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| <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).<br><br>*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>• Consider <b>audiology</b> test for hearing impaired patients and monitor all patients for ototoxicity through-out treatment.</li> <li>• <b>Cardiotoxicity:</b> caution in patients with prior history of coronary heart disease, arrhythmias and angina pectoris.</li> <li>• <b>ECG</b> baseline and during treatment as clinically indicated.</li> <li>• <b>Renal impairment:</b> <ul style="list-style-type: none"> <li>• C+G should be used to measure CrCl prior to cycle 1.</li> <li>• If CrCl &lt;60ml/min then obtain EDTA result.</li> <li>• If CrCl 30-59ml/min consider dose reduction of cisplatin or consider carboplatin.</li> <li>• If CrCl &lt; 50 ml/min dose reduce capecitabine (see SPC)</li> <li>• Regimen contraindicated if CrCl &lt;30ml/min.</li> </ul> </li> <li>• <b>Hepatic Impairment:</b> no recommended dose adjustment in hepatic impairment.</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 if neuts <math>\geq 1</math> and PLT <math>\geq 75</math> continue with treatment.</li> <li>○ If neuts 0.5 - &lt;1 or PLT 50 - &lt;75 or any episode of neutropenic sepsis during the previous cycle stop chemotherapy until recovery. Restart with 25% dose reduction cisplatin and capecitabine.</li> <li>○ If neuts &lt;0.5 and/or PLT &lt;50 stop chemotherapy until recovery. Restart with 50% dose reduction cisplatin and capecitabine.</li> <li>○ Given that this is potentially curative treatment, consider the use of GCSF in the management of neutropenia.</li> </ul> </li> <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> </ul> |

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|--------------------|--------------|---|-----------------------|
| Protocol No        | UGI-072      | Kent and Medway SACT Protocol<br>Disclaimer: No responsibility will be accepted for the accuracy of this information when used elsewhere. |                       |
| Version            | V1           | Written by  | M.Archer              |
| Supersedes version | New protocol | Checked by  | C.Waters<br>O.Adebayo |
| Date               | 21.03.2023   | Authorising consultant (usually NOG Chair)  | M.Cominos             |

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|-------------------|--|
|                   | <ul style="list-style-type: none"> <li>• <b>Drug interactions (for comprehensive list refer to BNF/SPC):</b> <ul style="list-style-type: none"> <li>○ <b>Capecitabine:</b> must not be given with concurrent sorivudine or derivatives (e.g. brivudine), see SPC.<br/>Monitor PT and INR regularly in patients taking coumarin-derivative anticoagulants.<br/>Monitor phenytoin levels with concomitant use.<br/>Caution with folinic acid or folic acid – potential for increased toxicity. Avoid concomitant allopurinol.</li> <li>○ <b>Cisplatin:</b> Caution when used concurrently with other nephrotoxic or ototoxic drugs.</li> <li>○ Caution in patients receiving phenytoin, levels may be affected.</li> </ul> </li> <li>• 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.</li> </ul> |
| <b>References</b> | KMCC proforma UGI-010 V5 SPC accessed online 05.08.2022 SCOPE 2 trial protocol V8  |

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

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

| Day | Drug                 | Dose  | Route | Infusion Duration   | Administration                                |
|-----|----------------------|---|-------|---|---|
| 1   | Sodium Chloride 0.9% | 1000ml  | IV    | 2hours  | + 20mmol KCL + 10mmol Mg <sup>2+</sup>        |
|     | Mannitol 10%         | 200ml   | IV    | 15min   |   |
|     | Ondansetron          | <75yrs 16mg<br>>=75yrs 8mg                            | IV    | 15min   | Sodium Chloride 0.9% 50ml                     |
|     | Dexamethasone        | 8mg   | PO    |   |   |
|     | <b>CISPLATIN</b>     | <b>60mg/m<sup>2</sup></b>                             | IV    | 2 hours   | In Sodium Chloride 0.9% 1000ml                |
|     | Furosemide           | 40mg  | IV/PO |   | If urine output <100ml/hr or weight gain >1kg |
|     | Sodium Chloride 0.9% | 1000ml  | IV    | 2 hours   | + 20mmol KCL + 10mmol Mg <sup>2+</sup>        |
|     | Sodium Chloride 0.9% | 500ml   | IV    | 1 hour  | Or 500ml water, orally                        |
|     | *(Furosemide)        | 40mg  | IV/PO | <b>*only if required</b>  | If patient remains in a 2L positive balance   |
| TTO | Drug                 | Dose  | Route | Directions  |   |
|     | <b>CAPECITABINE*</b> | <b>1250mg/m<sup>2</sup>/day</b><br>In 2 divided doses | PO    | <b>For 21 days continuously.</b><br><br>Take within 30 mins after food and approximately every 12 hours.<br>Available as 500mg & 150mg. |   |
|     | 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.<br><br>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.**

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