

<b>Indication</b>	For the treatment of adenocarcinoma, undifferentiated cancer or squamous cell carcinoma of the oesophagus.
<b>Treatment Intent</b>	Radical
<b>Frequency and number of cycles</b>	2 cycles of primary chemotherapy given every 21 days, followed by 2 x 21 day cycles of chemotherapy given concurrently with radiotherapy (50Gy/25 fractions). *NB close monitoring towards the end of radiotherapy is required, if necessary capecitabine may be discontinued on completion of radiotherapy.
<b>Monitoring Parameters pre-treatment</b>	<ul style="list-style-type: none"> <li>• <b>Virology screening:</b> All new patients referred for systemic anti-cancer treatment should be screened for hepatitis B and C and the result reviewed prior to the start of treatment. Patients not previously tested who are starting a new line of treatment, should also be screened for hepatitis B and C. Further virology screening will be performed following individual risk assessment and clinician discretion.</li> <li>• <b>DPD testing:</b> DPD testing must be undertaken in all patients before starting treatment; the result must be checked before treatment is started.</li> <li>• <b>Cardiotoxicity:</b> <ul style="list-style-type: none"> <li>○ Caution in patients with prior history of coronary heart disease, arrhythmias and angina pectoris.</li> <li>○ ECG baseline and during treatment as clinically indicated.</li> </ul> </li> <li>• EDTA should be used to measure GFR prior to cycle 1 or 2.</li> <li>• C+G may be used to estimate CrCl if delay in obtaining EDTA result.</li> <li>• <b>Monitor FBC, LFT's and U&amp;Es</b> prior to start of treatment, at each cycle and weekly FBC during chemoradiotherapy (cycles 3 and 4). <ul style="list-style-type: none"> <li>○ Prior to the start of treatment neuts <math>\geq 1.5</math> and PLT <math>\geq 100</math>.</li> <li>○ During treatment: <ul style="list-style-type: none"> <li>○ If neuts 1 - <math>&lt;1.5</math> and PLT 75-99 discuss with consultant.</li> <li>○ If neuts 0.5 - <math>&lt;1</math> or PLT 50 - <math>&lt;75</math> or any episode of neutropenic sepsis during the previous cycle stop chemotherapy until recovery. Restart with 25% dose reduction of capecitabine and carboplatin.</li> <li>○ If neuts <math>&lt;0.5</math> and/or PLT <math>&lt;50</math> stop chemotherapy until recovery. Restart with 50% dose reduction of capecitabine and carboplatin.</li> <li>○ Given that this is potentially curative treatment, consider the use of GCSF in the management of neutropenia.</li> </ul> </li> </ul> </li> <li>• <b>Hepatic impairment:</b> no recommended dose adjustment in hepatic impairment.</li> <li>• <b>Renal impairment:</b> <ul style="list-style-type: none"> <li>○ Regimen contraindicated if CrCl <math>&lt;30</math>ml/min.</li> <li>○ If CrCl <math>&lt; 50</math> ml/min dose reduce capecitabine. d/w consultant, consider 25% dose reduction of capecitabine (see SPC).</li> </ul> </li> <li>• <b>Infusion-related reactions:</b> <ul style="list-style-type: none"> <li>○ <b>Carboplatin:</b> Mild/moderate reactions (grade 1-2): If symptoms resolve after treatment with hydrocortisone and chlorphenamine, the infusion may be restarted at 50% rate for 30 mins, then, if no further reaction, increase to 100% rate. If symptoms do not resolve after treatment with hydrocortisone and chlorphenamine, do not restart the infusion. At consultant's discretion, patients may be rechallenged at a later date with additional prophylaxis. In the event of further reaction (grade 1-3), stop infusion and consider alternative treatment.</li> <li>○ Severe (grade 3): Do not restart infusion. Consider alternative treatment.</li> </ul> </li> </ul>

Protocol No	UGI-076	Kent and Medway SACT Protocol Disclaimer: No responsibility will be accepted for the accuracy of this information when used elsewhere.	
Version	2	Written by	M.Archer
Supersedes version	1	Checked by	C.Waters V2 A.Ho V1 V2 minor change pharmacist approval only
Date	25.04.2023	Authorising consultant (usually NOG Chair)	S.Forner V1

	<p>Anaphylaxis (grade 4): Follow anaphylaxis protocol. Discontinue permanently and consider alternative treatment.</p> <ul style="list-style-type: none"> <li>• <b>Dose Modification:</b> Interrupt capecitabine in the event of <math>\geq</math> grade 2 non-haematological toxicity (with the exception of side effects such as alopecia, alteration in taste etc, considered to be not serious) until resolution of toxicity to grade 0-1. Dose reduction should be considered if grade 3 or 4 non-haematological toxicity or repeat appearance of grade 2 (except N&amp;V and alopecia). Delay until resolution of toxicity to <math>\leq</math> grade 1.</li> <li>• <b>Skin reactions:</b> Capecitabine can induce severe skin reactions such as Stevens-Johnson syndrome and Toxic Epidermal Necrolysis. Patients should be informed of the possibility of such reactions and informed to seek urgent medical advice should any symptoms of a severe skin reaction occur. Treatment should be permanently discontinued in affected patients.</li> <li>• <b>Common drug interactions (for comprehensive list refer to BNF/SPC):</b> <ul style="list-style-type: none"> <li>○ <b>Carboplatin:</b> Caution when used concurrently with other nephrotoxic or ototoxic drugs.</li> <li>○ <b>Capecitabine</b> must not be given with concurrent sorivudine or derivatives (e.g brivudine), see SPC. Monitor PT and INR regularly in patients taking coumarin-derivative anticoagulants. Monitor phenytoin levels with concomitant use. Caution with folic acid or folic acid – potential for increased toxicity. Avoid concomitant allopurinol.</li> </ul> </li> <li>• <b>Driving and operating machinery:</b> Capecitabine may cause dizziness, fatigue and nausea. Patients should be aware this may affect their ability to drive or operate machinery.</li> <li>• For oral self-administration: refer to local Trust policy on oral anti-cancer medicines and supply Patient Information Leaflet and Macmillan information sheet.</li> </ul>
<b>References</b>	KMCC proforma UGI-007 V4 and UGI-071 draft protocol. UGI NOG 22.11.2022. SCOPE 2 trial protocol V8

NB For funding information, refer to CDF and NICE Drugs Funding List

Protocol No	UGI-076	Kent and Medway SACT Protocol Disclaimer: No responsibility will be accepted for the accuracy of this information when used elsewhere.	
Version	2	Written by	M.Archer
Supersedes version	1	Checked by	C.Waters V2 A.Ho V1 V2 minor change pharmacist approval only
Date	25.04.2023	Authorising consultant (usually NOG Chair)	S.Forner V1

**Cycle 1-4: 21-day cycle (cycle 3 and 4 current with radiotherapy)**

Day	Drug	Dose	Route	Infusion duration	Administration
1	Dexamethasone	8mg	PO		
	Ondansetron	<75yrs 16mg >/=75yrs 8mg	IV	15min	Sodium Chloride 0.9% 50ml
	<b>CARBOPLATIN (AUC= 5)</b>	<b>DOSE = AUC x (GFR + 25) Max dose 700mg</b>	IV	30min	In Glucose 5% 500ml
TTO	Drug	Dose	Route	Directions	
.	<b>CAPECITABINE*</b>	<b>1250mg/m<sup>2</sup>/day</b>  In 2 divided doses	PO	<b>Continuously for 21 days.</b>  Take within 30 mins after food and approximately every 12 hours.  Available as 500mg and 150mg tablets	
	Dexamethasone	6mg	PO	OM for 3 days	
	Metoclopramide	10mg	PO	10mg TDS for 3 days and then 10mg up to 3 times a day as required. Do not take for more than 5 days continuously.	

**\*NB close monitoring towards the end of radiotherapy is required, if necessary capecitabine may be discontinued on completion of radiotherapy.**

Protocol No	UGI-076	Kent and Medway SACT Protocol Disclaimer: No responsibility will be accepted for the accuracy of this information when used elsewhere.	
Version	2	Written by	M.Archer
Supersedes version	1	Checked by	C.Waters V2 A.Ho V1 V2 minor change pharmacist approval only
Date	25.04.2023	Authorising consultant (usually NOG Chair)	S.Forner V1