medial border: From Angle of Louis superiorly along midline to chin.
- Superior border: Inferior to jaw (to allow access to right internal jugular vein and SCM)
- Laterally: From inferior to ear down to right shoulder

• Inferior: From lateral shoulder medially to Angle of Louis

Abdomen: Blue adhesive drapes used inferiorly keeping pubis and right anterior inferior iliac spine in field. This is critical as the far inferior/lateral RLQ needs to be in the field for optimal port placement if the patient needs a jejunostomy.

Six Blue Adhesive Drapes around perimeter of field (on top of 1000 drapes)

If central venous port: Transverse sheet TURNED HEAD-TO-FOOT turned at an angle to keep deltopectoral groove and SCM within the field

Laparoscopy drape skewed to inferior and right to keep pubis and right ASIS in the field.

Turn on Bair Hugger only AFTER drapes in place

#### Operation

If a central venous port is placed, the port is performed first. See IJ Port

**Abdominal access** is obtained in one of two ways:

- Veress needle inserted in left upper quadrant just inferior to costal margin.
   Abdominal entry with 5mm optical port at lateral border of rectus just superior to umbilicus
- Infraumbilical approach using modified Hasson technique. If the peritoneum is not easily entered, a Veress needle is used to insufflate, followed by incision of the fascia with a 15 blade, and a 5mm optical port

The insufflation pressure is decreased to 4mmHg and the abdomen vented to drop the pressure. A 30 degree scope is passed inferior to the falciform ligament into the left upper quadrant over the lateral segment of liver. The post of the scope is positioned to the left, allowing visualization of the lesser curvature of the stomach with the end of the scope near the left aspect of the falciform.

A site for placement of the gastrostomy is selected on the skin, using a 22Ga needle as a finder. The site is marked, then infiltrated with local anesthetic and a 5mm transverse incision made.

Endoscopy is performed and the following noted:

Level in cm from the incisors, of the most proximal area of Barrett's esophagus. Level in cm from the incisors of the GE junction Appearance of the GE junction on retroflexed view. Extent of invasion of the tumor into the cardia or fundus.

A bite block is positioned (unless the patient is edentulous). The endoscope (usually a neonatal scope) is introduced into the esophagus and the video capture started. If the tumor will not allow passage of the scope, do not force the scope.

Once the scope is passed into the stomach, the fundus and duodenal bulb are suctioned.

Insufflation is then reduced to 4mm and the stomach insufflated to find the optimal location for tube placement which will minimize tension and will avoid

injury to the right gastroepiploic artery. The reduced laparoscopic insufflation pressure allows endoscopic insufflation of the stomach.

Gastrostomy tube placement is done either by Pull or Seldinger technique.

#### Seldinger Gastrostomy

Four T-fasteners are then used to affix the stomach to the anterior abdominal wall. These are arranged at 8:00, 10:00, 2:00, 4:00 relative to the proposed tube site. T-fasteners are not placed inferior to the tube site to avoid injury to right gastroepiploic vessels.

The J wire is then passed into the stomach, followed by the dilators, up to 20Fr. A 16Fr MIC gastrostomy tube is introduced and the balloon inflated with 5mL of *water*.

#### **Pull Gastrostomy**

A 20Fr PULL PEG tube kit is opened. The snare and tube are passed to the upper operator. The snare is passed through the scope and opened in anticipation of the passage of the wire.

The angiocath from the kit is placed into the stomach through the abdominal wall. Once the snare has grasped the angiocath, the needle is withdrawn and the split wire passed through the Angiocath into the stomach. The snare is adjusted to grasp the split wire, which is pulled out through the mouth.

The laparoscopic port site is considered clean. The PEG tube, once it is pulled through the mouth, is considered dirty. The right abdomen is covered with a towel to protect the laparoscopic site.

The recording is now stopped. The split wire is joined to the PEG tube, which is pulled into place by the abdominal operator. The tapered portion of the tube will be the first source of resistance, which may require firm traction. The split wire (and PEG tube) is dropped of the table to the patient's left.

Once the tapered portion of the tube is through the abdominal wall skin, the next point of resistance will be the bumper of the PEG tube passing through the tumor. In general, if a 5mm neonatal scope can pass the tumor, a 20Fr PEG tube can pass as well.

The tube is pulled into position and the measurement at the skin noted. The stomach is aspirated by the upper operator and insufflation is resumed at 8mmHg. Tension on the PEG tube is adjusted to allow apposition of the gastric serosa to the abdominal wall. If the tube is not easily apposed to the abdominal wall, T fasteners ("GI Anchors") must be employed.

The scope is withdrawn and the hypopharynx suctioned. The liquid from the 'First Step' disinfectant is suctioned through the scope, followed by water.

The abdomen is desufflated and the port sites closed with 4-0 Monocryl.

## Esophagectomy 1 Stage

#### 22.1 Indications

MI Ivor Lewis ("One Stage") esophagectomy is the most common approach to esophagectomy. The patient is positioned in 'corkscrew' position to allow simultaneous access to the abdomen and chest by prepping both abdomen and chest.

### 22.2 Room Prep

Double-decker Back Table

EGD Cart (check w/ Salo regarding adult vs neonatal scope)

Doppler box with foot pedal

BK Ultrasound (with laparoscopic probe)

Walter 'Long Arm' retractor

Salo Positioning Cart:

- Four black side-rail clamps
- Four rectangular lateral positioners
- · Yassargil socket
- Well-leg holder (used as an arm holder)

Harmonic Scalpel generator box

Critical supplies (chek prior to pt in room)

- Stapler 25mm DSTXL
- Orvil 25mm
- Stapler 21mm DSTXL

- Orvil 21mm
- $\bullet\,$  Echelon 60mm stapler with 10 gold loads and 2 gray loads
- Gel Port

22.3. POSITION 81

#### Anesthesia

- Dual-lumen ETT tube (taped to left)
- Arterial line (left arm will be on arm board)
- Anesthesia may place paraspinal muscle blocks on right side.

#### 22.3 Position

Supine on blue foam pad.

Mark upper midline with skin marker

Foley catheter (with Criticor temp sensor)

Bovie pad

Lower-body Bair Hugger at level of thighs

Hair clipped from abdomen, right chest, and right axilla.

Pre-existing jejunostomy

- Prep into field
- Remove any eschar at site
- Secure to skin inferiorly with 0 silk

Shoulders are shifted to the right in preparation for 'corkscrew' positioning

Lateral positioners positioned with pad extending from greater trochanter inferiorly.

Velcro strap over thighs and over lateral positioners

Yassargil socket attached to headpiece of bed on left side.

Well-leg holder attached to Yassargil socket and used to support right forearm (which will cross body). Right arm crosses body and is supported on well-leg holder.

Lateral positioner placed posterior to spine and scapula to support right chest, allowing access for thoracoscopy.

Arm holder is dropped towards the floor enough that right arm is brought forward.

#### Prep

Chloroprep (two sticks) of abdomen, right chest, right axilla. Particular attention to prepping as far as possible to the left lateral side and the right lateral side. Nipple prepped into field.

#### Drape

Proximal right arm draped with 1000 (clear adhesive) drape.

Two blue plastic "U" drapes with center of the "U" on either lateral side with tails forming the perimeter of the field Trauma drape. All of right chest and axilla is kept within the field, as is the lateral aspect of the left upper quadrant. The field does not need to extend inferior to the umbilicus. In general, it is usually possible to keep all of these area in the field without cutting the drape, except in very large patients. Ioban strips (4") aound the periphery of the surgical field after the trauma drape. Laparoscopic cords (gas, light cord, camera) to tower at left shoulder. Suction irrigator brought off field. Laparoscopic LigaSure (2 bars) and Bovie (30/30).

#### 22.3.1 Time Out

Operation header on consent

Tumor location and likelihood of division of esophagus from abdomen (twophase operation) vs division of esophagus from chest (four-phage operation).

#### Blood:

- Surgeon's expectation of blood loss
- Availability of blood (type/screen vs type/cross).

#### Comorbidities:

Cardiopulmonary disease. If echo, ejection fraction and aortic valve area (if abnormal)

Beta blockage: Note whether patient on home beta blockade and if so, whether home medication was taken the morning of surgery.

Anticipated Intraop Problems:

- Possibility of tension left pneumothorax due to carbon dioxide entry into left chest during mediastinal dissection
- Expectation of ventilatory difficulties due to carbon dioxide entry into the right chest Gastric mobilization Greater Curvature

#### 22.4 Gastric Mobilization

7cm upper midline incision for handport. Incision is centered over pylorus. Gel-Port inserted, and abdomen insufflated to 15mmHg. Two 5mm ports placed LUQ. Medial LUQ 5mm port placed in angle between left costal margin and superior edge of GelPort ring. Lateral LUQ 5mm port placed as far lateral as possible. Depending upon visualization, a third port may be required between these two and somewhat more inferior.

If feasible, division of gastrocolic omentum starts by delivering transverse colon into GelPort and dividing ligament with cautery and LigaSure in the avascular plane just cephalad to transverse colon. Dissection proceeds as far proximal and

distal and feasible. It is important to avoid damage to the right gastroepiploic artery.

Colon is returned to abdomen and gastricolic ligament divided going using LigaSure, taking care to avoid the colon and the right gastroepiploic artery. For patients with a bulky omentum, it may be helpful to place an additional port in the left mid-quadrant for the camera to facilitate dissection of omentum off the transverse colon. Stomach is retracted to the patient's right with the back of the left hand, placing the gastrocolic ligament (and short gastric arteries) on stretch. Left gastroepippoic artery divided with LigaSure near its origin. Short gastric arteries divided with Ligasure close to spleen. As superior aspect of short gastrics is reached, the dissection plane shifts medially to create a tunnel towards the base of the left crus. This places the most superior short gastric vessels on stretch and facilitates their division. Once all short gastric vessels divided, peritoneum tethering fundus to diaphragm is incised, and fundus brushed medially.

#### 22.4.1 Distal mobilization

Attention is directed to mobilizing the lateral aspect of the duodenum. This can either be accomplished with the camera through a LUQ port and a 45 degree camera or by placing an additional 5mm port in the RLQ. Hook cautery is used to incise the connective tissue lateral and posterior to the duodenum. The gastrocolic ligament is now dissected distally, taking care to preserve the integrity of the right gastroepiploic vessels. An areolar plane generally exists between the fat pad containing the right gastroepiploic vessels and that containing the transverse mesocolon vessels.

#### 22.4.2 Left gastric artery

Lymph nodes around the celiac axis and left gastric artery are now dissected. The extend of dissection depends upon the tumor location and the presence of nodes her either based upon imaging or palpation. Dissection begins on the superior edge of the pancreas, and proceeds superiorly to the right crus. The left gastric (coronary) vein is usually located to the left of the arery and is divided with the LigaSure. In a two-phase approach, the left gastric artery is now divided with a 30mm gray load (2.0mm) Echelon linear stapler. For patients with mid-esophageal tumors, (for whom a four-phase approach is used), if there is any question about the resectability of the tumor, division of the left gastric gastric artery is generally deferred until the second abdominal phase.

#### 22.4.3 Mediastinal dissection

The esophagus is now dissected circumferentially at the gastroesophageal junction. The peritoneum overlying the diaphragm is incised, and circumferential dissection of esopahgus performed. On the left side, it is helpful to distract the left crus laterally with a Prestige clamp placed on the left crus The right

pleura is widely entered in otder to both facilitate the thoracic dissection of the esophagus and placing the conduit into the right chest in preparation for the final thoracic phase. Division of Esophagus (Two-Phase only) In a four-phase approach, the esophagus is divided from the right chest during the first (of two) thoracic phases. In a two-phase approach, the esophagus is divided from the abdomen by reaching up into the mediastinum to divide the esophagus above the tumor. This is only feasible for tumors of the gastroesophageal junction. In a two-phase approach, the esophagus is divided with a 60mm Medium-Thich (Gold) Echelon TriStaple stapler.

#### 22.4.4 Division of Esophagus (Four-Phase)

Penrose Drain (Four-Phase only) In a four-phase approach, the esophagus is divided from the chest. In order to facilitate the thoracic dissection, a 1/4 penrose drain is tied around the distal esophagus and the drain is slid cephalad into the mediastinum. The 'tails' of the drain are directed into the right chest so that they can be grasped from the right chest during the thoracic phase and can be used to provide traction on the esophagus

#### 22.4.5 Entry into right chest

The pneumoperitoneum in the abdomen is vented and the gas pressure turned down to 8mmHg in preparation for the thoracic phase. The bed is rotated to the left 20 degrees. A site for entry in to the left chest is selected just posterior to the tip of the scapula. An incision is made here and a 5mm optical port with a 5mm 0 degree scope used to enter the chest. The chest is insufflated at 8mmHg of carbon dioxide, which helps to both copllapase the lung and depress the diaphragm. Two 12mm ports are placed. The more superior is placed lateral and superior to the nipple. The inferior port is placed just lateral to the diaphragmatic reflection. It is ciritcal to avoid injury to the diaphragm (and liver) with the inferior port placement. A mini-thoracotomy incision is placed along the mid-axillary line, frequently in the same interspace as the inferior/anterior 12mm port. The chest is entered just superior to the rib and the intercostal muscles divided with the LigaSure device to allow the ribs to separate. A narrow Deaver retractor is used to guage the space between the ribs, as the width of the retractor approximates the diameter of the 25mm stapler. A 5mm 'U' port is generally placed as high as possible midway between the scapular tip port and the anterior/superior 12mm port t0o the 4 right 5

Division of Esophagus (Two-Phase) In patients with low-lying tumors, it is possible to divide the esophagus above the tumor from the abdominal approach. This evaluation is facilitated by review of the properative endoscopy (and EUS), and the PET scan, particularly the PET obtained prior to neoadjuvant chemoradiation.

After dissection of the mediastinum, a Echelon stapler with a 60mm Medium-Thick (Gold) load is inserted through a 12mm port either placed either through

the GelPort or by upsizing the most lateral LUQ

Construction of Conduit The distal esophagus and stomach is exteriorized through the GelPort. The lesser curvature vessels are divided with the LigaSure 7-9cm cephalad from the pylorus. The stomach is now placed on stretch along the greater curvature. An Echelon Medium-Thick (Gold) stapler is used to construct a 5-6cm wide gastric tube. In constructing the conduit in a patient with a tumor of the GE inction invading into the cardia, it is important to be certain that the staple line to construct the conduit stays clear of the tumor. Patients with tumors invading the cardia are at risk of a positive distal margin, meaning that microscopic tumor may be left in the wall of the gastric conduit. To make things more complicated, in patients with low-lying esophageal or GE junction tumors, not all of the length of the conduit are needed in order to reach to the level of the esophageal transection. After construction of the anastomosis, the 'extra' cephalad portion of the conduit (near the angle of His) are excised as the 'additional gastric margin.' In order to distinguish which portions of the conduit staple line which will be used to replace the esophagus and those which will be included in the 'additional gastric margin', both sides of the gastric conduit staple line are marked with sutures designated 'A', 'B', 'C', etc proceeding from the Angle of His to the antrum.

The distal esophagus and GE junction are now sent for frozen section.

Feeding jejunostomy The ligament of Trietz is identified and the jejunum is identified 20cm distal and marked with a directional suture. A site is selected to the left of the handport incision. A 16Fr Cook Introducer Kit is used to pass a 16Ga needle, followed by a J wire, through the left rectus muscle. A skin incision 4mm in length is made adjacent to the J wire. The 16Fr dilator and sheath are now passed through the rectus muscle and the dilator and wire removed. A 14Fr Jejunostomy Tube 0301-14 is selected and the 'wings' trimmed off with a scalpel. The jejunostomy tube is passed through the peelaway sheath, which is removed.

A pursestring suture 1.5cm in diameter is placed on the antimesenteric border of the jejunum. The pursestring is started and ends on the lateral aspect. A second 16Fr Cook Introduced Kit is used to introduce the J wire through the center of the pursestring.

Placement of Drains Two (or three) 19Fr full-fluted Blake drains (72230) are placed:

JP1: placed into the left pleura through the hiatus. Drain is brought out through the most lateral 5mm port site on the LUQ JP2: placed into the right pleura through the hiatus. Drain is brought out through the next most medial 5mm LUQ port site JP3: (optional) placed in the abdomen posterior to the left lateral segment of the liver and brought out through the most medial 5mm LUQ port site Transposition of Conduit The gastric conduit is now placed into the right pleura through the mediastinum, with the assistance of a laparoscopic Babcock and gentle pressure on the greater curvature with the fingers.

Entrance into the R chest (Two Phase) For two-phase operations, ports are now placed into the right chest, as stated above . In similar fashion, the inferior pulmonary ligament in dissected and the lung reflected anterior.

Anastomosis The right chest is entered and the right lung reflected anterior with the paddle placed in the superior/anterior port. The gastric conduit is placed into the mediastinum by tucking it medially from the right pleura into the posterior mediastinum, in order to allow the conduit to take the most direct path from the hiatus to the proximal esophagus. Gentle superior tension is now applied to the conduit in order to eliminate redundancy. The paddle retractor is now moved from the anterior/inferior port to the anterior/superior 12mm port and the lung reflected anterior and inferior.

OrVil The OrVil device is now used to place a EEA anvil into the distal esophagus. In general, a 25mm size is selected, unless the patient has particularly small frame, in which case a 21mm size is used. Two stay sutures of 2-0 silk on RB-1 needles are placed in the center of the esophageal staple line 3mm apart. A Harmonic scalpel is used to divide the staple line. The OrVil device is passed through the mouth and is passed through the fenestration in the staple line. The anvil portion of the OrVil is oriented so that the rounded portion is placed against the roof of the mouth. The OrVil is guided into the hypopharynx by pulling on the the tube end of the device. As the anvil approaches the hypopharynx, the jaw is pulled forward to allow passage of the anvil.

The shaft of the anvil is brought through the esophageal staple line, and the tube disconnected from the anvil by cutting the blue sutures.

The superior end of the conduit is opened along the staple line and the DST XL stapler (matching the diameter of the OrVil) introduced through the Alexis device. The stapler shaft is placed into the open end of the gastric conduit and the conduit pulled over it ('sock over shoe'). The stapler spike is brought out through the greater curvature. The anvil is grasped with a Maryland grasper placed through a superior 5mm port and the two components of the stapler are mated and the stapler tightened and fired. The knob of the stapler is rotated two turns counter-clockwise until a click is felt, at which time the anvil will flip. The stapler is withdrawn and the donuts examined and sent for pathologic exam

The anastomosis is completed by firing a Echelon Medium-Thick (Gold) linear) across the conduit cephalad to the anastomosis. The excess conduit is sent for pathologic exam as 'additional gastric margin.'

NG Tube A Covidien Salem Sump 18Fr nasogastric tube is passed by the anesthetist. A laparoscopic BK ultrasound is used to monitor the passage of the NG tube through the esophagus and into the gastric conduit. The NG tube is passed to the level that all for dots are outside the nose, with the 4th dot at the nares. The NG tube is secured with an AMT bridle.

Chest Tube A 28Fr Blake chest tube is placed through the anterior/inferior

 $12\mathrm{mm}$  port and is positioned into the posterior mediastinum. JP2 is placed near the gastric conduit. The right lung is re-inflated.

Closure The stapler access port incision is closed with 0 Vicryl to approximate the serratus muscle. The incisions are closed with 4-0 Monocryl followed by Dermabond.

## Lymph Node Bioppsy

When lymph node biopsies are obtain for a possible diagnosis of lymphoma, it is important that the pathology specimen be sent fresh and labeled for "lymphoma workup"

Workup for lymphoma requires flow cytomety, which requires special handling of of the fresh specimen. Preparaton for flow cytometry also requires that a technician be available to process the specimen. For biopsies which occur on weekends or evenings, it is critical to contact pathology before the case to be certain that specimen handling for flow cytometry can be performed.

Esophageal Cancer

## EsoCa SCORE - JR

Junior Resident SCORE - Esophageal Neoplasms

Junior Resident SCORE - Esophagectomy

#### Anatomy

#### **Epidemiology and Prevention**

- Worldwide distribution
- Risk factors

#### Presentation

- Symptoms
- Nutritional consequences

#### Diagnosis and Staging

- Physical findings
- Role of PET scan
- Role of EUS
- TNM staging system
- Treatment Categories
  - Superficial
  - Localized
  - Locally-advanced
  - Metastatic

#### Operative Treatment

- Indications for surgery
- Operative anatomy
- Gastric tube construction
- Ivor Lewis

- $\bullet$  Transhiatal
- McKeown

#### Complications

- Postoperative hypovolemia
- $\bullet$  Chylothorax
- Anastomotic leak chest
- Anastomotic leak neck
- Atrial fibrillation
- Aspiration
- Vocal cord paralysis
- $\bullet$  Stomach ischemia

#### Intraop Decision-making

- Liver metastasis
- Peritoneal lesion

#### Nonoperative management

- Palliative Radiation
- Esophageal stents

## EsoCa Objectives - Chief

#### Chief Resident SCORE

#### **Epidemiology and Prevention**

- Worldwide distribution
- Risk factors for squamous cell carcinoma
- Risk factors for adenocarcinoma

#### Presentation

- Symptoms
- Nutritional consequences

#### Diagnosis and Staging

- Physical findings
- Role of PET scan
- Role of EUS
- TNM staging system
- Treatment Categories
  - Superficial
  - Localized
  - Locally-advanced
  - Metastatic

#### Multidisciplinary Management

- High-grade dysplasia
- Endoscopic therapy
- Primary surgical therapy
- Trimodality therapy

#### Operative Mangement

- $\bullet~$  Ivor Lewis
- $\bullet$  Transhiatal
- Left thoracoabdominal
- McKeown
- Alternative conduits
- $\bullet\,$  Complications and their management

#### References

- CROSS Trial Behind the Knife Podcast
- MAGIC Trial
- FREGAT
- Dutch TIME trial

## Esophageal Overview

Esophageal cancers can be grouped into 4 treatment categories:

- Superficial  $\rightarrow$  Endoscopic therapy
- Localized  $\rightarrow$  Primary surgery
- Locally Advanced  $\rightarrow$  Trimodality therapy
- Metastatic  $\rightarrow$  Systemic therapy

Patients with minimal dysphagia, no weight loss, and small (<3cm length) tumors are evaluated with endoscopic ultrasound:

- If uT1 on EUS and <2cm in size, endoscopic mucosal resection yields more information and may be therapeutic for tumors with negative margins and without high-risk features.
- If uT2N0 on EUS, and PET scan shows a small tumor (MTV <10cm<sup>3</sup>), primary surgery is preferred in patients who are good surgical risks
- If T3 or N+ on EUS, if PET shows no metastatic disease, trimodality therapy is optimal)

Patients with dysphagia to solids or weight loss or tumor length >3cm are unlikely to have T1-2 tumors and can be evaluated with PET scan.

- If PET shows disease confined to the esophagus and regional nodes, trimodality therapy (chemoradiation followed by surgery) is optimal.
- If PET shows metastatic disease, patients are eligible for palliative chemotherapy with radiation for treatment of symptoms of dysphagia.
- If PET shows extra-regional lymph node disease, patient is at high risk for distant disease and can be treated with induction chemotherapy followed by chemoradiation and surgical evaluation.

Esophageal Surgery

## Staging

The staging workup begins once a diagnosis is made on endoscopy.

The first step is to make a preliminary determination whether the tumor is early stage (and can be treated with endoscopy or primary surgery) or later stage (and treated with chemoradiation followed by surgery or with)

The diagnostic studies needed for these treatment groups are different, so the workup can be make more efficient by sorting patients at presentation in to two groups:

Patients with minimal dysphagia, no weight loss, and tumors with less than 3cm cranio-caudal extent have a reasonable change of being T1 or T2 tumors. Tumors <3cm in length are much more likely to represent T1-2 lesions than those  $\geq$  3cm(Hollis et al., 2017)

Superficial and Localized tumors generally present with minimal dysphagia or weight loss. These tumors may present with bleeding, or dysphagia without weight loss. For these patients, determining the precise T stage is important in their workup, so **endoscopic ultrasound** is the most frequent staging study after diagnosis.

Locally-advanced or metastatic tumors tend to present with dysphagia and weight loss. At first approximation, these tumors are usually clinical T3 lesions, and the important bifurcation in their treatment is the presence or absence of metastatic disease. For patients with dysphagia and weight loss, **PET** is the most frequent initial staging study after diagnosis.

Patients who present with dysphagia are likely to have T3 or T4 disease, which is generally treated with neoadjuvant chemoradiation followed by surgery. Data from Memorial Sloan Kettering [Ripley 226] among 61 patients with esophageal cancer who presented with dysphagia, 54 (89%) were found on EUS to have uT3-4 tumors. On the other hand, among 53 patients without dysphagia, 25

(47%) were uT1-2, and were potentially candidates for primary surgery. Their conclusion was that EUS could be omitted from the workup of patients with dysphagia, but is useful in patients without dysphagia.

PET can be helpful in evaluating patients who may have T1-2 disease, and might be candidates for primary surgical therapy. A comparison of PET and EUS [malik,claxton,1] showed that uT1-2 tumors had median metabolic tumor volume (MTV) of  $6.7 \mathrm{cm}^3$ , compared with uT3-4 tumors, with a median SUV of  $35.7 \mathrm{cm}^3$ .

## Superficial EsoCa

Superficial esophageal cancer is usually asymptomatic, which means that the diagnosis is generally made in the context of surveillance for Barrett's esophagus.

Nodular Barrett's esophagus can be best evaluation with endoscopic mucosal resection, which can provide further staging information if an adenocarcinoma is found, such as depth of invasion, differentiation, and lymphovascular invasion.

Larger lesions should first be evaluated with endoscopic ultrasound (EUS)?

EUS is less sensitive for T1 lesions (Bergeron et al., 2014) -> use EMR for diagnosis (Maish and DeMeester, 2004)

(Should nodular Barrett's be evaluated with EUS prior to EMR?)

T1a tumors have a low risk of nodal metastasis (Dunbar and Spechler, 2012)

### 29.1 Endscopic Mucosal Resection (EMR)

For patients with nodular Barrett's esophagus or small tumors judged to be T1 by endoscopic ultrasound, endoscopic mucosal resection (EMR) can be diagnostic and potentially curative. (Thomas et al., 2009)

EMR also helps establish the difference between T1a and T1b compared with pathology (Worrell et al., 2018)

EMR is likely sufficient for small tumors with favorable patholgic factors(Pech et al., 2014) (Nurkin et al., 2014):

- Size less than 2cm
- Lateral and deep margins clear
- Absence of lymphovascular invasion
- Well- or moderately- differentiated

EMR: (Soetikno et al., 2005)

See MOlina JTCVS 153:1206

EMR for high-grade dysplasia (Shaheen et al., 2009)

EMR for low-grade dysplasia (Phoa et al., 2014) resulted in 25% riskd reduction in progression go HGD.

Endoscopic submucosal dissection is a technique for deeper endoscopic removal of esophageal lesions using endoscopic cautery, which dissects through the submucosa. ESD has a higher rate of curative resection (Cao et al., 2009) albiet at the cost of prolonged operative times and increased risk of complications such a bleeding. (Repici et al., 2010)

ESD takes more time and has higher R0 resection rate but similar recurrence erate at 2 eyars (Terheggen et al., 2017)

Need for RFA of Barrett's after EMR: (Haidry et al., 2013) Combination therapy with EMR and RFA results in lower rate of recurence than EMR alone.(Pech et al., 2008)

RFA for Barrett's national registry (Ganz et al., 2008)

### Localized EsoCa

#### 30.1 T1b Tumors

#### **30.2** T2N0 Tumors

Multiple studies have failed to show the additional benefit of chemotherapy or chemoradiation for pT2N0M0 esophageal cancer patients treated with radiation.

Neoadjuvant chemo not likely to be helpful for early stage disease - FFCD 9901 [Mariette 2416] enrolled patients with T1-2 or T3N0 tumors to chemoradiation followed by surgery versus surgery alone. The majority of the tumors (72%) were squamous cell carcinoma. Postoperative mortality was significally increased in the chemoradiation arm (11.1% vs 3.4%).

Meta-analysis of 5265 patients in 10 studies showed that while neoadjuvant therapy was associated with a reduction in positive margin rate, there was no difference in terms of recurrence or survival.[MOta 176]

French trial FREGAT(Markar et al., 2016)

Retrospective review of the National Cancer DataBase failed to demonstrate a difference in survival of cT2N0M0 esophageal cancer with or without preoperative chemoradiation. (Speicher et al., 2014)

A retrospective report from Johns Hopkins examined outcomes of T2N0 squamous cell carcinoma patients and showed equivalent outcomes for primary surgery vs neoadjuvant chemoradiation followed by surgery (Zhang et al., 2012)

### 30.3 Staging of T2N0 Tumors

The challenge for treatment decision-making is the limited sensitivity of endoscopic ultrasound in ruling out pT3 or pN+ disease. In other words, if a patient who is thought to have cT2N0 disease undergoes resection, and is found on pathology to have pT3 or pN $^+$  disease, this would dictate the need for post-operative chemoradiation. In general, chemoradiation after esophagectomy is difficult for patients to tolerate, with a \_\_\_\_\_ % chance of failure to complete therapy.

Data from the Cleveland Clinic looked at 53 patients judged to be T2N0 by endoscopic ultrasound (uT2N0) were treated with primary surgery. Pathologic examination showed that 17 (37%) were understaged by endoscopic ultrasound, and were pathologic (pT3) in 4 or node positive (pN $^+$ ) in 13 cases. These patients were treated with postoperative adjuvant chemoradiation.(Rice et al., 2007)

It is critical, therefore, in patients for whom primary surgery is contemplated, to attempt to identify those with occult T3 or N+ disease.

Patients who appear to have limited stage disease benefit from evaluation with a combination of

See also PMID:25047477

(MTV)

(Tumor Length)

(dysphagia)

##Primary Surgery {#primary\_surgery}

NCCN recommends PET scanS

Most common sites of metastasis are liver, lung, bones, adrenal.

PET detects occult metastasis in 10-20% of cases (Kato et al., 2002, Kim et al. (2009)). Among 129 patients with esophageal cancer, PET detected additional sites of disease in 41% and changed management in 38% (Chatterton et al., 2009)

PET for restaging detects interval development of metastatic disease in 8-17% of cases (van Vliet et al., 2008)

## Locally Advanced EsoCa

Tumors that are T2N<sup>+</sup>M0 or T3NxM0 are considered locally-advanced. The high rate of failure with surgery alone has led to development of adjunctive therapies.

#### 31.1 Trimodality Therapy

Trimodality therapy consists of chemoradiation followed by surgery.

CROSS trial randomized 364 patients with resectable esophageal and gastroesophageal junction tumors (75% adenocarcinoma) to neoadjuvant chemoradiation consisting of 4,140 cGy of radiation with concurrent carboplatin and paclitaxel or surgery alone. (van Hagen et al., 2012) Clinical node-positive disease was present in 16%. Pathologic complete response was seen in 23% of adenocarcinoma and 49% of squamous cell carcinomas. Median overall survival was 49 months after trimodality vs 24 months after surgery alone (p=0.003). Squamous cell carcinomas appeared to have particular benefit, with a hazard ratio of 0.42 for squamous cell vs 0.74 for adenocarcinoma. Median survival was improved for adenocarcinoma from 27.1 months to 43.2 months, but the median survival for squamous cell increased from 27.1months to 81.6 months for squamous cell. Rate of R0 resection was higher with chemoradiation (92% vs 69% p<0.001) and local recurrence rates lower (14% vs 34% P<0.001), and peritoneal recurrence lower (4% vs 14% P<0.001). Despite the relatively low dose of radiation, in-field recurrences were less than 5%. The primary cause of failure was distant disease (31%) and local/regional failure (14%). (Oppedijk et al., 2014)

Alternative to carbotaxol is FOLFOX (SOG trial (Leichman et al., 2011))

Ongoing PROTECT trial compares FOLFOX to paclitaxel and carboplatin (Messager et al., 2016)

See also Definitive ChemoRT

#### 31.1.1 Neoadjuvant chemoRT for SCCA

NeoCRTEC5010 (Yang et al., 2018)

Meta-abalysis of chemoRT vs chemo (Zhao et al., 2018)

#### 31.1.2 Neoadjuvant chemotheraphy followed by surgery

POET Trial (Pre-Operative therapy in Esophageal adenocarcinoma Trial) treated patients with adenocarcinoma of the gastroesophageal junction with either neoadjuvant chemotherapy (5-FU, leucovorin, cisplatin) followed by surgery or induction chemotherapy with the same agents, followed by chemoradiation (4000cGy with concurrent cisplatin and etoposide). The study failed to meet its accrual goal, but there was a suggestion of improved 3-year survival with preoperative chemoradiation (47.4% vs 27.7% p=0.07) as well as improved local control (76.5% vs 59%). In addition, chemoradiation was associated with a higher pathologic complete response rate (15.6% vs 2%)(Stahl et al., 2009). A meta-analysis of 33 randomized trials further suggested a greater benefit from neoadjuvant chemoradiation followed by surgery compared with neoadjuvant chemotherapy followed by surgery(Pasquali et al., 2017) and a similar meta-analysis (Sjoquist et al., 2011)

#Active Surveillance

EGD is poor predictor of pCR (?)

#### 31.2 GE Junction

(Siewert et al., 2006)

# 31.3 Induction chemotherapy followed by ${ m chemoRT}$

See NCCN pages M-25 and M-26

Stahl (Stahl et al., 2009) randomized patients to preoperative chemotherapy (A) vs preoperative chemotherapy followed by preoperative chemoradiation (B). Higher pcR rate in arm B (15.6% vs 2%) and ypN0 resection (64.4% vs 37.7%).

### 31.4 Postoperative chemoradiation

Intergroup-0116 (Macdonald et al., 2001) (Smalley et al., 2012) treated 556 patients with adenocarcinoma of the stomach or GE junction with surgery along vs surgery followed by postoperative chemoradiation. After a median followup of over 5 years, median overall survival iin the surgery alone group was 27 months vs 36 months in the postoperative chemoradiation group (p=0.005) Decrease in

local failure as the first site of failure in the chemoradiation group (19% versus 29%).

Chemoradiation afte resect dion of GE junction tumors (Kofoed et al., 2012) among a group of 211 patients with GE junction a denocarcinoma with positive lymph nodes with improved 3-year disease-free survival  $(37\%~{\rm s}~24\%).$ 

### Chemoradiation

\*\*This section addresses chemoradiation as primary (definitive) therapy for esophageal cancer. See also Trimodality Therapy

### 32.1 Phase II Studies

Experience with patients who refuse surgery or are medically unfit:

MD Anderson report of 61 patients out of 622 tri-modality-eligible patients who refused surgery after cCR. 5-year overall survival was 58%. 13 developped local recurrence during surveillance, and 12 had successful salvage esophagectomy (Taketa et al., 2012). The same group compared those who underwent surgery vs definitive chemoradiation in a propensity-matched fashion(Taketa et al., 2013). Irish study of 56 patients aged 70 or older treated with neoadjuvant chemoradiation followed by selective surgery. Median survival was 28 months overall, 47 months for those with cCR, 61 months for primary resection, 46 months for cCRs who did not undergo resection, and 29 months for those with salvage esophagectomy(Furlong et al., 2013).

Castoro (Castoro et al., 2013)

preSANO(Chirieac et al., 2005) Clinical Response evaluation after chemoRT for esophageal cancer with PET and EGD.

### 32.2 ChemoRT vs Trimodality therapy

The sensitivity of squamous cell carcinoma of the esophagus to chemoradiation has raised the question whether

Stahl Locally advanced squamous cell carcinoma randomized to induction chemotherapy (cisplatin, etopiside, 5FU with leuocovrin) followed by chemora-

diation (4000cGy with concurrent ciplatin and etopiside) followed by surgery compared with induction chemotherapy followed by chemoradiation (6400cGy with concurrent cisplatin and etopiside).(Stahl et al., 2005) progression-free survival was better in the trimodality group (64.3% vs 40.7%) Treatment-related morality was substantial in the surgery arm (13% vs 4%). This would be considered an excessive rate of operative mortality by modern standards. Unsurprisingly, there was no difference in overall survival between groups, in part because the surgical group had an excess 9% mortality rate from treatment. Two-year survival in the surgery arm was 40% vs 35% in the definitive chemoradiation arm.

In the French FFCD trial, 444 patients with carcinoma of the esophagus (90% squamous cell) were treated with two cycles of 5-FU and cisplatin with concurrent radiation. (Bedenne et al., 2007) Patients with a partial or complete clinical response to chemoradiation were randomized to either surgery or a boost of radiation. Patients who did not respond to chemoradiation were treated with surgery and were eliminated from the study. Only 259 of the original 444 patients (59%) went on to randomization, with the remainder (those not responding to chemoradiation) treated with surgery. Of the randomized group, median survival was 17.7months in the surgery arm versus 19.3months in the definitive chemoradiation arm. Like the Stahl study, treatment-related mortality in the surgical arm was high (9% versus 1%).

#Active Surveillance

EGD is poor predictor of pCR (?)

# Radiation

RTOG 94-05 clinical trial (Minsky et al., 2002)

## Esophagectomy

Three general approaches exist for surgical therapy.

Trans-thoracic or Ivor Lewis esophagectomy (Visbal et al., 2001) removes the intrathoracic portion of the esophagus and constructs an anastomosis within the chest. The approach include an abdominal phase, during which an esophageal substitute is constructed (usually from stomach). A thoracic phase then removes the intrathoracic esophagus and constructs an anastomosis within the chest cavity.

A McKeown esophagectomy utilizes three surgical fields: abdomen, right chest, and neck. The right chest approach allows dissection of peri-esophageal lymph nodes, and the cervical incision allows removal of the total esophagus. (McKeown, 1976) This approach is useful for tumors which involve the proximal thoracic esophagus, to ensure a negative margin. The cervical anastomosis carries a higher risk of anastomotic leak than a thoracic anastomosis, although the morbidity of a cervical anastomosis leak is less serious than that of a leak of a thoracic anastomosis.

A transhiatal esohpagectomy approaches the esophagus from the abdomen through the hiatus and from neck. By blunt dissection the esophagus is freed up without the need for thoracotomy. An esophageal substute is then brought from the abdomen to the neck through the mediastinum(Orringer and Sloan, 1978) (Orringer et al., 2007)<! – Orringer Ann Surg 2007 –> The operation is designed to avoid the pulmonary toxicity of the right chest approach. On the other hand, the blunt nature of the mediastinal dissection means that fewer lymph nodes are harvested than with a trans-thoracic approach.

Randomized trial of transthoracic esophagectomy with extended lymph node dissection versus transhiatal esohpagectomy showed fewer pulmonary complications with the transhiatal approach. (Hulscher et al., 2002) Fewer lymph nodes were havested with a transhiatal approach. A post-hoc analysis showed that

among patients with 1-8 positive lymph nodes, survival with improved with the extended lymph node dissection. (Omloo et al., 2007)

Minimally-invasive approaches to esophagectomy are now common, with evidence for less perioperative morbidity than an open approach (Biere et al., 2012) (Zhou et al., 2015)

Randomized trial of a hybrid MIE (with laparoscopy and thoracotomy) was associated with lower postoperative complications than open esophagectomy (Mariette et al., 2019)

High volume centers have lower mortality for esophageal cancer than low-volume centers. (Birkmeyer et al., 2003) (Wouters et al., 2009)

#### 34.0.1 GE Junction Adenocarcinoma

Siwert III lesions are considered gastric cancers (Rusch, 2004) (Siewert et al., 2006)

Laparoscopy may be helpful in Siewert III tumors (de Graaf et al., 2007)

#### 34.0.2 Preoperative Evaluation

Dysphagia can be scored according to Mellow et al (?):

- 0 No dysphagia
- 1 Dysphagia to normal solids
- 2 Dysphagia to soft solids (ground beef, poultry,fish)
- 3 Dysphagia to solids and liquids
- 4 Inability to swallow saliva

### 34.1 Minimally-invasive Esophagectomy

Higher lymph node yield with MIE vs open approach (?)

### 34.2 Early Recovery Pathways

**ERAS Society Guidlines** 

### 34.3 Colon Interposition

Left colon technique based upon blood supply (?)

### 34.4 Salvage esophagectomy

(Markar et al., 2014)

(Swisher et al., 2002)

### Metastatic EsoCa

### 35.1 Palliative radiation

Palliative radiation vs chemoradiation (Penniment et al., 2018)

Radiation along favored over chemoradiation in the palliaitve setting (Penniment et al., 2018)

# 35.2 Chemoradiation vs chemotherapy in Stage IV

(Guttmann et al., 2017)

### 35.3 Stents for malignant disease

(Vakil et al., 2001)

Review of guidelines 2010 Am Society GI (Sharma et al., 2010)

# Survivorship

# 36.1 Nutritional consequences of esophagectomy

### 36.1.1 Vitamin D deficiency

Vitamin D deficiency is defined as serum 25(OH)D levels below 20ng/mL

• Replacement with 2000 IU Vitamin D3 daily (Khan and Fabian, 2010)

Vitamin D insufficiency is defined as serum 25(OH)D level 20-29ng/mL

• Replacement with 1000-2000 IU Vitamin D3 daily

NIH ODS Vitamin D info

(Baker et al., 2016)

Weight loss (Martin and Lagergren, 2009) (Ouattara et al., 2012)

### 36.2 Cardiac toxicity of radiation

(Beukema et al., 2015) (Frandsen et al., 2015) (Gharzai et al., 2016)

#### 36.3 Surveillance

T1a treated with endoscopic resection EGD every 3 mo for first year, then every 6 months for second year, then annually(Shaheen et al., 2016)

**Tib treated with endoscoic resection** EGD every 3 mon for first year, then every 4-6 months for seond year, then annually CT chest/abdomen every 12 months for up to 3 years (as clinically indicated)

T1b treated with esophagectomy EGD every 3-6 months for first 2 years, then annually for 3 more years. CT every 6-9 months for first 2 years, then annually up to 5 years.

### Stage II or III treated with chemoradiation.

These patients are at risk for local recurrence (Sudo et al., 2014) and some may be candidates for salvage esophagectomy. Most relapses (95%) occur within 24 months. See also (Taketa et al., 2014)

### Locally-advanced treated with trimodality therapy

Local/regional relapses are uncommon. (Dorth et al., 2014) (Oppedijk et al., 2014) (Sudo et al., 2013) => NCCN does not recommend EGD. 90% of relapses occur within 36 months of surgery.

CT every 6 months up to 2 years (if patient is a candidate for additional curative-intent therapy)

# Gastric Cancer

## Gastric Ca SCORE

#### Gastric Cancer SCORE

#### Anatomy

- Lymph node stations
- D1 vs D2 vs D3 nodal basins

### **Epidemiology**

- Worldwide distribution
- Risk factors

### Diagnosis and Staging

- Nodal staging number vs location
- Role of laparoscopy

#### Operative Treatment

• Reconstruction options

### Non-operative Treatments

- Palliative chemotherapy
- Stents

Gastrectomy SCORE

Operative Anatomy

Preoperative Preparation

Operative Details

Complications

- Anastomotic leak
- Bleeding
- Obstruction
- Afferent Loop Syndrome
- Dumping
- Alkaline reflux gastritis

#### GIST SCORE

#### GI stromal tumors

- Pathology: Cell of origin/IHC
- Operative treatment

#### Gastric carciniods

- Pathology Cell of origin
- Type I
- Type II
- Type III
- Role for endoscopic therapy
- Surgery
  - Tumor resection
  - Antrectomy

#### Gastric lymphoma

- Low-grade vs high-grade MALT vs non-MALT
- Treatment: operative/non-op
- Indications for surgery

#### Gastric Polyps

- Indications for resection
- Non-operative treatment of hyperplastic polyps
- SMAD4 mutations

Superficial Gastric

## Locally-Advanced Gastric

Locally-advanced gastric cancer (T3 or N<sup>+</sup>) is generally treated with some form of adjuvant therapy, which has been shown to improve upon the outcomes with surgery alone.

### 39.1 Preoperative Chemotherapy

FLOT chemotherapy (Al-Batran et al., 2019) with 5FU, oxaliplatin and Taxotere superior for neoadjuvant chemotherapy compared with epirubicin, oxaliplatin and 5FU (MAGIC regimen)

MAGIC study randomized 503 patients to perioperative 'sandwich' therapy consisting of epirubicin, cisplatin, and 5-FU versus surgery alone. In the perioperative chemotherapy group, 4 cycles were administered prior to surgery, and 4 cycles afterwards. Tumors of the esophagus or gastroesophageal junction comprised 26% of the study population. While over 90% of patients assigned to the chemotherapy arm completed their preoperative chemotherapy, only 66% completed their postoperative therapy. Survival at 5 years was 36% in the perioperative chemotherapy group, compared with 24% in the surgery group (p<0.001).(Cunningham et al., 2006)

CLASSIC clinical trial randomized 1033 patients with stage II or III gastric cancer after D2 gastrectomy to 6 months of adjuvant chemotherapy versus surgery alone. Three-year survival was improved in the chemotherapy group (74% v 59%).(Bang et al., 2012)

The FFCD trial randomized patients to preoperative chemotherapy with 2 or 3 cycles of cisplatin and 5-FU versus surgery alone. Tumors of the lower esophagus or gastroesophageal junction comprised 75% of the study population. Survival at 5 years was longer in the chemotherapy group (38%) versus 24% in the surgery alone group (p=0.02).(Ychou et al., 2011)

RESONANCE-II trial of Oxaliplatin + S-1 (SOX) (Wang et al., 2021)

### 39.2 Postoperative chemotherapy

CLAASIC trial (Noh et al., 2014) (Bang et al., 2012) patients with II or IIIB gastric cancer received gastrectomy with D2 node dissection randomized to postoperative chemotherapy with capecitabine and oxaliplatin. Chemotherapy group had improved 3-year DFS (74% vs 59% P<.0001)

S-1 Adjuvant chemotherapy: (?)

### 39.3 Postoperative chemoradiation

Intergroup 0116 trial (Macdonald et al., 2001) (Smalley et al., 2012) Surgical quality control was poor, as 90% were treated a limited lymph node dissection. Long-term followup, however (Smalley et al., 2012) showed a persistent benefit of postoperative chemoradiation.

ARTIST trial 450 patients treated with a D1  $\alpha$  gastrectomy were randomized to adjuvant capcitibine and cisplatin versus chemoradiation consisting of two cycles of capcitabine/oxalipaltin followed by chemoradiation followed by chemotherapy. Overall 3- year survival did differ between groups (78.2% vs 74.2% p =0.86). A post-hoc analysis of patients with positive nodes showed a beneficial effect of chemoradiation (77.5% v 72.3% p=0.365).(Lee et al., 2012)

CRITICS trial treated all patients with preoperative chemoterhapy followed by surgery. Postoperative patients were then randomized between additional chemotherapy versus chemoradiation.

### 39.4 Preoperative chemoradiation

(Ajani et al., 2006)

# Hereditary Diffuse Gastric Cancer

HDGC is a syndrome characterized by early-onset diffuse gastric cancer and increased risk of lobular breast cancer.

Hereditary diffuse gastric cancer was first reported in a New Zealand Maori cohort. The genetic cause was subsequently found to the a deletion in the CDH1 gene.

#### 40.0.1 Penetrance estimates for lifetime risk of DGC

The original estimates of lifetime risk of gastric cancer in those with pathogenic CDH1 mutations were made from a population of patients with strong familiy histories of DGC(Hansford et al., 2015), in this group, the cumulative risk of gastric cancer by 80 years of age was estimated to be to be 70% in men and 56% in women.

The lifetime risk of gastric cancer in more modern cohorts (Roberts et al., 2019) (Xicola et al., 2019), in whom a minority of patients have a family history of gastric cancer, are much lower: 42% in men and 33% in women

# Gastrectomy

### 41.1 Proximal gastrectomy

Proximal gastrectomy for small proximal tumors can be performed with a dual-tract reconstruction to take advantage of the reservoir capacity of the distal stomach without the risk of gastroesophageal reflux.

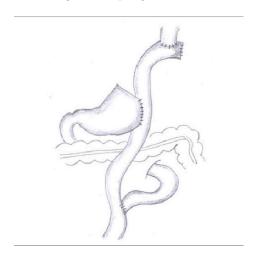


Figure 41.1: half-size image

### Gastric GIST

From NCCN Guidelines

### 42.1 Genetics

- Mutation of KIT receptor tyrosine kinase in 80%
  - KIT exon 9 mutations have lower response rate to imatinib and shorter progression-free survival than exon 11 mutations (especially at dose of 400mg daily)
- Mutation in PDGRFA receptor tyrosine kinase in 5-10%. Most mutations respond to imatinib except D842V (which may respond to avapritinib)
- No mutation of KIT or PDGFRA in 10-15%. Most will have functional inactivation of SDH complex and evidenced by lack of SDHB on IHC. These patients should be tested for germline SDH mutations. These tumors tend to occur in the stomach in younger patients.

#### Tyrosine kinase inhibitors:

- imatinib first line therapy
- Sunitinib treatment is indicated for patients with imatinib-resistant tumors or imatinib intolerance.
- Regorafenib is indicated for patients with disease progression on imatinib or sunitinib.
- Ripretinib is indicated for patients who have received prior treatment with 3 or more kinase inhibitors, including imatinib.

# Colon Cancer

### ColonCa SCORE

Colon Cancer SCORE

### Physiology

- APC
- $\bullet$  K-ras
- p53
- microsatellite instability
- CIMP

### **Epidemiology**

• Sporadic vs hereditary vs familial

#### Screening

- Colonoscopy
- Flex sig
- Occult blood testing
- Fecal DNA testing

### **Hereditary Syndromes**

- Genes | Presentation | Screening
  - Lynch Syndrome
  - APC

 $Screening\ for\ average\ risk$ 

Screening for elevated risk

### Endoscopy

• Types of polyps

### Staging

- CT scan
- $\bullet$  PET scan
- Staging

### **Adjuvant Therapy**

 $\bullet\,$  Indications for adjuvant chemo

### Operative Treatment

- Types of colectomy
- Dx large bowel obstruction

#### Resources

ASCRS Guidelines for treatment of Colorectal Cancer 2022

# ColonCa Genetics

### Resources

ASCRS Guidelines 2017

# Partial Colectomy SCORE

Partial Colectomy CORE

Indications

Operative Anatomy

### Preop Prep

- Antibiotics
- Stoma marking
- Ureteral stents

### Complications

- Anastomotic leak
- Ureteral injury

#### Resources

ASCRS Guidelines for Early Recovery

# Colostomy SCORE

 ${\bf Colostomy~SCORE}$ 

#### Indications

• Colostomy vs Ileostomy

#### Preop Prep

• Stoma marking

Operative Anatomy

Intraop Decision-making

# Total Colectomy SCORE

 ${\bf Total~Colectomy~SCORE}$ 

#### Indications

- Subtotal Colectomy
- Proctocolectomy

Operative Anatomy

Intraop Decision-making

# Stage I Colon Cancer

48.1 Malignant colon polyp

### Stage IV Colon Cancer

#### 49.1 Stent vs colostomy

CREST trial(CReST Collaborative Group, 2022) stent as bridge to surgery in obstructing left colon cancers. 246 patients randomized to colostomy vs stent as bridge to surgery. Lower rate of stoma formation in stent group.

Average patency of stent duration is 120 days

### 49.2 Colon resection in stage IV colon cancer

MSKCC: Colon surgery patients with bilobar liver metastasis treated with chemotherapy with primary tumor in place. Surgical intervention needed in 7% of cases (Poultsides and Paty, 2011)

### 49.3 Immunotherapy for dMMR (MSI high)

Colorectal cancers deficient in mismatch-repair proteins (dMMR) respond poorly to 5FU-based chemotherapy but do respond to immune-checkpoint inhibitors ( $\alpha$ -PD-1, ( $\alpha$ PD-L1, ( $\alpha$ -CTLA-4).

Review of immune checkpoint inhibitors in colorectal cancer

Blockade of the PD-1 (Programmed Death-1) ligand with monoclonal antibody as shows efficacy in colorectal cancer, especially those deficient in mismatch repair protein expression, as manifested by microsatellite instability (MSI-High) on pathologic exam.

Keynote-177 trial (André et al., 2020) randomized 307 patients with MSI-High dMMR colorectal cancer patients to receive either pembrolizumab or 5-FU based chemeotherapy and found superior survival with pembro.

### 49.4 Peritoneal Colon Cancer

Prodige - Cytoreductive surgery vs cytoreductive surgery + HPIEC

### Colectomy

#### 50.1 Extended Node dissection

Short-term outcomes of complete mesocolic excision versus D2 dissection in patients undergoing laparoscopic colectomy for right colon cancer (RELARC): a randomised, controlled, phase 3, superiority tria

Short-term outcomes of a multicentre randomized clinical trial comparing D2 versus D3 lymph node dissection for colonic cancer (COLD trial). Karachun A, Panaiotti L, Chernikovskiy I, Achkasov S, Gevorkyan Y, Savanovich N, Sharygin G, Markushin L, Sushkov O, Aleshin D, Shakhmatov D, Nazarov I, Muratov I, Maynovskaya O, Olkina A, Lankov T, Ovchinnikova T, Kharagezov D, Kaymakchi D, Milakin A, Petrov A. Br J Surg. 2020 Apr;107(5):499-508. doi: 10.1002/bjs.11387. Epub 2019 Dec 24. PMID: 31872869 Clinical Trial.

# Chemotherapy

### 51.1 Neoadjuvant Chemotherapy

Seymour MT, Morton D. FOxTROT: an international randomized controlled trial in 1052 patients (pts) evaluating neoadjuvant chemotherapy (NAC) for colon cancer. J Clin Oncol. 2019 May;37(15 Suppl):3504-3504.

## Appendiceal

#### 52.1 Mucinous Lesions

Up to Date - Appendiceal Mucinous Neoplasms

#### 52.2 Categories:

• Low grade mucinous neoplasm - Epithelial cells with dysplasia which make mucin. LMN cells are non-invasive yet have implanted within the peritoneum (similar to endometriosis).

Mucocele - Non-ruptured LAMIN

Well-differentiated mucinous adenocarcinoma. Treatment is controversial. Neoadjuvant chemotherapy vs HIPEC.

Poorly-differentiated adenocarcinoma

# **Rectal Cancer**

### RectalCa SCORE

Rectal CA SCORE

### 53.1 Anatomy

-Venous drainge

#### 53.2 Presentation

- Assessment of sphincter function
- 53.3 Operative Tx
- 53.3.1 Total mesorectal excision
- 53.3.2 Transanal excision
- 53.3.3 Isolated liver mets

### 53.4 Adjuvant therapy

- $\bullet$  Standard course chemoRT
- Swedish preop RT
- Total Neoadjuvant Therapy

# Objectives - APR/Exent

APR/Exent SCORE

#### 54.1 Indications

-Contraindications

### 54.2 Operative Anatomy

### 54.3 Preop Prep

- Genetics consultation
- Neoadjuvant therapy

### 54.4 Key Steps

### 54.5 Intraop Decisions

- Intraop radiation
- ullet Vascular reconstruction
- VRAM flap

### 54.6 Complications

- Surgical site infection
- Missed enterotomy
- Perineal wound dehiscence

- Abdominal wound dehiscence
- Urethral injury
- BeedingOstomy necrosis

## Rectal Cancer Staging

#### 55.1 MRI staging

Rectal cancer is preferentially evaluated with MRI. At Atrium, this is ordered as "MRI Pelvis without Contrast Rectal Protocol"

Key findings

- Circumferential resection margin
- Extra mesorectal vascular invasion
- Mesorectal lymph nodes
- Extra-mesorectal lymph nodes

MRI has supplanted endoscopic ultrasound (EUS) as the study of choice for staging rectal cancer.

MERCURY group trial demonstrated the predictive value of MRI for rectal cancer(Taylor et al., 2014) (MERCURY Study Group, 2006)

#### 55.2 EUS

Endoscopic ultrasound has now been replaced by MRI for initial staging, in part due to intraoperater variability

# Rectal Ca Surgery

Importance of total mesorectal excision was championed by Bill Heald (Heald and Ryall, 1986) who emphasized sharp dissection of the mesorecum outside of the visceral fasial envelope. In addition he advocated dissection of the totality of the mesorectum distal to the tumor to avoid leaving behind nodes within the mesorectum distal to the tumor(Quirke et al., 1986) (?) (?).

### Rectal Adjuvant Therapy

### 57.1 Surgery $\Rightarrow$ CRT

The original approach to adjuvant therapy for rectal cancer was surgery followed by chemoradiation.

#### 57.2 $CRT \Rightarrow Surgery$

The next innovation to adjuvant therapy for rectal cancer was chemoradiation prior to surgery, followed by adjuvant chemotherapy.

German Rectal Cancer Study (AIO-94) compared preop and postop chemoradiation in locally-advanced rectal cancer (Sauer et al., 2001). Among 823 patients, local recurrence was 6% in the preoperative group vs 13% in the postop group (p=0.0006). Overall survival was not different, but preop chemoradiation was less toxic. Pathologic complete response rate in the preoperative group as 8%.

NSABP R-03 clinical trial (Roh et al., 2009) randomized 267 patients with T3 or T4 or node-positive rectal cancer to preop vs postop chemoradiation. Disease-free survival was better in the preoperative group, wiht no difference in overall survival. Pathologic complete response rate in the preop group was 15%. The trial did not meets its accrual targets.

Dutch Rectal Cancer Trial (CKVO 95-04): Demonstrated the benefit of preop radiation in combination with TME(Kapiteijn et al., 2001). Radiation reduced local recurrence form 10.9% to 5.6%

The Dutch trial found a reduction in local recurrence form 10.9% to 5.6% with preoperative radiation with no difference in survival(Kapiteijn et al., 2001). A later update(van Gijn et al., 2011) The Medical Research Council C07 trial compared preoperative short-course radiation with selective post-operative chemora-

diation and found no difference in local recurrence (4.4% vs 10.6%) and no difference in overall survival. (Sebag-Montefiore et al., 2009)

#### 57.3 Short-course RT

An alternative to preoperative chemoradiation over 6 weeks is to administer radiation alone properative over a five day period.

Swedish Rectal Cancer Trial compared surgery alone with preoperative short-course therapy consisting of 5 doses of 500cGy of radiation without chemotherapy administered in one week prior to surgery. Local recurrence was 9% in the therapy group vs 26% in the control group, with an improvement in overall survival of 38% vs 80%(Swedish Rectal Cancer Trial et al., 1997). Of note, this trial was performed in the era prioro to the widespread use of total mesorectal excision.

Stockholm III trial showed that short-course radiation therapy performed 4-8 weeks prior to surgery resulted in imporved rates of pathologic complete response (12% vs 2%) compared with short-course radiation therapy performed the week prior to surgery (Pettersson et al., 2015)

#### 57.4 Total Neoadjuvant

A more modern approach has been to administer chemotherapy in addition to chemoradiation prior to surgery, Total Neoadjuvant Therapy (TNT)

RAPIDO trial randomized 920 patients with T4 or node-positive disease to long-course chemoradiation followed by surgery vs short-course radiation followed by chemotherapy and surgery. The pCR rate was significantly higher in the short course/chemotherapy/surgery group (28% vs 14%) and disease-specific surival at 3years was higher (30% vs 24%).(Bahadoer et al., 2021) (van der Valk et al., 2020)

PRODIGE 23 randomized 461 patients with T3 or T4 recta cancers to long-course radiation followed by surgery vs induction chemotherapy, long-course radiation followed by surgery. Up-front chemotherapy was associates with increased 3-year survival (76% vs 69%) and an increase in rate of pathologic complete response of 28% vs 12%.(Conroy et al., 2021)

STELLAR trial (Jin et al., 2022) TNT with short-course radiation (5Gy x 5 days) + CAPEOX chemotherapy vs long-course chemoRT with capecitibine. No differences in relapse-free survival or metastasis free survival. Overall survival better with TNT (p=0.03). On subgroup analysis, no group appeared to have more benefit. pCR rate 17% with TNT vs 13.9% with conventional chemoradiation. Positive margins (R1) occurred in 8.5% of TNT patients vs 12.5% in the conventional chemoRT group.

OPRA clinical trial (?). 324 rectal cancer patients staged with MRI randomized to INCT (induction chemo followed by chemoRT) vs CNCT (Chemoradiation followed by chemotherapy). Patients with a response were offered watch and wait. Patients without a response were treated with surgery. No difference in disease-free survival or overall survival or metastasis-free survival. 304 patients were restaged and only 26% were recommended to have surgery. Among 225 patients in watch and wait, somewhat more patients with INCT had recurrences (40% of 105 = 42 patients vs 27% of 102 = 32 patients). More organ preservation at 3 years with CNCT (60% CNCT vs 47% INCT). See also (?)

#### 57.5 Selective Adjuvant

Selective adjuvant therapy approaches reserves adjuvant therapy for high-risk patients, and treats low-risk patients with surgery alone.

MERCURY study group has examined selective approaches to adjuvant chemoradiation in low-risk patients with rectal cancer. (?), (?)

German OCUM group (?) (?)

Canadian [Quicksilver Trial] (https://jamanetwork.com/journals/jamaoncology/fullarticle/2730134)

#### 57.6 Neoadjuvant ImmunoTx

Mismatch repair protein deficient (MSI-high) colorectal cancer (due to either Lynch Syndrome or BRAF-1 mutation) responds pooly to chemotherapy but responds well to immunotherapy with PD-L1 blockade

MSKCC trial of neoadjuvant PD-L1 blockade with 6 months of dostarlimab for 12 patients with MMR-deficient (MSI-high) rectal cancer resulted in 100% clinical response rate (Cercek et al., 2022). No patients were subsequently treated with either chemoradiation or surgery as originally planned.

PICC trial (Hu et al., 2022) Chinese trial of PD-L1 blockade with neoadjvant toripalimab + colecoxib vs toripalimab for 3 months in 34 patients with MMR-deficient locally-advanced (T3/4 or N+) colorectal cancer. All patients were then treated with surgery. Pathologic complete response in 88% of dual-therapy patients and 65% of monotherapy. Grade 3 toxicity in 1/34 patients. Previous neoadjuvant chemo in 25%. Rectal cancer in 18%.

NICHE trial (Chalabi et al., 2024) 115 patients with dMMR locally-advanced colon cancer treated with a neoadjuvant immunotherapy. Pathologic complete response in 68%. No recurrences with a median follow up of 26 months.

# Neoadjuvant Chemotherapy

The PROSPECT clinical trial (?) randomized patients with T2/3 recctal cancer to neoadjuvant therapy with chemoradiation vs FOLFOX. 585 in the FOLFOX group and 543 in the chemoradiotherapy group. FOLFOX was noninferior to chemoradiotherapy for disease-free survival and the groups were similar with respect to overall survival. In the FOLFOX group, 53 patients (9.1%) received preoperative chemoradiotherapy (if the primary tumor decreased in size by <20% or if FOLFOX was discontinued because of side effects) and 8 (1.4%) received postoperative chemoradiotherapy.

# Non-operative management of Rectal Cancer

MSKCC published an experience of 113 patients with cCR after chemoradiation for rectal cancer who elected non-operative management (?). Among this group, 22 developed a local recurrence, and 8% developed metastatic disease.(?)

A new paradigm for rectal cancer: Organ preservation Introducing the International Watch & Wait Database (IWWD) (Beets et al., 2015)

```
Habr-Gama A.
Gama-Rodrigues J.
Sao Juliao G.P.
et al.
```

Local recurrence after complete clinical response and watch and wait in rectal cancer after neoadjuvant chemoradiation: impact of salvage therapy on local disease control. Int J Radiat Oncol Biol Phys. Mar 15 2014; 88: 822-828 View in Article

```
Maas M.
Beets-Tan R.G.
Lambregts D.M.
et al.
```

Wait-and-see policy for clinical complete responders after chemoradiation for rectal cancer. J Clin Oncol. Dec 10 2011; 29: 4633-4640 View in Article

```
Scopus (791)
PubMed
Crossref
Google Scholar
```

#### 176CHAPTER 59. NON-OPERATIVE MANAGEMENT OF RECTAL CANCER

Appelt A.L. Ploen J. Harling H. et al.

High-dose chemoradiotherapy and watchful waiting for distal rectal cancer: a prospective observational study. Lancet Oncol. Aug 2015; 16: 919-927 View in Article

Scopus (392)
PubMed
Abstract
Full Text
Full Text PDF
Google Scholar

Smith J.D. Ruby J.A. Goodman K.A. et al.

Surveillance after neoadjuvant therapy in advanced rectal cancer with complete clinical response can have comparable outcomes to total mesorectal excision. Int J Colorectal Dis. Jun 2015; 30:769-774

# Anal Squamous Cell Carcinoma

NCCN Guidelines

#### 60.1 Chemoradiation

Chemoradiation is now the standard for anal squamous cell carcinoma of the anal canal and for perianal cancers (except for small T1 lesions). Nigro protocol is the standard approach. (Nigro et al., 1983)

#### 60.1.1 Restaging after chemoRT

Based on the results of the ACT-II study, it may be appropriate to follow patients who have not achieved a complete clinical response with persistent anal cancer up to 6 months following completion of radiation therapy and chemotherapy as long as there is no evidence of progressive disease during this period of follow-up. Persistent disease may continue to regress even at 26 weeks from the start of treatment.(James et al., 2013)

# Sarcoma

## Chapter 61

# Soft Tissue Sarcomas

## 61.1 Desmoid Tumors

Review of desmoid tumors: Global Consensus Guidelines

## 61.2 Retroperitoneal

### **Preop Radiation**

STRASS trial randomized 266 patients to preop radiation followed by surgical resection vs surgery alone. At a median followup of 43 months, there was no difference in recurrence-free survival. Serious adverse affects were more common in the preop radiation group (24% vs 10%). One patients in the radiation group died of treatment-related toxicity (gastrocolic fistula), compared with none in the surgery alone group (Bonvalot et al., 2020). See commentary: (Cardona, 2020)

STRASS2: Ongoing trial of neoadjuvant chemotherapy.

#### 61.3 Peritoneal mesothelioma

Review (Bridda et al., 2007)

Surgical Oncology Clinics review (Li and Alexander, 2018)

# **Bibliography**

- Ajani, J. A., Winter, K., Okawara, G. S., Donohue, J. H., Pisters, P. W. T., Crane, C. H., Greskovich, J. F., Anne, P. R., Bradley, J. D., Willett, C., and Rich, T. A. (2006). Phase II trial of preoperative chemoradiation in patients with localized gastric adenocarcinoma (RTOG 9904): quality of combined modality therapy and pathologic response. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 24(24):3953–3958.
- Al-Batran, S.-E., Homann, N., Pauligk, C., Goetze, T. O., Meiler, J., Kasper, S., Kopp, H.-G., Mayer, F., Haag, G. M., Luley, K., Lindig, U., Schmiegel, W., Pohl, M., Stoehlmacher, J., Folprecht, G., Probst, S., Prasnikar, N., Fischbach, W., Mahlberg, R., Trojan, J., Koenigsmann, M., Martens, U. M., Thuss-Patience, P., Egger, M., Block, A., Heinemann, V., Illerhaus, G., Moehler, M., Schenk, M., Kullmann, F., Behringer, D. M., Heike, M., Pink, D., Teschendorf, C., Löhr, C., Bernhard, H., Schuch, G., Rethwisch, V., von Weikersthal, L. F., Hartmann, J. T., Kneba, M., Daum, S., Schulmann, K., Weniger, J., Belle, S., Gaiser, T., Oduncu, F. S., Güntner, M., Hozaeel, W., Reichart, A., Jäger, E., Kraus, T., Mönig, S., Bechstein, W. O., Schuler, M., Schmalenberg, H., Hofheinz, R. D., and FLOT4-AIO Investigators (2019). Perioperative chemotherapy with fluorouracil plus leucovorin, oxaliplatin, and docetaxel versus fluorouracil or capecitabine plus cisplatin and epirubicin for locally advanced, resectable gastric or gastro-oesophageal junction adenocarcinoma (FLOT4): a randomised, phase 2/3 trial. Lancet (London, England), 393(10184):1948–1957.
- André, T., Shiu, K.-K., Kim, T. W., Jensen, B. V., Jensen, L. H., Punt, C., Smith, D., Garcia-Carbonero, R., Benavides, M., Gibbs, P., de la Fouchardiere, C., Rivera, F., Elez, E., Bendell, J., Le, D. T., Yoshino, T., Van Cutsem, E., Yang, P., Farooqui, M. Z. H., Marinello, P., Diaz, L. A., and KEYNOTE-177 Investigators (2020). Pembrolizumab in Microsatellite-Instability-High Advanced Colorectal Cancer. The New England Journal of Medicine, 383(23):2207–2218.
- Bahadoer, R. R., Dijkstra, E. A., van Etten, B., Marijnen, C. A. M., Putter, H., Kranenbarg, E. M.-K., Roodvoets, A. G. H., Nagtegaal, I. D., Beets-Tan, R. G. H., Blomqvist, L. K., Fokstuen, T., Ten Tije, A. J., Capdevila,

J., Hendriks, M. P., Edhemovic, I., Cervantes, A., Nilsson, P. J., Glimelius, B., van de Velde, C. J. H., Hospers, G. A. P., and RAPIDO collaborative investigators (2021). Short-course radiotherapy followed by chemotherapy before total mesorectal excision (TME) versus preoperative chemoradiotherapy, TME, and optional adjuvant chemotherapy in locally advanced rectal cancer (RAPIDO): a randomised, open-label, phase 3 trial. *The Lancet. Oncology*, 22(1):29–42.

- Baker, M., Halliday, V., Williams, R., and Bowery, D. (2016). A systematic review of the nutritional consequences of esophagectomy. *Clinical nutrition* (*Edinburgh, Scotland*), 35(5):987–997. Publisher: Clin Nutr.
- Bang, Y.-J., Kim, Y.-W., Yang, H.-K., Chung, H. C., Park, Y.-K., Lee, K. H., Lee, K.-W., Kim, Y. H., Noh, S.-I., Cho, J. Y., Mok, Y. J., Kim, Y. H., Ji, J., Yeh, T.-S., Button, P., Sirzén, F., Noh, S. H., and CLASSIC trial investigators (2012). Adjuvant capecitabine and oxaliplatin for gastric cancer after D2 gastrectomy (CLASSIC): a phase 3 open-label, randomised controlled trial. Lancet (London, England), 379(9813):315–321.
- Bedenne, L., Michel, P., Bouché, O., Milan, C., Mariette, C., Conroy, T., Pezet, D., Roullet, B., Seitz, J.-F., Herr, J.-P., Paillot, B., Arveux, P., Bonnetain, F., and Binquet, C. (2007). Chemoradiation followed by surgery compared with chemoradiation alone in squamous cancer of the esophagus: FFCD 9102. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 25(10):1160–1168.
- Beets, G. L., Figueiredo, N. L., Habr-Gama, A., and van de Velde, C. J. H. (2015). A new paradigm for rectal cancer: Organ preservation: Introducing the International Watch & Wait Database (IWWD). European Journal of Surgical Oncology: The Journal of the European Society of Surgical Oncology and the British Association of Surgical Oncology, 41(12):1562–1564.
- Bergeron, E. J., Lin, J., Chang, A. C., Orringer, M. B., and Reddy, R. M. (2014). Endoscopic ultrasound is inadequate to determine which T1/T2 esophageal tumors are candidates for endoluminal therapies. *The Journal of Thoracic and Cardiovascular Surgery*, 147(2):765–771: Discussion 771–773.
- Beukema, J. C., van Luijk, P., Widder, J., Langendijk, J. A., and Muijs, C. T. (2015). Is cardiac toxicity a relevant issue in the radiation treatment of esophageal cancer? Radiotherapy and Oncology: Journal of the European Society for Therapeutic Radiology and Oncology, 114(1):85–90.
- Biere, S. S., van Berge Henegouwen, M. I., Maas, K. W., Bonavina, L., Rosman, C., Garcia, J. R., Gisbertz, S. S., Klinkenbijl, J. H., Hollmann, M. W., de Lange, E. S., Bonjer, H. J., van der Peet, D. L., and Cuesta, M. A. (2012). Minimally invasive versus open oesophagectomy for patients with oesophageal cancer: a multicentre, open-label, randomised controlled trial. Lancet, 379(9829):1887–92. Type: Journal Article.
- Birkmeyer, J. D., Stukel, T. A., Siewers, A. E., Goodney, P. P., Wennberg,

D. E., and Lucas, F. L. (2003). Surgeon volume and operative mortality in the United States. *The New England Journal of Medicine*, 349(22):2117–2127.

- Bonvalot, S., Gronchi, A., Le Péchoux, C., Swallow, C. J., Strauss, D., Meeus, P., van Coevorden, F., Stoldt, S., Stoeckle, E., Rutkowski, P., Rastrelli, M., Raut, C. P., Hompes, D., De Paoli, A., Sangalli, C., Honoré, C., Chung, P., Miah, A., Blay, J. Y., Fiore, M., Stelmes, J.-J., Dei Tos, A. P., Baldini, E. H., Litière, S., Marreaud, S., Gelderblom, H., and Haas, R. L. (2020). Preoperative radiotherapy plus surgery versus surgery alone for patients with primary retroperitoneal sarcoma (EORTC-62092: STRASS): a multicentre, open-label, randomised, phase 3 trial. *The Lancet. Oncology*, 21(10):1366–1377.
- Bridda, A., Padoan, I., Mencarelli, R., and Frego, M. (2007). Peritoneal Mesothelioma: A Review. *Medscape General Medicine*, 9(2):32.
- Cao, Y., Liao, C., Tan, A., Gao, Y., Mo, Z., and Gao, F. (2009). Meta-analysis of endoscopic submucosal dissection versus endoscopic mucosal resection for tumors of the gastrointestinal tract. *Endoscopy*, 41(9):751–757.
- Cardona, K. (2020). The STRASS trial: an important step in the right direction. *The Lancet. Oncology*, 21(10):1257–1258.
- Castoro, C., Scarpa, M., Cagol, M., Alfieri, R., Ruol, A., Cavallin, F., Michieletto, S., Zanchettin, G., Chiarion-Sileni, V., Corti, L., and Ancona, E. (2013). Complete clinical response after neoadjuvant chemoradiotherapy for squamous cell cancer of the thoracic oesophagus: is surgery always necessary? Journal of Gastrointestinal Surgery: Official Journal of the Society for Surgery of the Alimentary Tract, 17(8):1375–1381.
- Cercek, A., Lumish, M., Sinopoli, J., Weiss, J., Shia, J., Lamendola-Essel, M.,
  El Dika, I. H., Segal, N., Shcherba, M., Sugarman, R., Stadler, Z., Yaeger,
  R., Smith, J. J., Rousseau, B., Argiles, G., Patel, M., Desai, A., Saltz, L. B.,
  Widmar, M., Iyer, K., Zhang, J., Gianino, N., Crane, C., Romesser, P. B.,
  Pappou, E. P., Paty, P., Garcia-Aguilar, J., Gonen, M., Gollub, M., Weiser,
  M. R., Schalper, K. A., and Diaz, L. A. (2022). PD-1 Blockade in Mismatch
  Repair-Deficient, Locally Advanced Rectal Cancer. The New England Journal
  of Medicine, 386(25):2363-2376.
- Chalabi, M., Verschoor, Y. L., Tan, P. B., Balduzzi, S., Van Lent, A. U., Grootscholten, C., Dokter, S., Büller, N. V., Grotenhuis, B. A., Kuhlmann, K., Burger, J. W., Huibregtse, I. L., Aukema, T. S., Hendriks, E. R., Oosterling, S. J., Snaebjornsson, P., Voest, E. E., Wessels, L. F., Beets-Tan, R. G., Van Leerdam, M. E., Schumacher, T. N., van den Berg, J. G., Beets, G. L., and Haanen, J. B. (2024). Neoadjuvant Immunotherapy in Locally Advanced Mismatch Repair-Deficient Colon Cancer. The New England Journal of Medicine, 390(21):1949–1958.
- Chatterton, B. E., Ho Shon, I., Baldey, A., Lenzo, N., Patrikeos, A., Kelley, B., Wong, D., Ramshaw, J. E., and Scott, A. M. (2009). Positron emission

tomography changes management and prognostic stratification in patients with oesophageal cancer: results of a multicentre prospective study. *European Journal of Nuclear Medicine and Molecular Imaging*, 36(3):354–361.

- Chirieac, L. R., Swisher, S. G., Ajani, J. A., Komaki, R. R., Correa, A. M., Morris, J. S., Roth, J. A., Rashid, A., Hamilton, S. R., and Wu, T.-T. (2005). Posttherapy pathologic stage predicts survival in patients with esophageal carcinoma receiving preoperative chemoradiation. *Cancer*, 103(7):1347–1355.
- Conroy, T., Bosset, J.-F., Etienne, P.-L., Rio, E., François, ., Mesgouez-Nebout, N., Vendrely, V., Artignan, X., Bouché, O., Gargot, D., Boige, V., Bonichon-Lamichhane, N., Louvet, C., Morand, C., de la Fouchardière, C., Lamfichekh, N., Juzyna, B., Jouffroy-Zeller, C., Rullier, E., Marchal, F., Gourgou, S., Castan, F., Borg, C., and Unicancer Gastrointestinal Group and Partenariat de Recherche en Oncologie Digestive (PRODIGE) Group (2021). Neoadjuvant chemotherapy with FOLFIRINOX and preoperative chemoradiotherapy for patients with locally advanced rectal cancer (UNICANCER-PRODIGE 23): a multicentre, randomised, open-label, phase 3 trial. *The Lancet. Oncology*, 22(5):702–715.
- CReST Collaborative Group (2022). Colorectal Endoscopic Stenting Trial (CReST) for obstructing left-sided colorectal cancer: randomized clinical trial. British Journal of Surgery, 109(11):1073–1080.
- Cunningham, D., Allum, W. H., Stenning, S. P., Thompson, J. N., Van de Velde, C. J. H., Nicolson, M., Scarffe, J. H., Lofts, F. J., Falk, S. J., Iveson, T. J., Smith, D. B., Langley, R. E., Verma, M., Weeden, S., Chua, Y. J., and MAGIC Trial Participants, n. (2006). Perioperative chemotherapy versus surgery alone for resectable gastroesophageal cancer. The New England Journal of Medicine, 355(1):11–20.
- de Graaf, G. W., Ayantunde, A. A., Parsons, S. L., Duffy, J. P., and Welch, N. T. (2007). The role of staging laparoscopy in oesophagogastric cancers. European Journal of Surgical Oncology: The Journal of the European Society of Surgical Oncology and the British Association of Surgical Oncology, 33(8):988–992.
- Dorth, J. A., Pura, J. A., Palta, M., Willett, C. G., Uronis, H. E., D'Amico, T. A., and Czito, B. G. (2014). Patterns of recurrence after trimodality therapy for esophageal cancer. *Cancer*, 120(14):2099–2105.
- Dunbar, K. B. and Spechler, S. J. (2012). The risk of lymph-node metastases in patients with high-grade dysplasia or intramucosal carcinoma in Barrett's esophagus: a systematic review. *The American Journal of Gastroenterology*, 107(6):850–862; quiz 863.
- Frandsen, J., Boothe, D., Gaffney, D. K., Wilson, B. D., and Lloyd, S. (2015). Increased risk of death due to heart disease after radiotherapy for esophageal cancer. *Journal of Gastrointestinal Oncology*, 6(5):516–523.
- Furlong, H., Bass, G., Breathnach, O., O'Neill, B., Leen, E., and Walsh, T. N.

(2013). Targeting therapy for esophageal cancer in patients aged 70 and over. *Journal of Geriatric Oncology*, 4(2):107–113.

- Ganz, R. A., Overholt, B. F., Sharma, V. K., Fleischer, D. E., Shaheen, N. J., Lightdale, C. J., Freeman, S. R., Pruitt, R. E., Urayama, S. M., Gress, F., Pavey, D. A., Branch, M. S., Savides, T. J., Chang, K. J., Muthusamy, V. R., Bohorfoush, A. G., Pace, S. C., DeMeester, S. R., Eysselein, V. E., Panjehpour, M., Triadafilopoulos, G., and U.S. Multicenter Registry (2008). Circumferential ablation of Barrett's esophagus that contains high-grade dysplasia: a U.S. Multicenter Registry. *Gastrointestinal Endoscopy*, 68(1):35–40.
- Gharzai, L., Verma, V., Denniston, K. A., Bhirud, A. R., Bennion, N. R., and Lin, C. (2016). Radiation Therapy and Cardiac Death in Long-Term Survivors of Esophageal Cancer: An Analysis of the Surveillance, Epidemiology, and End Result Database. *PloS One*, 11(7):e0158916.
- Guttmann, D. M., Mitra, N., Bekelman, J., Metz, J. M., Plastaras, J., Feng, W., and Swisher-McClure, S. (2017). Improved Overall Survival with Aggressive Primary Tumor Radiotherapy for Patients with Metastatic Esophageal Cancer. Journal of Thoracic Oncology: Official Publication of the International Association for the Study of Lung Cancer, 12(7):1131–1142.
- Haidry, R. J., Dunn, J. M., Butt, M. A., Burnell, M. G., Gupta, A., Green, S.,
  Miah, H., Smart, H. L., Bhandari, P., Smith, L. A., Willert, R., Fullarton,
  G., Morris, J., Di Pietro, M., Gordon, C., Penman, I., Barr, H., Patel, P.,
  Boger, P., Kapoor, N., Mahon, B., Hoare, J., Narayanasamy, R., O'Toole,
  D., Cheong, E., Direkze, N. C., Ang, Y., Novelli, M., Banks, M. R., and Lovat, L. B. (2013). Radiofrequency ablation and endoscopic mucosal resection
  for dysplastic barrett's esophagus and early esophageal adenocarcinoma: outcomes of the UK National Halo RFA Registry. Gastroenterology, 145(1):87–95.
- Hansford, S., Kaurah, P., Li-Chang, H., Woo, M., Senz, J., Pinheiro, H., Schrader, K. A., Schaeffer, D. F., Shumansky, K., Zogopoulos, G., Santos, T. A., Claro, I., Carvalho, J., Nielsen, C., Padilla, S., Lum, A., Talhouk, A., Baker-Lange, K., Richardson, S., Lewis, I., Lindor, N. M., Pennell, E., MacMillan, A., Fernandez, B., Keller, G., Lynch, H., Shah, S. P., Guilford, P., Gallinger, S., Corso, G., Roviello, F., Caldas, C., Oliveira, C., Pharoah, P. D. P., and Huntsman, D. G. (2015). Hereditary Diffuse Gastric Cancer Syndrome: CDH1 Mutations and Beyond. JAMA oncology, 1(1):23–32.
- Heald, R. J. and Ryall, R. D. (1986). Recurrence and survival after total mesorectal excision for rectal cancer. *Lancet (London, England)*, 1(8496):1479–1482.
- Hollis, A. C., Quinn, L. M., Hodson, J., Evans, E., Plowright, J., Begum, R., Mitchell, H., Hallissey, M. T., Whiting, J. L., and Griffiths, E. A. (2017). Prognostic significance of tumor length in patients receiving esophagectomy for esophageal cancer. *Journal of Surgical Oncology*, 116(8):1114–1122.
- Hu, H., Kang, L., Zhang, J., Wu, Z., Wang, H., Huang, M., Lan, P., Wu, X., Wang, C., Cao, W., Hu, J., Huang, Y., Huang, L., Wang, H., Shi, L., Cai, Y.,

Shen, C., Ling, J., Xie, X., Cai, Y., He, X., Dou, R., Zhou, J., Ma, T., Zhang, X., Luo, S., Deng, W., Ling, L., Liu, H., and Deng, Y. (2022). Neoadjuvant PD-1 blockade with toripalimab, with or without celecoxib, in mismatch repair-deficient or microsatellite instability-high, locally advanced, colorectal cancer (PICC): a single-centre, parallel-group, non-comparative, randomised, phase 2 trial. The Lancet. Gastroenterology & Hepatology, 7(1):38–48.

- Hulscher, J. B. F., van Sandick, J. W., de Boer, A. G. E. M., Wijnhoven, B. P. L., Tijssen, J. G. P., Fockens, P., Stalmeier, P. F. M., ten Kate, F. J. W., van Dekken, H., Obertop, H., Tilanus, H. W., and van Lanschot, J. J. B. (2002). Extended transthoracic resection compared with limited transhiatal resection for adenocarcinoma of the esophagus. The New England Journal of Medicine, 347(21):1662–1669.
- James, R. D., Glynne-Jones, R., Meadows, H. M., Cunningham, D., Myint, A. S., Saunders, M. P., Maughan, T., McDonald, A., Essapen, S., Leslie, M., Falk, S., Wilson, C., Gollins, S., Begum, R., Ledermann, J., Kadalayil, L., and Sebag-Montefiore, D. (2013). Mitomycin or cisplatin chemoradiation with or without maintenance chemotherapy for treatment of squamous-cell carcinoma of the anus (ACT II): a randomised, phase 3, open-label, 2 × 2 factorial trial. The Lancet. Oncology, 14(6):516-524.
- Jin, J., Tang, Y., Hu, C., Jiang, L.-M., Jiang, J., Li, N., Liu, W.-Y., Chen, S.-L., Li, S., Lu, N.-N., Cai, Y., Li, Y.-H., Zhu, Y., Cheng, G.-H., Zhang, H.-Y., Wang, X., Zhu, S.-Y., Wang, J., Li, G.-F., Yang, J.-L., Zhang, K., Chi, Y., Yang, L., Zhou, H.-T., Zhou, A.-P., Zou, S.-M., Fang, H., Wang, S.-L., Zhang, H.-Z., Wang, X.-S., Wei, L.-C., Wang, W.-L., Liu, S.-X., Gao, Y.-H., and Li, Y.-X. (2022). Multicenter, Randomized, Phase III Trial of Short-Term Radiotherapy Plus Chemotherapy Versus Long-Term Chemoradiotherapy in Locally Advanced Rectal Cancer (STELLAR). Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 40(15):1681–1692.
- Kapiteijn, E., Marijnen, C. A., Nagtegaal, I. D., Putter, H., Steup, W. H., Wiggers, T., Rutten, H. J., Pahlman, L., Glimelius, B., van Krieken, J. H., Leer, J. W., van de Velde, C. J., and Dutch Colorectal Cancer Group (2001). Preoperative radiotherapy combined with total mesorectal excision for resectable rectal cancer. The New England Journal of Medicine, 345(9):638–646.
- Kato, H., Kuwano, H., Nakajima, M., Miyazaki, T., Yoshikawa, M., Ojima, H., Tsukada, K., Oriuchi, N., Inoue, T., and Endo, K. (2002). Comparison between positron emission tomography and computed tomography in the use of the assessment of esophageal carcinoma. *Cancer*, 94(4):921–928. tex.ids: kato921a.
- Khan, Q. J. and Fabian, C. J. (2010). How I Treat Vitamin D Deficiency. Journal of Oncology Practice, 6(2):97–101.
- Kim, T. J., Kim, H. Y., Lee, K. W., and Kim, M. S. (2009). Multimodality

assessment of esophageal cancer: preoperative staging and monitoring of response to the rapy. Radiographics: A Review Publication of the Radiological Society of North America, Inc, 29(2):403–421.

- Kofoed, S. C., Muhic, A., Baeksgaard, L., Jendresen, M., Gustafsen, J., Holm, J., Bardram, L., Brandt, B., Brenø, J., and Svendsen, L. B. (2012). Survival after adjuvant chemoradiotherapy or surgery alone in resectable adenocarcinoma at the gastro-esophageal junction. Scandinavian journal of surgery: SJS: official organ for the Finnish Surgical Society and the Scandinavian Surgical Society, 101(1):26-31.
- Lee, J., Lim, D. H., Kim, S., Park, S. H., Park, J. O., Park, Y. S., Lim, H. Y., Choi, M. G., Sohn, T. S., Noh, J. H., Bae, J. M., Ahn, Y. C., Sohn, I., Jung, S. H., Park, C. K., Kim, K.-M., and Kang, W. K. (2012). Phase III trial comparing capecitabine plus cisplatin versus capecitabine plus cisplatin with concurrent capecitabine radiotherapy in completely resected gastric cancer with D2 lymph node dissection: the ARTIST trial. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 30(3):268-273.
- Leichman, L. P., Goldman, B. H., Bohanes, P. O., Lenz, H. J., Thomas, C. R., Billingsley, K. G., Corless, C. L., Iqbal, S., Gold, P. J., Benedetti, J. K., Danenberg, K. D., and Blanke, C. D. (2011). S0356: a phase II clinical and prospective molecular trial with oxaliplatin, fluorouracil, and external-beam radiation therapy before surgery for patients with esophageal adenocarcinoma. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 29(34):4555–4560.
- Li, C. Y. and Alexander, H. R. (2018). Peritoneal Metastases from Malignant Mesothelioma. Surgical Oncology Clinics of North America, 27(3):539–549.
- Macdonald, J. S., Smalley, S. R., Benedetti, J., Hundahl, S. A., Estes, N. C., Stemmermann, G. N., Haller, D. G., Ajani, J. A., Gunderson, L. L., Jessup, J. M., and Martenson, J. A. (2001). Chemoradiotherapy after surgery compared with surgery alone for adenocarcinoma of the stomach or gastroesophageal junction. The New England Journal of Medicine, 345(10):725-730.
- Maish, M. S. and DeMeester, S. R. (2004). Endoscopic mucosal resection as a staging technique to determine the depth of invasion of esophageal adenocarcinoma. *The Annals of Thoracic Surgery*, 78(5):1777–1782.
- Mariette, C., Markar, S. R., Dabakuyo-Yonli, T. S., Meunier, B., Pezet, D.,
  Collet, D., D'Journo, X. B., Brigand, C., Perniceni, T., Carrère, N., Mabrut,
  J.-Y., Msika, S., Peschaud, F., Prudhomme, M., Bonnetain, F., and Piessen,
  G. (2019). Hybrid Minimally Invasive Esophagectomy for Esophageal Cancer.
  New England Journal of Medicine, 380(2):152–162. Publisher: Massachusetts
  Medical Society \_eprint: https://doi.org/10.1056/NEJMoa1805101.
- Markar, S. R., Gronnier, C., Pasquer, A., Duhamel, A., Beal, H., Théreaux, J., Gagnière, J., Lebreton, G., Brigand, C., Meunier, B., Collet, D., Mariette, C., and FREGAT working group FRENCH AFC (2016). Role of neoadjuvant

treatment in clinical T2N0M0 oesophageal cancer: results from a retrospective multi-center European study. European Journal of Cancer (Oxford, England: 1990), 56:59–68.

- Markar, S. R., Karthikesalingam, A., Penna, M., and Low, D. E. (2014). Assessment of short-term clinical outcomes following salvage esophagectomy for the treatment of esophageal malignancy: systematic review and pooled analysis. *Annals of Surgical Oncology*, 21(3):922–931.
- Martin, L. and Lagergren, P. (2009). Long-term weight change after oesophageal cancer surgery. *The British Journal of Surgery*, 96(11):1308–1314.
- McKeown, K. C. (1976). Total three-stage oesophagectomy for cancer of the oesophagus. *The British Journal of Surgery*, 63(4):259–262.
- MERCURY Study Group (2006). Diagnostic accuracy of preoperative magnetic resonance imaging in predicting curative resection of rectal cancer: prospective observational study. *BMJ (Clinical research ed.)*, 333(7572):779.
- Messager, M., Mirabel, X., Tresch, E., Paumier, A., Vendrely, V., Dahan, L., Glehen, O., Vasseur, F., Lacornerie, T., Piessen, G., El Hajbi, F., Robb, W. B., Clisant, S., Kramar, A., Mariette, C., and Adenis, A. (2016). Preoperative chemoradiation with paclitaxel-carboplatin or with fluorouracil-oxaliplatin-folinic acid (FOLFOX) for resectable esophageal and junctional cancer: the PROTECT-1402, randomized phase 2 trial. *BMC cancer*, 16:318.
- Minsky, B. D., Pajak, T. F., Ginsberg, R. J., Pisansky, T. M., Martenson, J., Komaki, R., Okawara, G., Rosenthal, S. A., and Kelsen, D. P. (2002). INT 0123 (Radiation Therapy Oncology Group 94-05) phase III trial of combined-modality therapy for esophageal cancer: high-dose versus standard-dose radiation therapy. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 20(5):1167-1174.
- Nigro, N., Seydel, H., Considine, B., Vaitkevicius, V., Leichman, L., and Kinzie, J. (1983). Combined preoperative radiation and chemotherapy for squamous cell carcinoma of the anal canal. *Cancer*, 51(10):1826. Publisher: Cancer.
- Noh, S. H., Park, S. R., Yang, H.-K., Chung, H. C., Chung, I.-J., Kim, S.-W., Kim, H.-H., Choi, J.-H., Kim, H.-K., Yu, W., Lee, J. I., Shin, D. B., Ji, J., Chen, J.-S., Lim, Y., Ha, S., Bang, Y.-J., and CLASSIC trial investigators (2014). Adjuvant capecitabine plus oxaliplatin for gastric cancer after D2 gastrectomy (CLASSIC): 5-year follow-up of an open-label, randomised phase 3 trial. *The Lancet. Oncology*, 15(12):1389–1396.
- Nurkin, S. J., Nava, H. R., Yendamuri, S., LeVea, C. M., Nwogu, C. E., Groman, A., Wilding, G., Bain, A. J., Hochwald, S. N., and Khushalani, N. I. (2014). Outcomes of endoscopic resection for high-grade dysplasia and esophageal cancer. Surgical Endoscopy, 28(4):1090–1095.
- Omloo, J. M. T., Lagarde, S. M., Hulscher, J. B. F., Reitsma, J. B., Fockens, P., van Dekken, H., Ten Kate, F. J. W., Obertop, H., Tilanus, H. W., and van

Lanschot, J. J. B. (2007). Extended transthoracic resection compared with limited transhiatal resection for adenocarcinoma of the mid/distal esophagus: five-year survival of a randomized clinical trial. *Annals of Surgery*, 246(6):992–1000; discussion 1000–1001.

- Oppedijk, V., van der Gaast, A., van Lanschot, J. J. B., van Hagen, P., van Os, R., van Rij, C. M., van der Sangen, M. J., Beukema, J. C., Rütten, H., Spruit, P. H., Reinders, J. G., Richel, D. J., van Berge Henegouwen, M. I., and Hulshof, M. C. C. M. (2014). Patterns of recurrence after surgery alone versus preoperative chemoradiotherapy and surgery in the CROSS trials. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 32(5):385–391.
- Orringer, M. B., Marshall, B., Chang, A. C., Lee, J., Pickens, A., and Lau, C. L. (2007). Two thousand transhiatal esophagectomies: changing trends, lessons learned. *Annals of Surgery*, 246(3):363–372; discussion 372–374.
- Orringer, M. B. and Sloan, H. (1978). Esophagectomy without thoracotomy. The Journal of Thoracic and Cardiovascular Surgery, 76(5):643–654.
- Ouattara, M., D'Journo, X. B., Loundou, A., Trousse, D., Dahan, L., Doddoli, C., Seitz, J. F., and Thomas, P.-A. (2012). Body mass index kinetics and risk factors of malnutrition one year after radical oesophagectomy for cancer. European Journal of Cardio-Thoracic Surgery: Official Journal of the European Association for Cardio-Thoracic Surgery, 41(5):1088–1093.
- Pasquali, S., Yim, G., Vohra, R. S., Mocellin, S., Nyanhongo, D., Marriott, P., Geh, J. I., and Griffiths, E. A. (2017). Survival After Neoadjuvant and Adjuvant Treatments Compared to Surgery Alone for Resectable Esophageal Carcinoma: A Network Meta-analysis. Annals of Surgery, 265(3):481–491.
- Pech, O., Behrens, A., May, A., Nachbar, L., Gossner, L., Rabenstein, T., Manner, H., Guenter, E., Huijsmans, J., Vieth, M., Stolte, M., and Ell, C. (2008). Long-term results and risk factor analysis for recurrence after curative endoscopic therapy in 349 patients with high-grade intraepithelial neoplasia and mucosal adenocarcinoma in Barrett's oesophagus. Gut, 57(9):1200–1206.
- Pech, O., May, A., Manner, H., Behrens, A., Pohl, J., Weferling, M., Hartmann, U., Manner, N., Huijsmans, J., Gossner, L., Rabenstein, T., Vieth, M., Stolte, M., and Ell, C. (2014). Long-term efficacy and safety of endoscopic resection for patients with mucosal adenocarcinoma of the esophagus. *Gastroenterology*, 146(3):652–660.e1. tex.ids: pechLongtermEfficacySafety2014.
- Penniment, M. G., De Ieso, P. B., Harvey, J. A., Stephens, S., Au, H.-J., O'Callaghan, C. J., Kneebone, A., Ngan, S. Y., Ward, I. G., Roy, R., Smith, J. G., Nijjar, T., Biagi, J. J., Mulroy, L. A., Wong, R., and TROG 03.01/CCTG ES.2 group (2018). Palliative chemoradiotherapy versus radiotherapy alone for dysphagia in advanced oesophageal cancer: a multicentre randomised controlled trial (TROG 03.01). The Lancet. Gastroenterology & Hepatology, 3(2):114–124.

Pettersson, D., Lörinc, E., Holm, T., Iversen, H., Cedermark, B., Glimelius, B., and Martling, A. (2015). Tumour regression in the randomized Stockholm III Trial of radiotherapy regimens for rectal cancer. *The British Journal of Surgery*, 102(8):972–978; discussion 978.

- Phoa, K. N., van Vilsteren, F. G. I., Weusten, B. L. A. M., Bisschops, R., Schoon, E. J., Ragunath, K., Fullarton, G., Di Pietro, M., Ravi, N., Visser, M., Offerhaus, G. J., Seldenrijk, C. A., Meijer, S. L., ten Kate, F. J. W., Tijssen, J. G. P., and Bergman, J. J. G. H. M. (2014). Radiofrequency ablation vs endoscopic surveillance for patients with Barrett esophagus and low-grade dysplasia: a randomized clinical trial. *JAMA*, 311(12):1209–1217. tex.ids: phoa1209a.
- Poultsides, G. A. and Paty, P. B. (2011). Reassessing the need for primary tumor surgery in unresectable metastatic colorectal cancer: overview and perspective. *Therapeutic Advances in Medical Oncology*, 3(1):35–42.
- Quirke, P., Durdey, P., Dixon, M. F., and Williams, N. S. (1986). Local recurrence of rectal adenocarcinoma due to inadequate surgical resection. Histopathological study of lateral tumour spread and surgical excision. *Lancet (London, England)*, 2(8514):996–999.
- Repici, A., Hassan, C., Carlino, A., Pagano, N., Zullo, A., Rando, G., Strangio, G., Romeo, F., Nicita, R., Rosati, R., and Malesci, A. (2010). Endoscopic submucosal dissection in patients with early esophageal squamous cell carcinoma: results from a prospective Western series. Gastrointestinal Endoscopy, 71(4):715–721.
- Rice, T. W., Mason, D. P., Murthy, S. C., Zuccaro, G., J., Adelstein, D. J., Rybicki, L. A., and Blackstone, E. H. (2007). T2N0M0 esophageal cancer. *J Thorac Cardiovasc Surg*, 133(2):317–24. Type: Journal Article.
- Roberts, M. E., Ranola, J. M. O., Marshall, M. L., Susswein, L. R., Graceffo, S., Bohnert, K., Tsai, G., Klein, R. T., Hruska, K. S., and Shirts, B. H. (2019). Comparison of CDH1 Penetrance Estimates in Clinically Ascertained Families vs Families Ascertained for Multiple Gastric Cancers. *JAMA oncology*, 5(9):1325–1331.
- Roh, M. S., Colangelo, L. H., O'Connell, M. J., Yothers, G., Deutsch, M., Allegra, C. J., Kahlenberg, M. S., Baez-Diaz, L., Ursiny, C. S., Petrelli, N. J., and Wolmark, N. (2009). Preoperative multimodality therapy improves disease-free survival in patients with carcinoma of the rectum: NSABP R-03. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 27(31):5124–5130.
- Rusch, V. W. (2004). Are cancers of the esophagus, gastroesophageal junction, and cardia one disease, two, or several? *Seminars in Oncology*, 31(4):444–449.
- Sauer, R., Fietkau, R., Wittekind, C., Martus, P., Rödel, C., Hohenberger, W., Jatzko, G., Sabitzer, H., Karstens, J. H., Becker, H., Hess, C., and Raab, R.

(2001). Adjuvant versus neoadjuvant radiochemotherapy for locally advanced rectal cancer. A progress report of a phase-III randomized trial (protocol CAO/ARO/AIO-94). Strahlentherapie Und Onkologie: Organ Der Deutschen Rontgengesellschaft ... [et Al], 177(4):173–181.

- Sebag-Montefiore, D., Stephens, R. J., Steele, R., Monson, J., Grieve, R., Khanna, S., Quirke, P., Couture, J., Metz, C. d., Myint, A. S., Bessell, E., Griffiths, G., Thompson, L. C., and Parmar, M. (2009). Preoperative radiotherapy versus selective postoperative chemoradiotherapy in patients with rectal cancer (MRC CR07 and NCIC-CTG C016): a multicentre, randomised trial. The Lancet, 373(9666):811–820. Publisher: Elsevier.
- Shaheen, N. J., Falk, G. W., Iyer, P. G., Gerson, L. B., and American College of Gastroenterology (2016). ACG Clinical Guideline: Diagnosis and Management of Barrett's Esophagus. *The American Journal of Gastroenterology*, 111(1):30–50; quiz 51.
- Shaheen, N. J., Sharma, P., Overholt, B. F., Wolfsen, H. C., Sampliner, R. E., Wang, K. K., Galanko, J. A., Bronner, M. P., Goldblum, J. R., Bennett, A. E., Jobe, B. A., Eisen, G. M., Fennerty, M. B., Hunter, J. G., Fleischer, D. E., Sharma, V. K., Hawes, R. H., Hoffman, B. J., Rothstein, R. I., Gordon, S. R., Mashimo, H., Chang, K. J., Muthusamy, V. R., Edmundowicz, S. A., Spechler, S. J., Siddiqui, A. A., Souza, R. F., Infantolino, A., Falk, G. W., Kimmey, M. B., Madanick, R. D., Chak, A., and Lightdale, C. J. (2009). Radiofrequency ablation in Barrett's esophagus with dysplasia. The New England Journal of Medicine, 360(22):2277–2288. tex.ids: shaheen2277a.
- Sharma, P., Kozarek, R., and Practice Parameters Committee of American College of Gastroenterology (2010). Role of esophageal stents in benign and malignant diseases. *The American Journal of Gastroenterology*, 105(2):258–273; quiz 274.
- Siewert, J. R., Stein, H. J., and Feith, M. (2006). Adenocarcinoma of the esophago-gastric junction. Scandinavian journal of surgery: SJS: official organ for the Finnish Surgical Society and the Scandinavian Surgical Society, 95(4):260–269.
- Sjoquist, K. M., Burmeister, B. H., Smithers, B. M., Zalcberg, J. R., Simes, R. J., Barbour, A., Gebski, V., and Australasian Gastro-Intestinal Trials Group (2011). Survival after neoadjuvant chemotherapy or chemoradiotherapy for resectable oesophageal carcinoma: an updated meta-analysis. *The Lancet. Oncology*, 12(7):681–692.
- Smalley, S. R., Benedetti, J. K., Haller, D. G., Hundahl, S. A., Estes, N. C., Ajani, J. A., Gunderson, L. L., Goldman, B., Martenson, J. A., Jessup, J. M., Stemmermann, G. N., Blanke, C. D., and Macdonald, J. S. (2012). Updated analysis of SWOG-directed intergroup study 0116: a phase III trial of adjuvant radiochemotherapy versus observation after curative gastric cancer

resection. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 30(19):2327–2333.

- Soetikno, R., Kaltenbach, T., Yeh, R., and Gotoda, T. (2005). Endoscopic mucosal resection for early cancers of the upper gastrointestinal tract. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 23(20):4490–4498.
- Speicher, P. J., Ganapathi, A. M., Englum, B. R., Hartwig, M. G., Onaitis, M. W., D'Amico, T. A., and Berry, M. F. (2014). Induction therapy does not improve survival for clinical stage T2N0 esophageal cancer. *Journal of Thoracic Oncology: Official Publication of the International Association for the Study of Lung Cancer*, 9(8):1195–1201. tex.ids: speicher1195a.
- Stahl, M., Stuschke, M., Lehmann, N., Meyer, H.-J., Walz, M. K., Seeber, S., Klump, B., Budach, W., Teichmann, R., Schmitt, M., Schmitt, G., Franke, C., and Wilke, H. (2005). Chemoradiation with and without surgery in patients with locally advanced squamous cell carcinoma of the esophagus. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 23(10):2310–2317.
- Stahl, M., Walz, M. K., Stuschke, M., Lehmann, N., Meyer, H.-J., Riera-Knorrenschild, J., Langer, P., Engenhart-Cabillic, R., Bitzer, M., Königsrainer, A., Budach, W., and Wilke, H. (2009). Phase III comparison of preoperative chemotherapy compared with chemoradiotherapy in patients with locally advanced adenocarcinoma of the esophagogastric junction. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 27(6):851–856.
- Sudo, K., Taketa, T., Correa, A. M., Campagna, M.-C., Wadhwa, R., Blum, M. A., Komaki, R., Lee, J. H., Bhutani, M. S., Weston, B., Skinner, H. D., Maru, D. M., Rice, D. C., Swisher, S. G., Hofstetter, W. L., and Ajani, J. A. (2013). Locoregional failure rate after preoperative chemoradiation of esophageal adenocarcinoma and the outcomes of salvage strategies. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 31(34):4306–4310.
- Sudo, K., Xiao, L., Wadhwa, R., Shiozaki, H., Elimova, E., Taketa, T., Blum, M. A., Lee, J. H., Bhutani, M. S., Weston, B., Ross, W. A., Komaki, R., Rice, D. C., Swisher, S. G., Hofstetter, W. L., Maru, D. M., Skinner, H. D., and Ajani, J. A. (2014). Importance of surveillance and success of salvage strategies after definitive chemoradiation in patients with esophageal cancer. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 32(30):3400–3405.
- Swedish Rectal Cancer Trial, Cedermark, B., Dahlberg, M., Glimelius, B., Påhlman, L., Rutqvist, L. E., and Wilking, N. (1997). Improved survival with preoperative radiotherapy in resectable rectal cancer. The New England Journal of Medicine, 336(14):980–987.

Swisher, S. G., Wynn, P., Putnam, J. B., Mosheim, M. B., Correa, A. M., Komaki, R. R., Ajani, J. A., Smythe, W. R., Vaporciyan, A. A., Roth, J. A., and Walsh, G. L. (2002). Salvage esophagectomy for recurrent tumors after definitive chemotherapy and radiotherapy. *The Journal of Thoracic and Cardiovascular Surgery*, 123(1):175–183.

- Taketa, T., Correa, A. M., Suzuki, A., Blum, M. A., Chien, P., Lee, J. H., Welsh, J., Lin, S. H., Maru, D. M., Erasmus, J. J., Bhutani, M. S., Weston, B., Rice, D. C., Vaporciyan, A. A., Hofstetter, W. L., Swisher, S. G., and Ajani, J. A. (2012). Outcome of trimodality-eligible esophagogastric cancer patients who declined surgery after preoperative chemoradiation. *Oncology*, 83(5):300–304. type: Journal Article tex.ids= taketa6.
- Taketa, T., Sudo, K., Correa, A. M., Wadhwa, R., Shiozaki, H., Elimova, E., Campagna, M.-C., Blum, M. A., Skinner, H. D., Komaki, R. U., Lee, J. H., Bhutani, M. S., Weston, B. R., Rice, D. C., Swisher, S. G., Maru, D. M., Hofstetter, W. L., and Ajani, J. A. (2014). Post-chemoradiation surgical pathology stage can customize the surveillance strategy in patients with esophageal adenocarcinoma. Journal of the National Comprehensive Cancer Network: JNCCN, 12(8):1139–1144.
- Taketa, T., Xiao, L., Sudo, K., Suzuki, A., Wadhwa, R., Blum, M. A., Lee, J. H., Weston, B., Bhutani, M. S., Skinner, H., Komaki, R., Maru, D. M., Rice, D. C., Swisher, S. G., Hofstetter, W. L., and Ajani, J. A. (2013). Propensity-based matching between esophagogastric cancer patients who had surgery and who declined surgery after preoperative chemoradiation. *Oncology*, 85(2):95–99.
- Taylor, F. G. M., Quirke, P., Heald, R. J., Moran, B. J., Blomqvist, L., Swift, I. R., Sebag-Montefiore, D., Tekkis, P., Brown, G., and Magnetic Resonance Imaging in Rectal Cancer European Equivalence Study Study Group (2014). Preoperative magnetic resonance imaging assessment of circumferential resection margin predicts disease-free survival and local recurrence: 5-year follow-up results of the MERCURY study. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 32(1):34-43.
- Terheggen, G., Horn, E. M., Vieth, M., Gabbert, H., Enderle, M., Neugebauer, A., Schumacher, B., and Neuhaus, H. (2017). A randomised trial of endoscopic submucosal dissection versus endoscopic mucosal resection for early Barrett's neoplasia. *Gut*, 66(5):783–793.
- Thomas, T., Singh, R., and Ragunath, K. (2009). Trimodal imaging-assisted endoscopic mucosal resection of early Barrett's neoplasia. *Surgical Endoscopy*, 23(7):1609–1613.
- Vakil, N., Morris, A. I., Marcon, N., Segalin, A., Peracchia, A., Bethge, N., Zuccaro, G., Bosco, J. J., and Jones, W. F. (2001). A prospective, randomized, controlled trial of covered expandable metal stents in the palliation of malig-

nant esophageal obstruction at the gastroesophageal junction. The American Journal of Gastroenterology, 96(6):1791–1796.

- van der Valk, M. J. M., Marijnen, C. A. M., van Etten, B., Dijkstra, E. A., Hilling, D. E., Kranenbarg, E. M.-K., Putter, H., Roodvoets, A. G. H., Bahadoer, R. R., Fokstuen, T., Ten Tije, A. J., Capdevila, J., Hendriks, M. P., Edhemovic, I., Cervantes, A. M. R., de Groot, D. J. A., Nilsson, P. J., Glimelius, B., van de Velde, C. J. H., Hospers, G. A. P., and Collaborative investigators (2020). Compliance and tolerability of short-course radiotherapy followed by preoperative chemotherapy and surgery for high-risk rectal cancer Results of the international randomized RAPIDO-trial. Radiotherapy and Oncology: Journal of the European Society for Therapeutic Radiology and Oncology, 147:75–83.
- van Gijn, W., Marijnen, C. A. M., Nagtegaal, I. D., Kranenbarg, E. M.-K., Putter, H., Wiggers, T., Rutten, H. J. T., Påhlman, L., Glimelius, B., van de Velde, C. J. H., and Dutch Colorectal Cancer Group (2011). Preoperative radiotherapy combined with total mesorectal excision for resectable rectal cancer: 12-year follow-up of the multicentre, randomised controlled TME trial. *The Lancet. Oncology*, 12(6):575–582.
- van Hagen, P., Hulshof, M. C. C. M., van Lanschot, J. J. B., Steyerberg, E. W., van Berge Henegouwen, M. I., Wijnhoven, B. P. L., Richel, D. J., Nieuwenhuijzen, G. a. P., Hospers, G. a. P., Bonenkamp, J. J., Cuesta, M. A., Blaisse, R. J. B., Busch, O. R. C., ten Kate, F. J. W., Creemers, G.-J., Punt, C. J. A., Plukker, J. T. M., Verheul, H. M. W., Spillenaar Bilgen, E. J., van Dekken, H., van der Sangen, M. J. C., Rozema, T., Biermann, K., Beukema, J. C., Piet, A. H. M., van Rij, C. M., Reinders, J. G., Tilanus, H. W., van der Gaast, A., and CROSS Group (2012). Preoperative chemoradiotherapy for esophageal or junctional cancer. *The New England Journal of Medicine*, 366(22):2074–2084.
- van Vliet, E. P. M., Heijenbrok-Kal, M. H., Hunink, M. G. M., Kuipers, E. J., and Siersema, P. D. (2008). Staging investigations for oesophageal cancer: a meta-analysis. *British Journal of Cancer*, 98(3):547–557.
- Visbal, A. L., Allen, M. S., Miller, D. L., Deschamps, C., Trastek, V. F., and Pairolero, P. C. (2001). Ivor Lewis esophagogastrectomy for esophageal cancer. The Annals of Thoracic Surgery, 71(6):1803–1808.
- Wang, X., Li, S., Sun, Y., Li, K., Shen, X., Xue, Y., Liang, P., Li, G., Chen, L., Zhao, Q., Li, G., Fu, W., Liang, H., Xin, H., Suo, J., Fang, X., Zheng, Z., Xu, Z., Chen, H., Zhou, Y., He, Y., Huang, H., Zhu, L., Yang, K., Ji, J., Ye, Y., Zhang, Z., Li, F., Wang, X., Tian, Y., Park, S., and Chen, L. (2021). The protocol of a prospective, multicenter, randomized, controlled phase III study evaluating different cycles of oxaliplatin combined with S-1 (SOX) as neoadjuvant chemotherapy for patients with locally advanced gastric cancer: RESONANCE-II trial. *BMC cancer*, 21(1):20.
- Worrell, S. G., Alicuben, E. T., Oh, D. S., Hagen, J. A., and DeMeester, S. R.

(2018). Accuracy of Clinical Staging and Outcome With Primary Resection for Local-Regionally Limited Esophageal Adenocarcinoma. *Annals of Surgery*, 267(3):484-488.

- Wouters, M. W. J. M., Karim-Kos, H. E., le Cessie, S., Wijnhoven, B. P. L., Stassen, L. P. S., Steup, W. H., Tilanus, H. W., and Tollenaar, R. a. E. M. (2009). Centralization of esophageal cancer surgery: does it improve clinical outcome? *Annals of Surgical Oncology*, 16(7):1789–1798.
- Xicola, R. M., Li, S., Rodriguez, N., Reinecke, P., Karam, R., Speare, V., Black, M. H., LaDuca, H., and Llor, X. (2019). Clinical features and cancer risk in families with pathogenic CDH1 variants irrespective of clinical criteria. *Journal of Medical Genetics*, 56(12):838–843.
- Yang, H., Liu, H., Chen, Y., Zhu, C., Fang, W., Yu, Z., Mao, W., Xiang, J., Han, Y., Chen, Z., Yang, H., Wang, J., Pang, Q., Zheng, X., Yang, H., Li, T., Lordick, F., D'Journo, X. B., Cerfolio, R. J., Korst, R. J., Novoa, N. M., Swanson, S. J., Brunelli, A., Ismail, M., Fernando, H. C., Zhang, X., Li, Q., Wang, G., Chen, B., Mao, T., Kong, M., Guo, X., Lin, T., Liu, M., Fu, J., and AME Thoracic Surgery Collaborative Group (2018). Neoadjuvant Chemoradiotherapy Followed by Surgery Versus Surgery Alone for Locally Advanced Squamous Cell Carcinoma of the Esophagus (NEOCRTEC5010): A Phase III Multicenter, Randomized, Open-Label Clinical Trial. Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology, 36(27):2796–2803.
- Ychou, M., Boige, V., Pignon, J.-P., Conroy, T., Bouché, O., Lebreton, G., Ducourtieux, M., Bedenne, L., Fabre, J.-M., Saint-Aubert, B., Genève, J., Lasser, P., and Rougier, P. (2011). Perioperative chemotherapy compared with surgery alone for resectable gastroesophageal adenocarcinoma: an FN-CLCC and FFCD multicenter phase III trial. *Journal of Clinical Oncology: Official Journal of the American Society of Clinical Oncology*, 29(13):1715–1721.
- Zhang, J. Q., Hooker, C. M., Brock, M. V., Shin, J., Lee, S., How, R., Franco, N., Prevas, H., Hulbert, A., and Yang, S. C. (2012). Neoadjuvant chemoradiation therapy is beneficial for clinical stage T2 N0 esophageal cancer patients due to inaccurate preoperative staging. *Ann Thorac Surg*, 93(2):429–35; discussion 436–7. tex.ids: zhang429a type: Journal Article.
- Zhao, X., Ren, Y., Hu, Y., Cui, N., Wang, X., and Cui, Y. (2018). Neoadjuvant chemotherapy versus neoadjuvant chemoradiotherapy for cancer of the esophagus or the gastroesophageal junction: A meta-analysis based on clinical trials. *PloS One*, 13(8):e0202185.
- Zhou, C., Zhang, L., Wang, H., Ma, X., Shi, B., Chen, W., He, J., Wang, K., Liu, P., and Ren, Y. (2015). Superiority of Minimally Invasive Oesophagectomy in Reducing In-Hospital Mortality of Patients with Resectable Oesophageal Cancer: A Meta-Analysis. *PloS One*, 10(7):e0132889. tex.ids= zhoue0132889a.