

ESMO Clinical Practice Guidelines

Resectable locally advanced oesophagogastric cancer

Clinical Case Presentation

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DISCLOSURES

Honoraria for advisory role to Bristol-Meier Squibb, Five Prime Therapeutics and Gritstone Oncology

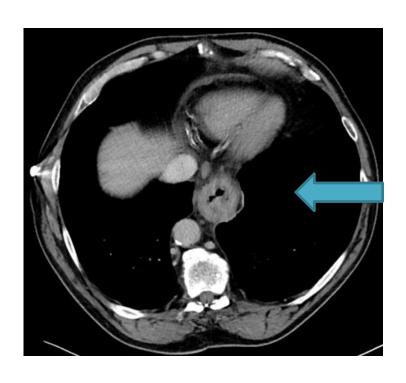


Patient history

- Presentation: 69 year-old male with 3 month history of progressive dysphagia and 7 kg weight loss, BMI 31, no history of reflux
- Medical history: Hypertension and past smoker (20 pack year history), good general health, ECOG PS=1
- Gastroscopy: circumferential tumour in lower oesophagus extending into upper part of gastroesophageal junction (Siewert I)
- Histology: moderately differentiated adenocarcinoma, HER2 positive



Staging investigations: CT



- CT demonstrates:
 - Type I gastroesophageal junctional cancer
 - Possible infiltration of left pleura
 - Lymph nodes in peri-oesophageal tissue

CT staging: cT4N2M0



Investigations

Q1. Which investigations are recommended prior to making a treatment decision?

- 1. Laparoscopy
- 2. Endoscopic ultrasound (EUS)
- 3. PET
- 4. All of the above



Investigations

Q1. Which investigations are recommended prior to making a treatment decision?

- 1. Laparoscopy (optional, if tumour infiltrates the cardia)
- 2. Endoscopic ultrasound (advisable)
- 3. PET (if available, will detect metastases in up to 15%)
- 4. All of the above <



Staging investigations



PET demonstrates

- 6cm thickening of the oesophagus
- No evidence of metastatic disease

EUS – cT4aN2



Q2. What is the current recommended treatment approach?

- 1. Surgery alone
- 2. Surgery then adjuvant chemotherapy or chemoradiotherapy
- 3. Perioperative chemotherapy
- 4. Perioperative chemotherapy + trastuzumab
- 5. Neoadjuvant chemoradiotherapy
- 6. Definitive chemoradiotherapy
- 7. Either perioperative chemotherapy or neoadjuvant chemoradiotherapy

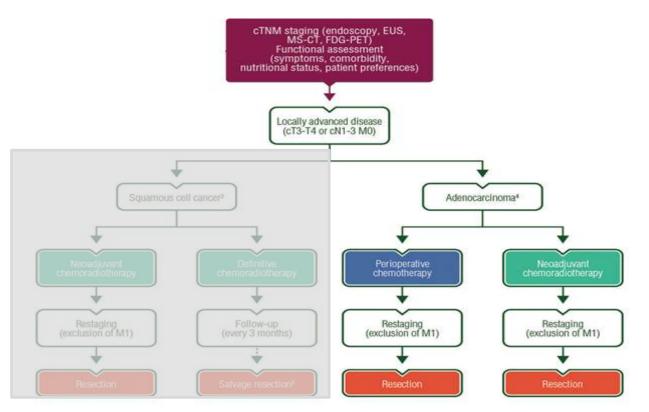


Q2. What is the current recommended treatment approach?

- 1. Surgery alone
- 2. Surgery then adjuvant chemotherapy or chemoradiotherapy
- 3. Perioperative chemotherapy plus surgery
- 4. Perioperative chemotherapy + trastuzumab
- 5. Neoadjuvant chemoradiotherapy (CRT) then surgery
- 6. Definitive chemoradiotherapy
- 7. Either perioperative chemotherapy or neoadjuvant chemoradiotherapy ✓



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EUS, endoscopic ultrasound; FDG-PET, fluorodeoxy glucose-positron-emission tomography; MS-CT, multiple-slice helical computed tomography; TNM, tumour, node and metastasis



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Neoadjuvant chemoradiotherapy and perioperative chemotherapy are both validated treatment options for gastroesophageal adenocarcinoma

Neoadjuvant CRT (CROSS)

- 5 year OS in CROSS (ADC+SCC) was 47% versus 33% (CRT versus surgery arms)
- Updated median OS for adenocarcinoma patients in CROSS was 43.2 months (24-9–61-4) versus 27-1 months (13-0-41-2) in the surgery alone arm (Shapiro 2015)

Perioperative chemotherapy (FLOT/FFCD/MAGIC)

- 5 year survival in MAGIC was 36% versus 23% (chemotherapy versus surgery arms); in FLOT4 predicted 5 years OS for FLOT arm was 45% (Al-Batran, ASCO 2017)
- Median overall survival for patients treated with perioperative FLOT was 50 months (38-na)

ADC, adenocarcinoma; CRT, chemoradiotherapy; OS, overall survival; SCC, squamous cell carcinoma



Key recommendations

- MDT discussion is mandatory
 - Upfront interdisciplinary planning of treatment
 - Consideration of performance status, co-morbidities
 - Correct nutritional status; jejunostomy is preferred to stent
- Surgery
 - High volume centre



- Treated with 3 cycles of ECX chemotherapy
- One episode of febrile neutropenia
- Post treatment CT: response, tumour now T3
- Lymph nodes reduced in size: staging cT3N2M0
- Surgical margin felt not at risk



Q3. What is the current recommended treatment approach?

- 1. Another three cycles of pre-operative ECX?
- 2. Switch to preoperative FLOT chemotherapy?
- 3. Switch to pre-operative chemoradiation?
- 4. Proceed with radical transthoracic Ivor-Lewis-oesophago-gastrectomy?
- 5. Watch-and-wait, re-assessment after three months?



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- 1. Another three cycles of pre-operative ECX?
- 2. Switch to preoperative FLOT chemotherapy?
- 3. Switch to pre-operative chemoradiation?
- 4. Proceed with radical transthoracic Ivor-Lewis-oesophago-gastrectomy? ✓
- 5. Watch-and-wait, re-assessment after three months?



- MDT decision was proceed to surgery
- Ivor Lewis oesophagogastrectomy performed
 - Good recovery post-operatively, weight stabilised
- Pathology showed UICC/AJCC
 - T3 tumour, 2/26 lymph nodes positive
 - R0 resection (negative margins)
 - Tumour regression: Mandard grade 3



Post-operative treatment

Q4. What is the recommended treatment now?

- 1. No further treatment
- 2. Continue same chemotherapy as pre-operatively
- 3. Switch chemotherapy
- 4. Adjuvant chemoradiotherapy



Post-operative treatment

Q4. What is the recommended treatment now?

- 1. No further treatment
- 2. Continue same chemotherapy as pre-operatively (if patient is fit for this) ✓
- 3. Switch chemotherapy
- 4. Adjuvant chemoradiotherapy