

Gastric and gastroesophageal metastatic ciSplatin fluorouracil and trastuzumab

ID: 1217 v.7

Endorsed

Essential Medicine List

▲ Fluoropyrimidine overdose or overexposure:

Fluoropyrimidine overdose or overexposure may result in severe or life-threatening toxicity. An antidote is available and is highly effective if given within 96 hours. Read more about fluoropyrimidine overdose or overexposure.

Check for clinical trials in this patient group. Link to Australian Clinical Trials website

The anticancer drug(s) in this protocol <u>may</u> have been included in the ADDIKD guideline. Dose recommendations in kidney dysfunction have yet to be updated to align with the ADDIKD guideline. Recommendations will be updated once the individual protocol has been evaluated by the reference committee. For further information refer to the ADDIKD guideline. To assist with calculations, use the <u>eviQ Estimated Glomerular Filtration Rate (eGFR) calculator</u>.

International Consensus Guideline for Anticancer Drug Dosing in Kidney Dysfunction (ADDIKD)

Click here



Related pages:

2022

- Gastric and gastroesophageal metastatic ciSplatin capecitabine and trastuzumab
- · Gastric and gastroesophageal metastatic FOLFOX6 (modified) and trastuzumab
- · Gastric and gastroesophageal metastatic CAPOX (XELOX) (capecitabine and oxaliplatin) and trastuzumab

Treatment schedule - Overview

Cycle 1

Drug	Dose	Route	Day
Trastuzumab	8 mg/kg (loading dose only)	IV infusion	1
ciSplatin	80 mg/m ²	IV infusion	1
Fluorouracil	4,000 mg/m 2 (equivalent to 1000 mg/m 2 /day)	CIV via pump over 96 hours*	1

Cycle 2 to 6

Drug	Dose	Route	Day
Trastuzumab	6 mg/kg (subsequent doses)	IV infusion	1
ciSplatin	80 mg/m ²	IV infusion	1
Fluorouracil	4,000 mg/m² (equivalent to 1000 mg/m²/day)	CIV via pump over 96 hours*	1

Cycle 7 and further cycles

Drug	Dose	Route	Day
Trastuzumab	6 mg/kg (subsequent doses)	IV infusion	1

^{*}Although in the original trial fluorouracil was administered over 120 hours, it is felt that a 96 hour infusion is more commonly used.

However, a shorter infusion may result in increased gastrointestinal toxicity.

Frequency: 21 days

Cycles: Continuous until disease progression or unacceptable toxicity (in the clinical trial cisplatin, fluorouracil and

trastuzumab were given for 6 cycles, after which trastuzumab monotherapy should be continued in patients who

are responding to treatment)

Drug status: Trastuzumab is PBS authority

Cisplatin and fluorouracil are on the PBS general schedule

Cost: ~ \$660

Treatment schedule - Detail

The supportive therapies (e.g. antiemetics, premedications, etc.), infusion times, diluents, volumes and routes of administration, if included, are listed as defaults. They may vary between institutions and can be substituted to reflect individual institutional policy.

Antiemetics if included in the treatment schedule are based upon recommendations from national and international guidelines. These are **defaults only** and may be substituted to reflect individual institutional policy. Select here for recommended doses of alternative antiemetics.

Cycle 1

Day 1		
Netupitant	300 mg (P0)	60 minutes before chemotherapy (fixed dose preparation with palonosetron)
Palonosetron	0.5 mg (PO)	60 minutes before chemotherapy (fixed dose preparation with netupitant)
Dexamethasone	12 mg (PO)	60 minutes before chemotherapy
Trastuzumab	8 mg/kg (IV infusion)	in 250 mL sodium chloride 0.9% over 90 minutes (loading dose; cycle 1 only)
ciSplatin	80 mg/m ² (IV infusion)	in 1000 mL sodium chloride 0.9% over 60 minutes
Fluorouracil	4,000 mg/m ² (CIV)	via ambulatory infusion pump over 96 hours (equivalent to 1000 mg/m²/day)

Day 2 to 4		
Dexamethasone	8 mg (PO)	ONCE a day (or in divided doses) with or after food

Cycle 2 to 6

Day 1		
Netupitant	300 mg (PO)	60 minutes before chemotherapy (fixed dose preparation with palonosetron)
Palonosetron	0.5 mg (P0)	60 minutes before chemotherapy (fixed dose preparation with netupitant)
Dexamethasone	12 mg (PO)	60 minutes before chemotherapy
Trastuzumab	6 mg/kg (IV infusion)	in 250 mL sodium chloride 0.9% over 30 minutes (if the initial loading dose was well tolerated)
ciSplatin	80 mg/m ² (IV infusion)	in 1000 mL sodium chloride 0.9% over 60 minutes
Fluorouracil	4,000 mg/m ² (CIV)	via ambulatory infusion pump over 96 hours (equivalent to 1000 mg/m²/day)

Day 2 to 4		
Dexamethasone	8 mg (PO)	ONCE a day (or in divided doses) with or after food

Cycle 7 and further cycles

Day 1		
Trastuzumab	6 mg/kg (IV infusion)	in 250 mL sodium chloride 0.9% over 30 minutes (if the initial loading dose was well tolerated)

Although in the original trial fluorouracil was administered over 120 hours, it is felt that a 96 hour infusion is more commonly used. However, a shorter infusion may result in increased gastrointestinal toxicity.

Frequency: 21 days

Cycles: Continuous until disease progression or unacceptable toxicity (in the clinical trial cisplatin, fluorouracil and

trastuzumab were given for 6 cycles, after which trastuzumab monotherapy should be continued in patients who

are responding to treatment)

Indications and patient population

Indications:

• HER-2 positive (IHC 3+ or FISH +ve) advanced adenocarcinoma of the stomach or gastro-oesophageal junction.

Exclusions:

- left ventricular ejection fraction (LVEF) less than 45%
- moderate/severe renal impairment (creatinine clearance less than 60 mL/min).

Cautions:

- · pre existing neuropathies
- significant hearing impairment/tinnitus.

Clinical information

Safety alert fluoropyrimidines	Fluoropyrimidines can be administered by different routes and schedules with each method having associated increased risk of certain side effects. Fluoropyrimidine overdose or overexposure is a rare but potentially life threatening side effect of this drug class and can occur by any route of administration. An antidote is available and highly effective if given within 96 hours. Read more about the medication safety alert for infusional fluorouracil and fluoropyrimidine overdose or overexposure
Venous access	Central venous access device (CVAD) is required to administer this treatment. Read more about central venous access device line selection
Hypersensitivity/infusion related reaction	Although hypersensitivity with trastuzumab is common, severe hypersensitivity reactions are uncommon. Use with caution in patients with dyspnoea at rest from pulmonary/cardiac conditions as increased risk of infusion related symptoms. Read more about Hypersensitivity reaction
Emetogenicity HIGH	Suggested default antiemetics have been added to the treatment schedule, and may be substituted to reflect institutional policy. Ensure that patients also have sufficient antiemetics for breakthrough emesis: Metoclopramide 10 mg three times a day when necessary (maximum of 30 mg/24 hours, up to 5 days) OR Prochlorperazine 10 mg PO every 6 hours when necessary. Read more about preventing anti-cancer therapy induced nausea and vomiting

Cardiac toxicity associated with HER-2 directed agents	Patients receiving HER-2 directed agents are at an increased risk of cardiotoxicity e.g. asymptomatic decrease in the left ventricular ejection fraction (LVEF) and congestive heart failure (CHF).
	In patients with a LVEF less than 45% and/or symptomatic heart failure HER-2 directed therapy should be avoided, except in the metastatic setting when breast cancer is life-threatening and where a cardiologist is also involved.
	Concurrent anthracycline and HER-2 directed therapy is not recommended for extended periods of time.
	Baseline and 3 monthly cardiac function tests are required during treatment. In the metastatic setting, after the first 12 months of therapy, if there are no cardiac complications, the frequency of cardiac assessments may be reduced at the discretion of the treating clinician unless there has been recent exposure to anthracyclines.
	Read more about cardiac toxicity associated with HER-2 targeted agents
Cardiac toxicity	Cardiac toxicity is a serious complication that can occur during treatment with fluorouracil. Patients treated with fluorouracil, especially those with a prior history of cardiac disease or other risk factors, should be carefully monitored during therapy.
	Read more about cardiac toxicity associated with anti-cancer drugs
Dihydropyrimidine dehydrogenase (DPD) enzyme deficiency	Rare, life-threatening toxicities such as mucositis, neutropenia, neurotoxicity and diarrhoea have been reported following administration of fluoropyrimidines (e.g. fluorouracil and capecitabine). Severe unexplained toxicities require investigation prior to continuing with treatment. Testing for DPD enzyme deficiency is available in Australia but not currently reimbursed.
	Read more about dihydropyrimidine dehydrogenase (DPD) enzyme deficiency
Diarrhoea	Antidiarrhoeals (e.g. loperamide) are usually prescribed with this treatment. Read more about treatment induced diarrhoea
Hydration	Hydration helps to prevent cisplatin-induced nephrotoxicity.
,	The default regimen is appropriate for patients with normal electrolytes, kidney function, fluid status etc. and should be adjusted according to individual requirements.
	Read more about cisplatin hydration regimens
Peripheral neuropathy	Assess prior to each treatment. If a patient experiences grade 2 or greater peripheral neuropathy, a dose reduction, delay, or omission of treatment may be required; review by medical officer before commencing treatment.
	Read more about peripheral neuropathy
	Link to chemotherapy-induced peripheral neuropathy screening tool
Ototoxicity	Ototoxicity may occur with platinum-based therapy; patients should be monitored for signs and symptoms. Platinum compounds should be used with caution in patients with pre-existing conditions or risk factors.
	Ototoxicity may become more severe in patients being treated with other drugs with
	nephrotoxic potential e.g. aminoglycosides.
	An audiometry test should be performed if symptoms develop. Read more about ototoxicity - tinnitus and hearing loss
Nutrition risk HIGH	Consider a dietitian review in week 1, with weekly reviews as necessary.
Blood toots	Read more about biosimilar drugs on the Biosimilar Awareness Initiative page
Blood tests	FBC, EUC, LFTs, calcium and magnesium at baseline and prior to each cycle. FBC prior to each treatment. INR as clinically indicated.
Hepatitis B screening and prophylaxis	Routine screening for HBsAg and anti-HBc is NOT usually recommended for patients receiving this treatment.
	Read more about hepatitis B screening and prophylaxis in cancer patients requiring cytotoxic and/or immunosuppressive therapy

Vaccinations	Live vaccines are contraindicated in cancer patients receiving immunosuppressive therapy and/or who have poorly controlled malignant disease. Refer to the recommended schedule of vaccination for immunocompromised patients, as outlined in the Australian Immunisation Handbook. Read more about COVID-19 vaccines and cancer.
Fertility, pregnancy and lactation	Cancer treatment can have harmful effects on fertility and this should be discussed with all patients of reproductive potential prior to commencing treatment. There is a risk of foetal harm in pregnant women. A pregnancy test should be considered prior to initiating treatment in females of reproductive potential if sexually active. It is important that all patients of reproductive potential use effective contraception whilst on therapy and after treatment finishes. Effective contraception methods and adequate contraception timeframe should be discussed with all patients of reproductive potential. Possibility of infant risk should be discussed with breastfeeding patients. Read more about the effect of cancer treatment on fertility

Dose modifications

Evidence for dose modifications is limited, and the recommendations made on eviQ are intended as a guide only. They are generally conservative with an emphasis on safety. Any dose modification should be based on clinical judgement, and the individual patient's situation including but not limited to treatment intent (curative vs palliative), the anti-cancer regimen (single versus combination therapy versus chemotherapy versus immunotherapy), biology of the cancer (site, size, mutations, metastases), other treatment related side effects, additional co-morbidities, performance status and patient preferences. Suggested dose modifications are based on clinical trial findings, product information, published guidelines and reference committee consensus. The dose reduction applies to each individual dose and not to the total number of days or duration of treatment cycle unless stated otherwise. Non-haematological gradings are based on Common Terminology Criteria for Adverse Events (CTCAE) unless otherwise specified. Renal and hepatic dose modifications have been standardised where possible. For more information see dosing considerations & disclaimer.

The dose recommendations in kidney dysfunction (i.e.renal impairment) displayed may not reflect those in the ADDIKD guideline and have been included for historical reference only. Recommendations will be updated once the individual protocol has been evaluated by the reference committee, with this version of the protocol then being archived. Clinicians are expected to refer to the ADDIKD guideline prior to prescribing in kidney dysfunction.

International Consensus Guideline for Anticancer Drug Dosing in Kidney Dysfunction (ADDIKD).

Note: all dose reductions are calculated as a percentage of the starting dose.

Haematological toxicity	
ANC x 10 ⁹ /L (pre-treatment blood test)	
1.0 to less than 1.5	Refer to local institutional guidelines; it is the view of the expert clinicians that treatment should continue if patient is clinically well.
0.5 to less than 1.0	Delay treatment until recovery
less than 0.5	Delay treatment until recovery and consider reducing cisplatin and fluorouracil by 25% for subsequent cycles
Febrile neutropenia	Delay treatment until recovery and consider reducing cisplatin and fluorouracil by 25% for subsequent cycles
Platelets x 10 ⁹ /L (pre-treatment blood test)	
75 to less than 100	The general recommendation is to delay, however if the patient is clinically well it may be appropriate to continue treatment; refer to treating team and/or local institutional guidelines.
50 to less than 75	Delay treatment until recovery
less than 50	Delay treatment until recovery and consider reducing cisplatin and fluorouracil by 25% for

Haematological toxicity	
	subsequent cycles

Renal impairment		
eGFR (CKI-EPI or MDRD) or eCrCl (Cockcroft Gault) (mL/min) *		
greater than or equal to 70	No dose modifications necessary	
50 to less than 70	Reduce cisplatin by 25%	
30 to less than 50	Reduce fluorouracil by 25% and cisplatin by 50%	
less than 30	Reduce fluorouracil by 50% and omit cisplatin or withhold chemotherapy	

 $^{{\}it *Each method has its limitations; refer to \textit{Nephrotoxicity associated with cisplatinfor more information.}}$

Hepatic impairment	
Hepatic dysfunction	
Mild	No dose modifications necessary
Moderate	Reduce fluorouracil by 25%
Severe	Reduce fluorouracil by 50%

Peripheral neuropathy	
Grade 2, Grade 3 or Grade 4	Omit cisplatin

Mucositis and stomatitis		
Grade 2	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1st occurrence: No dose reduction 2nd occurrence: Reduce fluorouracil by 25% 3rd occurrence: Reduce fluorouracil by 50% 4th occurrence: Omit fluorouracil	
Grade 3 or Grade 4	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1st occurrence: Reduce fluorouracil by 50% 2 nd occurrence: Omit fluorouracil	

<u>Diarrhoea</u>	
Grade 2	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1st occurrence: No dose reduction 2nd occurrence: Reduce fluorouracil 25% 3rd occurrence: Reduce fluorouracil by 50% 4th occurrence: Omit fluorouracil
Grade 3 or Grade 4	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1 st occurrence: Reduce fluorouracil by 50% 2 nd occurrence: Omit fluorouracil

Hand foot syndrome (link to Hand foot syndrome (Palmar-plantar erythrodysaesthesia))		
Grade 2	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1 st occurrence: No dose reduction 2 nd occurrence: Reduce fluorouracil 25% 3 rd occurrence: Reduce fluorouracil by 50%	

Hand foot syndrome (link to Hand foot syndrome (Palmar-plantar erythrodysaesthesia))		
	4 th occurrence: Omit fluorouracil	
Grade 3	Delay treatment until toxicity has resolved to Grade 1 or less and reduce the dose for subsequent cycles as follows: 1 st occurrence: Reduce fluorouracil by 50% 2nd occurrence: Omit fluorouracil	

Missed doses of trastuzumab	
By 1 week or less	No dose modification necessary. Give trastuzumab as soon as possible, i.e. do not wait until the next planned cycle.
By more than 1 week	Reload trastuzumab with a dose of 8 mg/kg. Subsequent doses of 6 mg/kg should then be given every 3 weeks, according to the previous cycle.

Interactions

Drug interactions in eviQ protocols are under review and being updated to align with current literature. Further site-wide updates and changes will occur in due course. References & Disclaimer

The drug interactions shown below are not an exhaustive list. For a more comprehensive list and for detailed information on specific drug interactions and clinical management, please refer to the specific drug product information and the following key resources:

- MIMS interactions tab (includes link to a CYP-450 table) (login required)
- Australian Medicines Handbook (AMH) interactions tab (login required)
- Micromedex Drug Interactions (login required)
- Cancer Drug Interactions
- Cytochrome P450 Drug Interactions

Cisplatin		
	Interaction	Clinical management
Nephrotoxic drugs (e.g. aminoglycosides, amphotericin, contrast dye, frusemide, NSAIDs)	Additive nephrotoxicity	Avoid combination or monitor kidney function closely
Ototoxic drugs (e.g. aminoglycosides, frusemide, NSAIDs)	Additive ototoxicity	Avoid combination or perform regular audiometric testing
Neurotoxic drugs (e.g. vincristine, paclitaxel)	Additive neurotoxicity	Monitor closely for neuropathy if combination used
Paclitaxel	Administration schedule may influence the development of myelosuppression	Minimise toxicity by administering paclitaxel first in regimens using the combination
Carbamazepine, phenytoin, valproate	Decreased antiepileptic plasma levels	Monitor antiepileptic serum levels and seizure frequency for efficacy; adjust dosage as appropriate or select alternative antiepileptic (e.g. clonazepam, diazepam, lorazepam)

Fluorouracil		
	Interaction	Clinical management
Folic acid	Increased toxicity of fluorouracil due to stabilisation of its bond to thymidylate synthetase (folic acid is a precursor of folinic acid/leucovorin)	Advise patients not to take folic acid supplements (inc. multivitamins) around the time of receiving treatment with fluorouracil
Metronidazole, tinidazole	Increased toxicity of fluorouracil due to reduced clearance	Avoid combination or monitor for fluorouracil toxicity
Warfarin and other drugs metabolised by CYP2C9 (e.g. warfarin, phenytoin etc.)	Increased effect/toxicity of these drugs due to inhibition of CYP2C9 by fluorouracil resulting in reduced clearance	Avoid combination or monitor for increased effect/toxicity of these drugs (e.g. for bleeding/elevated INR with warfarin, elevated phenytoin serum levels or signs of toxicity such as ataxia, tremor etc.)
Allopurinol	Reduced efficacy of fluorouracil possible due to reduced conversion to the active metabolites	Avoid combination or monitor for reduced fluorouracil efficacy

Trastuzumab		
	Interaction	Clinical management
Cardiotoxic drugs (e.g. anthracyclines cyclophosphamide)	Additive cardiotoxicity	Monitor cardiac function closely in patients who have previously been treated with cumulatively cardiotoxic drugs
Paclitaxel	Increased toxicity of trastuzumab possible due to reduced clearance	Monitor for trastuzumab toxicity (esp. cardiotoxicity)

NK-1 antagonist e.g. aprepitant, fosaprepitant, netupitant		
annegomor org. upropriamily rooupro	Interaction	Clinical management
Dexamethasone	Increased effects/toxicity of dexamethasone due to inhibition of its metabolism via CYP3A4	Reduce dose of antiemetic dexamethasone by approximately 50% when adding a NK-1 antagonist. For protocols that already recommend a NK-1 antagonist, the dose reduction of antiemetic dexamethasone has already been taken into account. If dexamethasone is part of the chemotherapy protocol, dose reduction as per the product information is not routinely recommended in clinical practice and no additional dexamethasone is required for antiemetic cover.
Warfarin	Reduced anticoagulant efficacy of warfarin due to increased clearance (aprepitant induces CYP2C9). *Note interaction only applicable to aprepitant/fosaprepitant	INR should be monitored in the 2 week period, particularly at 7 to 10 days following the administration of aprepitant/ fosaprepitant
Combined oral contraceptive	Reduced contraceptive efficacy due to increased clearance. *Note interaction only applicable to aprepitant/ fosaprepitant	Alternative non-hormonal methods should be used during and for 1 month after stopping aprepitant/ fosaprepitant
CYP3A4 inducers (e.g. carbamazepine, phenytoin, phenobarbitone, rifampicin, St John's wort etc.)	Reduced efficacy of NK-1 antagonist possible due to increased clearance	Avoid combination or monitor for decreased antiemetic effect. Consider using an alternative antiemetic regimen
CYP3A4 inhibitors (e.g. azole antifungals, clarithromycin, erythromycin, grapefruit juice, ritonavir etc.)	Increased toxicity of NK-1 antagonist possible due to reduced clearance	Avoid combination or monitor for increased adverse effects of NK-1 antagonist (e.g. headache, hiccups, constipation)
Drugs metabolised by CYP3A4 (e.g. etoposide, imatinib, irinotecan, midazolam, paclitaxel, vinblastine, vincristine etc.)	Increased effects/toxicity of these drugs possible due to inhibition of CYP3A4 by NK-1 antagonist	Avoid combination or monitor for increased toxicity especially with orally administered drugs

General		
	Interaction	Clinical management
Warfarin	Anti-cancer drugs may alter the anticoagulant effect of warfarin.	Monitor INR regularly and adjust warfarin dosage as appropriate; consider alternative anticoagulant.
Direct oral anticoagulants (DOACs) e.g. apixaban, rivaroxaban, dabigatran	Interaction with both CYP3A4 and P-gp inhibitors /inducers. DOAC and anti-cancer drug levels may both be altered, possibly leading to loss of efficacy or toxicity (i.e. increased bleeding).	Apixaban: avoid concurrent use with strong CYP3A4 and P-gp inhibitors. If treating VTE, avoid use with strong CYP3A4 and P-gp inducers. Rivaroxaban: avoid concurrent use with strong CYP3A4 and P-gp inhibitors. Dabigatran: avoid combination with strong P-gp inducers and inhibitors. If concurrent use is unavoidable, monitor closely for efficacy/toxicity of both drugs.
Digoxin	Anti-cancer drugs can damage the lining of the intestine; affecting the absorption of digoxin.	Monitor digoxin serum levels; adjust digoxin dosage as appropriate.
Antiepileptics	Both altered antiepileptic and anti- cancer drug levels may occur, possibly leading to loss of efficacy or toxicity.	Where concurrent use of an enzyme-inducing antiepileptic cannot be avoided, monitor antiepileptic serum levels for toxicity, as well as seizure frequency for efficacy; adjust dosage as appropriate. Also monitor closely for efficacy of the anti-cancer therapy.
Antiplatelet agents and NSAIDs	Increased risk of bleeding due to treatment related thrombocytopenia.	Avoid or minimise combination. If combination deemed essential, (e.g. low dose aspirin for ischaemic heart disease) monitor for signs of bleeding.
Serotonergic drugs, including selective serotonin reuptake inhibitors (SSRIs e.g. paroxetine) and serotonin noradrenaline reuptake inhibitors (SNRIs e.g. venlafaxine)	Increased risk of serotonin syndrome with concurrent use of 5-HT3 receptor antagonists (e.g. palonosetron, ondansetron, granisetron, tropisetron, dolasetron, etc.)	Avoid combination. If combination is clinically warranted, monitor for signs and symptoms of serotonin syndrome (e.g. confusion, agitation, tachycardia, hyperreflexia). For more information link to TGA Medicines Safety Update
Vaccines	Diminished response to vaccines and increased risk of infection with live vaccines.	Live vaccines (e.g. BCG, MMR, zoster and varicella) are contraindicated in patients on immunosuppressive therapy. Use with caution in patients on non-immunosuppressive therapy. For more information; refer to the recommended schedule of vaccination for cancer patients, as outlined in the Australian Immunisation Handbook

Administration - Cycle 1 to 6

eviQ provides safe and effective instructions on how to administer cancer treatments. However, eviQ does not provide every treatment delivery option, and is unable to provide a comprehensive list of cancer treatment agents and their required IV line giving set/filter. There may be alternative methods of treatment administration, and alternative supportive treatments that are also appropriate. Please refer to the individual

Day 1

Approximate treatment time: 5 hours (initial); 4 hours (subsequent)

Safe handling and waste management

Safe administration

General patient assessment prior to each day of treatment.

Peripheral neuropathy assessment tool

Any toxicity grade 2 or greater may require dose reduction, delay or omission of treatment and review by medical officer before commencing treatment.

Prime IV line(s).

Access TIVAD or CVAD.

Pre treatment medication

Verify antiemetics taken or administer as prescribed.

Verify dexamethasone taken or administer as prescribed.

(2) Treatment - Time out

Trastuzumab

- Trastuzumab is incompatible with glucose solutions. Ensure IV administration sets are flushed with sodium chloride 0.9% pre and post administration.
- Trastuzumab may be administered before or after chemotherapy.

Initial infusion - administer trastuzumab:

- via IV infusion over 90 minutes
- · observe patient for fever and chills or other infusion-related symptoms
- flush with ~50 mL of sodium chloride 0.9%
- · stop infusion at first sign of reaction:
 - if symptoms are mild and resolve when infusion is stopped, consider recommencing infusion after review by medical officer at a slower rate
 - o for severe reactions seek medical assistance immediately and do not restart infusion
- educate the patient about the possibility of delayed infusion-related symptoms.

Subsequent infusions - administer trastuzumab:

- if no previous hypersensitivity reaction administer via IV infusion over 30 minutes
- · observe patient for fever and chills or other infusion-related symptoms
- flush with ~50 mL of sodium chloride 0.9%
- · stop infusion at first sign of reaction:
 - if symptoms are mild and resolve when infusion is stopped, consider recommencing infusion after review by medical officer at a slower rate.
 - o for severe reactions seek medical assistance immediately and do not restart infusion
- educate the patient about the possibility of delayed infusion-related symptoms.

Ochemotherapy - Time out

Cisplatin

Commence prehydration for cisplatin:

- administer 10 mmol magnesium sulphate (MgSO₄) in 1000 mL sodium chloride 0.9% over 60 minutes
- followed by 200 mL of mannitol 20% over 15 minutes
 - o mannitol should be administered via a controlled infusion

- mannitol 10% may be used as per institutional policy; there is much variation in the use of mannitol and although there is no
 conclusive evidence that mannitol should be used, many sites have used it routinely without renal toxicity. The routine use of
 frusemide to increase urine flow is not recommended. Refer to your institutional guidelines and medical orders.
- ensure patient has passed urine prior to cisplatin administration as per institutional policy.

Administer cisplatin (irritant):

- · via IV infusion over 60 minutes
- flush with 100 mL of sodium chloride 0.9%.

Post hydration:

1000 mL sodium chloride 0.9% over 60 minutes.

Fluorouracil continuous infusion (irritant)

Connect pump containing fluorouracil and administer over the correct time for the amount of drug in the pump:

- A safety alert issued for administration of infusional fluorouracil
- · verify the correct rate of infusion via the ambulatory infusion pump
- read more information about the different ambulatory infusion pumps.

Continue safe handling precautions until 7 days after completion of drug(s)

Day 5

Approximate treatment time: 30 minutes

Safe handling and waste management

Disconnection of ambulatory infusion pump/infusor

Verify the ambulatory infusion pump/infusor is complete.

Disconnect the ambulatory infusion pump/infusor as per recommended procedure for type of pump/infusor.

Read more about ambulatory infusion pumps/infusors.

Deaccess TIVAD or CVAD.

Continue safe handling precautions until 7 days after completion of drug(s)

Discharge information

Antiemetics

· Antiemetics as prescribed.

Antidiarrhoeals

· Antidiarrhoeals as prescribed.

Patient information

· Ensure patient receives patient information sheet.

Infusion pumps

- CADD-Legacy® 1 ambulatory infusion pump patient information sheet.
- CADD-Legacy® Plus ambulatory infusion pump patient information sheet.
- CADD® Solis VIP ambulatory infusion pump patient information sheet.
- Elastomeric infusion system patient information sheet.

Administration - Cycle 7 and further cycles

eviQ provides safe and effective instructions on how to administer cancer treatments. However, eviQ does not provide every treatment delivery option, and is unable to provide a comprehensive list of cancer treatment agents and their required IV line giving set/filter. There may be alternative methods of treatment administration, and alternative supportive treatments that are also appropriate. Please refer to the individual

Day 1

Approximate treatment time: 60 minutes

Handling of monoclonal antibodies and waste management

Safe administration

General patient assessment prior to each day of treatment.

Any toxicity grade 2 or greater may require delay of treatment and review by medical officer before commencing treatment.

Prime IV line(s).

Insert IV cannula or access TIVAD or CVAD.

Pre treatment medication

Administer premedication only if previous hypersensitivity reaction.

② Treatment - Time out

Trastuzumab

• Trastuzumab is incompatible with glucose solutions. Ensure IV administration sets are flushed with sodium chloride 0.9% pre and post administration.

Administer trastuzumab:

- if no previous hypersensitivity reaction administer via IV infusion over 30 minutes
- · observe patient for fever and chills or other infusion-related symptoms
- flush with ~ 50 mL of sodium chloride 0.9%
- stop infusion at first sign of reaction:
 - if symptoms are mild and resolve when infusion is stopped, consider recommencing infusion after review by medical officer at a slower rate
 - o for severe reactions seek medical assistance immediately and do not restart infusion
- educate the patient about the possibility of delayed infusion-related symptoms.

If previous hypersensitivity reaction, infuse over 90 minutes following medical review.

Remove IV cannula and/or deaccess TIVAD or CVAD.

Discharge information

Patient information

· Ensure patient receives patient information sheet.

Side effects

The side effects listed below are not a complete list of all possible side effects for this treatment. Side effects are categorised into the approximate onset of presentation and should only be used as a guide.

Immediate (onset hours to days)		
Hypersensitivity reaction	Anaphylaxis and infusion related reactions can occur with this treatment. Read more about hypersensitivity reaction	
Novece and veniting	** *	
Nausea and vomiting	Read more about prevention of treatment induced nausea and vomiting	
Flu-like symptoms		
Taste and smell alteration	Read more about taste and smell changes	
Cardiotoxicity	Coronary artery spasm is a temporary, sudden narrowing of one of the coronary arteries that may present at any time during treatment with fluoropyrimidines. It most commonly manifests as angina.	

Early (onset days to weeks)	
Neutropenia	Abnormally low levels of neutrophils in the blood. This increases the risk of infection. Any fever or suspicion of infection should be investigated immediately and managed aggressively. Read more about immediate management of neutropenic fever
Thrombocytopenia	A reduction in the normal levels of functional platelets, increasing the risk of abnormal bleeding.
rmombocytopema	A reduction in the normal levels of functional platelets, increasing the risk of abhormal bleeding.
	Read more about thrombocytopenia
Oral mucositis	Erythematous and ulcerative lesions of the gastrointestinal tract (GIT). It commonly develops following chemotherapy, radiation therapy to the head, neck or oesophagus, and high dose chemotherapy followed by a blood and marrow transplant (BMT). Read more about oral mucositis
Diarrhoea	Read more about treatment induced diarrhoea
Oesophagitis	Inflammation of the mucosal lining of the oesophagus. It can progress to ulceration, haemorrhage, secondary infection and pain.
Anorexia	Loss of appetite accompanied by decreased food intake. Read more about anorexia
Fatigue	Read more about fatigue
Ocular changes	Symptoms may include eye pain, blurred vision, blepharitis, uveitis, optic neuritis, tear duct stenosis, conjunctivitis, hyperlacrimation, watery or dry eyes and photophobia.
Actinic keratoses flare	Pre-existing actinic keratoses (AKs) can become more inflamed and scaly as a result of immunosuppression. Read more about actinic keratoses flare
Photosensitivity	Increased sensitivity to ultraviolet (UV) light resulting in an exaggerated sunburn-like reaction accompanied by stinging sensations and urticaria.
Skin rash	Anti-cancer drugs can cause a number of changes in the skin with maculo-papular rash the most common type of drug-induced skin reaction. Read more about skin rash
Peripheral neuropathy	Typically symmetrical sensory neuropathy, affecting the fingers and toes, sometimes progressing to the hands and feet. It is associated with several classes of anti-cancer drugs. These include taxanes, platinum-based compounds, vinca alkaloids and some drugs used to treat multiple myeloma. Read more about peripheral neuropathy
Ototoxicity	Tinnitus and hearing loss may occur due to damage in the inner ear. Tinnitus is usually reversible, while hearing loss is generally irreversible. Hearing loss is dose-related, cumulative and may be worse in those with pre-existing hearing problems. Read more about ototoxicity - tinnitus and hearing loss
Nephrotoxicity	Renal dysfunction resulting from damage to the glomeruli, tubules or renal vasculature.
Hypomagnesaemia, hypokalaemia, hypocalcaemia	Abnormally low levels of magnesium, potassium and calcium in the blood.
Palmar-plantar erythrodysaesthesia (PPE) - hand-foot syndrome (HFS)	Bilateral erythema, tenderness, pain, swelling, tingling, numbness, pruritus, dry rash, or moist desquamation and ulceration of the palms and soles. It is also known as hand-foot syndrome (HFS). Symptoms appear to be dose dependent and palms are affected more than soles. Read more about hand-foot syndrome associated with chemotherapy

Late (onset weeks to months)		
Anaemia	Abnormally low levels of red blood cells (RBCs) or haemoglobin in the blood. Read more about anaemia	
Hyperpigmentation	Darkening of an area of skin caused by the overproduction of melanin.	
Nail changes	Hyperpigmentation, paronychia, onycholysis, splinter haemorrhage, pyogenic granuloma formation, subungal haematoma and subungal hyperkeratosis are some of the nail changes associated with anti-cancer drugs. Read more about nail toxicities	
Alopecia - partial	Hair thinning and/or patchy hair loss. Patients can also experience mild to moderate discomfort of the hair follicles, and rarely pain as the hair is falling out. Read more about alopecia and scalp cooling	
Pulmonary toxicity	Pulmonary toxicity may include damage to the lungs, airways, pleura and pulmonary circulation. Read more about pulmonary toxicity associated with anti-cancer drugs	

Delayed (onset months to years)		
Cardiotoxicity	Cardiotoxicity is a well recognised complication of HER-2 directed agents (e.g. trastuzumab, trastuzumab emtansine, pertuzumab). Mechanistically distinct from anthracycline-induced cardiotoxicity, it typically manifests as an asymptomatic decrease in the left ventricular ejection fraction (LVEF) and less commonly as congestive heart failure (CHF). Read more about cardiac toxicity associated with HER-2 targeted agents	

Evidence

The evidence supporting the use of this protocol comes from a phase 3, open-label, randomised controlled trial (ToGA) in which the clinical efficacy and safety of trastuzumab was investigated in combination with chemotherapy for first-line treatment of HER-2 positive advanced gastric or gastro-oesophageal junction cancer.¹

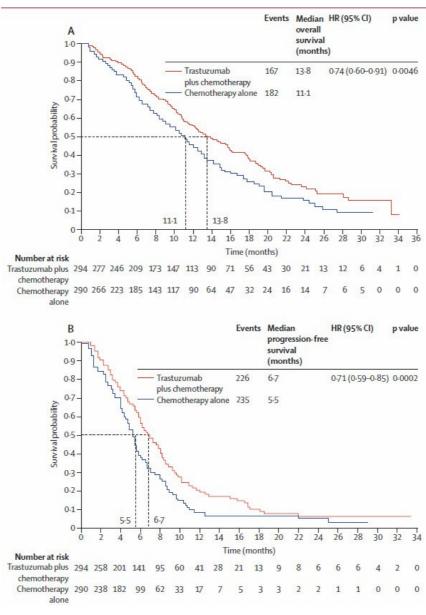
Patients were randomised to receive either trastuzumab plus chemotherapy (T+C) or chemotherapy alone (C). Chemotherapy consisted of either capecitabine $1000 \text{ mg/m}^2 \text{ BD D1-14}$ plus cisplatin $80 \text{ mg/m}^2 \text{ D1}$ every 3 weeks or infusional fluorouracil $800 \text{ mg/m}^2 \text{ D1-5}$ plus cisplatin $80 \text{ mg/m}^2 \text{ D1}$ every 3 weeks. Trastuzumab was given as 8 mg/kg for the first cycle followed by 6 mg/kg every 3 weeks. The chemotherapy regimen was chosen between 5-FU/cisplatin and capecitabine/cisplatin at the investigator's discretion and could be determined on an individual patient basis.

Between September 2005 and December 2008 584 patients were randomised and received at least one dose of study treatment – 294 in the C+T group, 290 in the C group alone. The primary endpoint was overall survival, secondary endpoints progression-free survival, time to progression, overall tumour response and safety.¹

Efficacy

At a median follow up of 17.1 to 18.6 months, median overall survival was 13.8 months in the C+T group and 11.1 months in the C group, HR 0.74. Overall tumour response rate, time to progression, and duration of response were significantly improved in the T+C group vs C alone. A post-hoc subgroup analysis of patients with high (IHC 2+ and FISH + or IHC 3+) levels of HER2 expression showed a median survival of 16.0 months in the T+C group vs 11.8 months in the C group alone, with a HR of 0.65.¹

(A) Median overall survival and (B) progression-free survival in the primary analysis population¹



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Toxicity

The toxicities in the two arms were comparable, except that a higher number of trastuzumab-treated patients had grade 3 or 4 diarrhoea (9% versus 4%) and an asymptomatic decrease in left ventricular ejection fraction (LVEF, 5% versus 1%). Only one patient developed grade 3 to 4 cardiac failure (versus two in the control group) and there were no differences in overall cardiac adverse events (6% in both groups). The other most common grade 3 or 4 adverse effects reported were nausea (7% in both groups), neutropenia (27% in C+T vs 30% in C), and anaemia (12% in C+T, 10% in C). Treatment-related mortality was 3% (10 deaths) in C+T group and 1% (3 deaths) in C group.¹

Toxicity¹

		Trastuzumab plus chemotherapy (n=294)		Chemotherapy alone (n=2	
	All grades	Grade 3 or 4	All grades	Grade 3 or 4	
Any adverse event	292 (99%)	201 (68%)	284 (98%)	198 (68%)	
Gastrointestinal disorders					
Nausea	197 (67%)	22 (7%)	184 (63%)	21 (7%)	
Vomiting	147 (50%)	18 (6%)	134 (46%)	22 (8%)	
Diarrhoea	109 (37%)	27 (9%)	80 (28%)	11 (4%)	
Constipation	75 (26%)	2 (1%)	93 (32%)	5 (2%)	
Stomatitis	72 (24%)	2 (1%)	43 (15%)	6 (2%)	
Abdominal pain	66 (22%)	7 (2%)	56 (19%)	5 (2%)	
Dysphagia	19 (6%)	7 (2%)	10 (3%)	1 (<1%)	
Blood and lymphatic system diso	rders				
Neutropenia	157 (53%)	79 (27%)	165 (57%)	88 (30%)	
Anaemia	81 (28%)	36 (12%)	61 (21%)	30 (10%)	
Thrombocytopenia	47 (16%)	14 (5%)	33 (11%)	8 (3%)	
Febrile neutropenia	15 (5%)	15 (5%)	8 (3%)	8 (3%)	
General, metabolic, and other dis	orders				
Anorexia	135 (46%)	19 (6%)	133 (46%)	18 (6%)	
Fatigue	102 (35%)	12 (4%)	82 (28%)	7 (2%)	
Hand-foot syndrome	75 (26%)	4 (1%)	64 (22%)	5 (2%)	
Weight decreased	69 (23%)	6 (2%)	40 (14%)	7 (2%)	
Asthenia	55 (19%)	14 (5%)	53 (18%)	10 (3%)	
Pyrexia	54 (18%)	3 (1%)	36 (12%)	0	
Renal impairment	47 (16%)	2 (1%)	39 (13%)	3 (1%)	
Mucosal inflammation	37 (13%)	6 (2%)	18 (6%)	2 (1%)	
Nasopharyngitis	37 (13%)	0	17 (6%)	0	
Chills	23 (8%)	1 (<1%)	0	0	
Hypokalaemia	22 (7%)	13 (4%)	13 (4%)	7 (2%)	
Dehydration	18 (6%)	7 (2%)	16 (6%)	5 (2%)	
Dyspnoea	9 (3%)	1 (<1%)	16 (6%)	5 (2%)	

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References

1 Bang, Y. J., E. Van Cutsem, A. Feyereislova, et al. 2010. "Trastuzumab in combination with chemotherapy versus chemotherapy alone for treatment of HER2-positive advanced gastric or gastro-oesophageal junction cancer (ToGA): a phase 3, open-label, randomised controlled trial." Lancet 376(9742):687-697.

History

Version 7

Date	Summary of changes
16/11/2021	Pulmonary toxicity added to side effects. Version number changed to V.7.

Version 6

Date	Summary of changes
21/04/2021	Administration section updated to include cycle 7 and further cycles information. Version number changed to V.6.

Version 5

Date	Summary of changes

Protocol discussed at the Medical Oncology Reference Committee meeting on 23/10/2020. The following changes have been made across all gastric and gastroesophageal metastatic chemotherapy and trastuzumab protocols:

- Treatment schedule Trastuzumab before chemotherapy.
- Administration Trastuzumab before chemotherapy.
- Patient Information (Your treatment) Trastuzumab before chemotherapy.

ID 3805 Gastric and gastroesophageal metastatic FOLFOX6 (modified) and trastuzumab and ID 3896 Gastric and gastroesophageal metastatic CAPOX (XELOX) (capecitabine and oxaliplatin) and trastuzumab added as related pages.

Version number increased to V.5. Next due for review in 2 years.

Version 4

25/11/2020

Date	Summary of changes
04/05/2020	Treatment schedule cycles changed to 'continuous until disease progression or unacceptable toxicity'. Cycle 7 and further cycles added to treatment schedule. Patient information your treatment section updated. Biosimilar trastuzumab added to clinical information. Day 1 approximate treatment time changed to 5 hours (initial), 4 hours (subsequent). Version number changed to V.4.
25/09/2020	Protocol reviewed electronically by the Medical Oncology Reference committee. Nil changes. Next review in 2 years.

Version 3

Date	Summary of changes
25/11/2011	New protocol taken to Medical Oncology Reference Committee meeting.
17/01/2012	Approved and published on eviQ.
10/07/2013	Dose modifications for cisplatin updated.
13/09/2013	Protocol reviewed by reference committee via email. No changes and next review in 2 years.
16/12/2015	Protocol reviewed electronically by Medical Oncology Reference committee. No changes and next review in 2 years.
16/10/2016	Patient information sheet updated to include more fluorouracil toxicity symptom warnings.
09/11/2016	The following changes made post Medical Oncology Reference Committee meeting held on 21 October 2016. Title changed from advanced to metastatic. Link to AGITG and ANZCTR added.
24/03/2017	Consensus of the Medical Oncology Reference Committee (via email discussion) to remove observation time frames from all trastuzumab protocols and replace with the statement "Observe patient for fever and chills or other infusion-related symptoms" as per current trastuzumab product information. Individual institutions may still implement/maintain local policies on monitoring time frames if they choose to do so.
31/05/2017	Transferred to new eviQ website. Version number changed to V.2. Antiemetic change: Netupitant/palonosetron combination has replaced aprepitant and a 5HT ₃ receptor antagonist in combination with dexamethasone for all highly emetogenic regimens. Hepatitis B screening changed to NOT recommended
16/02/2018	Protocol reviewed electronically by Medical Oncology Reference Committee. Fluoropyrimidine overdose or overexposure warning added. Review in 2 years
15/03/2018	Patient information sheet: title amended to "Stomach and gastroesophageal junction cancer"
10/05/2018	Haematological dose modifications updated as per consensus of the expert clinician group. DPD enzyme deficiency wording in clinical information updated. Fluoropyrimidine safety alert wording in clinical information updated. Version number changed to V.3.

The information contained in this protocol is based on the highest level of available evidence and consensus of the eviQ reference committee regarding their views of currently accepted approaches to treatment. Any clinician (medical oncologist, haematologist, radiation oncologist, medical physicist, radiation therapist, pharmacist or nurse) seeking to apply or consult this protocol is expected to use independent clinical judgement in the context of individual clinical circumstances to determine any patient's care or treatment. While eviQ endeavours to link to reliable sources that provide accurate information, eviQ and the Cancer Institute NSW do not endorse or accept responsibility for the accuracy, currency, reliability or correctness of the content of linked external information sources. Use is subject to eviQ's disclaimer available at www.eviQ.org.au

First approved: 17 January 2012
Last reviewed: 23 October 2020
Review due: 31 December 2022

The currency of this information is guaranteed only up until the date of printing, for any updates please check: https://www.eviq.org.au/p/1217

11 Jul 2023

Patient information - Stomach or gastroesophageal junction cancer metastatic - Cisplatin, fluorouracil and trastuzumab



Patient's name:

Your treatment

The treatment schedule below explains how the drugs for this treatment are given.

Cisplatin, fluorouracil, trastuzumab

This treatment cycle is repeated every 21 days. You will have 6 cycles of cisplatin, fluorouracil and trastuzumab. After 6 cycles you will stop cisplatin and fluorouracil and continue to receive trastuzumab once every 21 days. Your doctor will advise you of the number of treatments you will have.

Day	Treatment	How it is given	How long it takes
1	Trastuzumab (tras-TOOZ-ue-mab) Cisplatin (siss-PLAT-in)	By a drip into a vein	About 5 hours for the first treatment. If no reactions, subsequent treatment may be given over a shorter amount of time e.g. 4 hours
1 to 4	Fluorouracil (flure-oh-YOOR-a-sill)	By a pump slowly into a vein	Continuous for 4 days by pump at home
5	Disconnect pump		About 30 minutes

When to get help

Anticancer drugs (drugs used to treat cancer) can sometimes cause serious problems. It is important to get medical help immediately if you become unwell.

IMMEDIATELY go to your nearest hospital Emergency Department, or contact your doctor or nurse if you have any of the following at any time:	Emergency contact details Ask your doctor or nurse from your treating team who to contact if you have a problem
 a temperature of 38°C or higher chills, sweats, shivers or shakes shortness of breath uncontrolled vomiting or diarrhoea pain, tingling or discomfort in your chest or arms leaking from your pump you become unwell. 	Daytime: Night/weekend: Other instructions:

During your treatment immediately tell the doctor or nurse looking after you if you get any of the following problems:

- leaking from the area where the drugs are being given
- pain, stinging, swelling or redness in the area where the drugs are being given or at any injection sites
- a skin rash, itching, feeling short of breath, wheezing, fever, shivers, or feeling dizzy or unwell in any way (allergic reaction).

Other information about your treatment

Changes to your dose or treatment delays

Sometimes a treatment may be started at a lower dose or the dose needs to be changed during treatment. There may also be times when your treatment is delayed. This can happen if your doctor thinks you are likely to have severe side effects, if you get severe side effects, if your blood counts are affected and causing delays in treatment, or if you are finding it hard to cope with the treatment. This is called a dose reduction, dose change or treatment delay. Your doctor will explain if you need any changes or delays to your treatment and the reason why.

Blood tests and monitoring

Anti-cancer drugs can reduce the number of blood cells in your body. You will need to have regular blood tests to check that your blood cell count has returned to normal. If your blood count is low your treatment may be delayed until it has returned to normal. Your doctor or nurse will tell you when to have these blood tests. Tell your doctor if you are on an anticoagulant (medication used to treat or prevent blood clots) e.g. warfarin. You may need to have additional blood tests.

Pumps and central venous access devices (CVADs)

This treatment involves having chemotherapy through a pump. To have this, you will also need a central venous access device (CVAD). Your doctor or nurse will explain this to you. For more information see the eviQ patient information sheets on pumps and CVADs. At home you will need to look at your pump 3 to 4 times a day to check it is working. Your nurse will teach you how to do this.

Other medications given during this treatment

- Anti-sickness (anti-nausea) medication: you may be given some anti-sickness medication. Make sure you take this medication as your doctor or nurse tells you, even if you don't feel sick. This can help to prevent the sickness starting.
- Antidiarrhoeals: you may be given some medication to treat diarrhoea. Your doctor or nurse will tell you how and when to take your antidiarrhoeal medication.

Side effects

Cancer treatments can cause damage to normal cells in your body, which can cause side effects. Everyone gets different side effects, and some people will have more problems than others.

The table below shows some of the side effects you may get with this treatment. You are unlikely to get all of those listed and you may also get some side effects that have not been listed.

Tell your doctor or nurse about any side effects that worry you. Follow the instructions below and those given to you by your doctor or nurse.

Immediate (onset hours to days) • Allergic reactions are uncommon but can be life threatening. Allergic reaction • If you feel unwell during the infusion or shortly after it, or: o get a fever, shivers or shakes feel dizzy, faint, confused or anxious o start wheezing or have difficulty breathing have a rash, itch or redness of the face While you are in hospital: Tell your doctor or nurse immediately. After you leave: Contact your doctor or nurse immediately, or go to the nearest hospital **Emergency Department.** • You may feel sick (nausea) or be sick (vomit). Nausea and vomiting • Take your anti-sickness medication as directed even if you don't feel sick. • Drink plenty of fluids (unless you are fluid restricted). · Eat small meals more frequently. • Try food that does not require much preparation. • Try bland foods like dry biscuits or toast. • Gentle exercise may help with nausea. · Ask your doctor or nurse for eviQ patient information - Nausea and vomiting during cancer treatment. • Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you have uncontrolled vomiting or feel dizzy or light-headed. You may get: Flu-like symptoms a fever o chills or sweats o muscle and joint pain a cough headaches. Tell your doctor or nurse if you get any of the symptoms listed above. • Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you have a temperature of 38°C or higher. • You may find that food loses its taste or tastes different. Taste and smell changes • These changes are likely to go away with time. • Do your mouth care regularly. • Chew on sugar-free gum or eat sugar-free mints. • Add flavour to your food with sauces and herbs. · Ask your doctor or nurse for eviQ patient information - Taste and smell changes during cancer treatment.

Early (onset days to weeks)

Infection risk (neutropenia)

- This treatment lowers the amount of white blood cells in your body. The type of white blood
 cells that help to fight infection are called neutrophils. Having low level of neutrophils is
 called neutropenia. If you have neutropenia, you are at greater risk of getting an infection. It
 also means that your body can't fight infections as well as usual. This is a serious side effect,
 and can be life threatening.
- · Wash your hands often.
- Keep a thermometer at home and take your temperature regularly, and if you feel unwell.
- Do your mouth care regularly.
- Inspect your central line site (if you have one) daily for any redness, pus or swelling.
- · Limit contact with people who are sick.
- Learn how to recognise the signs of infection.
- Ask your doctor or nurse for eviQ patient information Infection during cancer treatment.
- Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you get any of the following signs or symptoms:
 - a temperature of 38°C or higher
 - o chills, shivers, sweats or shakes
 - o a sore throat or cough
 - uncontrolled diarrhoea
 - shortness of breath
 - a fast heartbeat
 - become unwell even without a temperature.

Low platelets (thrombocytopenia)

- This treatment lowers the amount of platelets in your blood. Platelets help your blood to clot. When they are low, you are at an increased risk of bleeding and bruising.
- Try not to bruise or cut yourself.
- · Avoid contact sport or vigorous exercise.
- Clear your nose by blowing gently.
- · Avoid constipation.
- Brush your teeth with a soft toothbrush.
- Don't take aspirin, ibuprofen or other similar anti-inflammatory medications unless your doctor tells you to.
- Tell your doctor or nurse if you have any bruising or bleeding.
- Tell your doctor or nurse immediately, or go to your nearest hospital Emergency Department if you have any uncontrolled bleeding.

Mouth pain and soreness (mucositis)

- You may have:
 - o bleeding gums
 - mouth ulcers
 - o a white coating on your tongue
 - o pain in the mouth or throat
 - difficulty eating or swallowing.
- Avoid spicy, acidic or crunchy foods and very hot or cold food and drinks.
- Try bland and soft foods.
- Brush your teeth gently with a soft toothbrush after each meal and at bedtime. If you normally floss continue to do so.
- Rinse your mouth after you eat and brush your teeth, using either:
 - 1/4 teaspoon of salt in 1 cup of warm water, or
 - 1/4 teaspoon of bicarbonate of soda in 1 cup of warm water
- Ask your doctor or nurse for eviQ patient information Mouth problems during cancer treatment.
- Tell your doctor or nurse if you get any of the symptoms listed above.

• You may get bowel motions (stools, poo) that are more frequent or more liquid. Diarrhoea • You may also get bloating, cramping or pain. • Take your antidiarrhoeal medication as directed by your doctor. Drink plenty of fluids (unless you are fluid restricted). · Eat and drink small amounts more often. • Avoid spicy foods, dairy products, high fibre foods, and coffee. Ask your doctor or nurse for eviQ patient information - Diarrhoea during cancer treatment. Tell your doctor or nurse immediately, or go to your nearest hospital Emergency Department if your diarrhoea is not controlled, you have 4 or more loose bowel motions per day, and if you feel dizzy or light-headed. • You may get heartburn or have difficult or painful swallowing. **Oesophagus inflammation** • Eat small meals that are high in protein and calories. (oesophagitis) • Avoid eating acidic, hot, salty or spicy foods, and drinking alcohol. Sit upright when eating. Ask to speak with a dietitian if you are having trouble eating. . Tell your doctor or nurse as soon as possible if you have the any of the symptoms listed above and they are suddenly getting worse. You may not feel like eating. Appetite loss (anorexia) • Try to avoid drinking fluids at meal times. • Try to eat small meals or snacks regularly throughout the day. • Try to eat food that is high in protein and calories. If you are worried about how much food you can eat, or if you are losing weight, ask to speak to a dietitian. • You may feel very tired, have no energy, sleep a lot, and not be able to do normal activities or Tiredness and lack of energy things you enjoy. (fatigue) • Do not drive or operate machinery if you are feeling tired. • Nap for short periods (only 1 hour at a time) • Prioritise your tasks to ensure the best use of your energy. Eat a well balanced diet and drink plenty of fluids (unless you are fluid restricted). • Try some gentle exercise daily. • Allow your friends and family to help. • Tell your doctor or nurse if you get any of the symptoms listed above. You may get: Eye problems eye pain red, sore or swollen eyes blurred vision watery or gritty eyes o changes in your eyesight o sensitivity to sunlight. • Protect your eyes from the weather (sun and wind) by wearing sunglasses, especially if you have lost your eyelashes. . Tell your doctor or nurse if you get any of the symptoms listed above. Eye drops may help with your symptoms. Your skin may become dry, and you may notice changes to areas of your skin that have been Skin changes exposed to the sun. Keep your skin moisturised with a cream such as sorbolene or aqueous cream. · Avoid direct sunlight. Protect your skin from the sun by wearing a wide-brimmed hat, sun-protective clothing, sunglasses and sunscreen of SPF 50 or higher. Tell your doctor or nurse if you notice any skin changes.

• After being out in the sun you may develop a rash like a bad sunburn. Skin that is more sensitive to • Your skin may become red, swollen and blistered. the sun (photosensitivity) · Avoid direct sunlight. Protect your skin from the sun by wearing sun-protective clothing, a wide-brimmed hat, sunglasses and a sunscreen of SPF 50 or higher. • Tell your doctor or nurse if you get any of the symptoms listed above. • You may get a red, bumpy rash and dry, itchy skin. Skin rash · Moisturise your skin with a gentle non-perfumed moisturising cream like sorbolene or aqueous cream. Do not scratch your skin. · Protect your skin from the sun by wearing sun-protective clothing, a wide-brimmed hat, sunglasses and sunscreen of SPF 50 or higher. • Talk to your doctor or nurse about other ways to manage your skin rash. • You may notice a change in the sensations in your hands and feet, including: Nerve damage (peripheral tingling or pins and needles neuropathy) o numbness or loss of feeling o pain. You may find it difficult to do everyday activities, such as doing up buttons or picking up small objects. • Test water temperature with your elbow when bathing to avoid burns. • Use rubber gloves, pot holders and oven mitts in the kitchen. • Wear rubber shoes or boots when working in the garden or garage. · Keep rooms well lit and uncluttered. Ask your doctor or nurse for eviQ patient information – Nerve problems during cancer treatment. • Tell your doctor or nurse if you get any of the symptoms listed above. · You may get ringing in your ears or loss of hearing. Hearing changes • You may have your hearing tested before and during your treatment. (ototoxicity) . Tell your doctor or nurse as soon as possible if you notice any changes to your hearing. • This treatment can cause changes to how your kidneys work. Kidney damage · You will have blood tests to make sure your kidneys are working properly. You may need to drink more fluids while you are having treatment. Your doctor or nurse will tell you if you need to do this. Tell your doctor or nurse as soon as possible if you notice that your urine changes colour or you don't need to empty your bladder as often. • This may be found from your routine blood tests and treated by your doctor. Low blood magnesium, • If it is severe you may get: potassium and calcium o muscle cramps or twitches levels (hypomagnesaemia, o numbness or tingling in your fingers, toes or around your mouth hypokalaemia, constipation hypocalcaemia) an irregular heartbeat sleepy, drowsy or confused . Tell your doctor or nurse as soon as possible if you get any of the signs or symptoms listed above

Hand-foot syndrome (palmar-plantar erythrodysaesthesia)

- The palms of your hands and soles of your feet may become:
 - red and hot
 - swollen
 - o painful and tender
 - o blistered.
- The skin in the area may also peel.
- Moisturise your hands and feet daily with sorbolene or aqueous cream.
- Keep your hands and feet clean and dry.
- Avoid hot water, instead use lukewarm water to bathe.
- Avoid direct sunlight.
- Avoid unnecessary walking, jogging or exercise.
- Wear cotton socks and avoid tight-fitting shoes.
- Tell your doctor or nurse as soon as possible if you notice any skin changes on your hands or feet.

Late (onset weeks to months) • You may feel dizzy, light-headed, tired and appear more pale than usual. Low red blood cells • Tell your doctor or nurse if you have any of these signs or symptoms. You might need a (anaemia) blood transfusion. Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you have any chest pain, trouble breathing, or feel like your heart is racing. • You may have darkening of your skin, especially in areas that are exposed to the sun. Skin colour changes • You may also notice darkening of your tongue, gums and over your finger joints. • These skin changes may fade over time. Protect your skin from the sun by wearing sun-protective clothing, a wide-brimmed hat, sunglasses and a sunscreen of SPF 50 or higher. · Your nails may: **Nail changes** grow more slowly become darker develop ridges or white lines o become brittle and flaky • In some cases, you may lose your nails completely. Keep your nails clean and short. • Avoid things like biting your fingernails, getting a manicure, pedicure or false nails. • Wear gloves when you wash the dishes, work in the garden, or clean the house. • Your hair may become dry and may break easily. Hair thinning · You may lose some of your hair. • Use a gentle shampoo and a soft hairbrush. • Take care with hair products like hairspray, hair dye, bleaches and perms. • Protect your scalp from the cold with a hat or scarf. • Protect your scalp from the sun with a hat and sunscreen of SPF 50 or higher. • Ask your doctor or nurse about the Look Good Feel Better program (www.lgfb.org.au) · Lung problems are rare, but can be serious. They may occur throughout treatment or after Lung problems the completion of treatment. You may get: shortness of breath fever · dry cough wheezing fast heartbeat o chest pain. • Your doctor will monitor how well your lungs are working during your treatment. Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you have chest pain or become short of breath.

Delayed (onset months to years)

Heart problems

- · You may get:
 - chest pain or tightness
 - shortness of breath
 - swelling of your ankles
 - o an abnormal heartbeat.
- · Heart problems can occur months to years after treatment.
- Tell your doctor if you have a history of heart problems or high blood pressure.
- Before or during treatment, you may be asked to have a test to see how well your heart is working.
- Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you get any of the symptoms listed above.

General advice for people having cancer treatment

Chemotherapy safety

- Learn how to keep you and your family safe while you are having anticancer drugs.
- · See our patient information sheet Chemotherapy safety at home.

Blood clot risk

- Cancer and anticancer drugs can increase the risk of a blood clot (thrombosis).
- Tell your doctor if you have a family history of blood clots.
- A blood clot can cause pain, redness, swelling in your arms or legs, shortness of breath or chest pain.
- If you have any of these symptoms go to your nearest hospital Emergency Department.

Medications and vaccinations

- Before you start treatment, tell your doctor about any medications you are taking, including vitamins or herbal supplements.
- Don't stop or start any medications during treatment without talking to your doctor and pharmacist first.
- Paracetamol is safe to take if you have a headache or other mild aches and pains. It is recommended that you avoid taking aspirin, ibuprofen and other anti-inflammatory type medications for pain while you are having treatment. However, if these medications have been prescribed by your doctor, do not stop taking them without speaking with your doctor.
- Vaccinations such as flu and tetanus vaccines are safe to receive while having treatment. Do not have any live vaccines during your treatment or for 6 months after it finishes. If you are unsure, check with your doctor before you have any vaccinations.
- People you live with should be fully vaccinated, including having live vaccines according to the current vaccination schedule. Extra
 care needs to be taken with hand washing and careful disposal of soiled nappies for infants who have recently received the
 rotavirus vaccine.

Other medical and dental treatment

- If you go to hospital or any other medical appointment (including dental appointments), always tell the person treating you that you are receiving anticancer drugs.
- · Before you have any dental treatment, talk to your doctor.

Diet

- · While you are receiving this treatment it is important that you try to maintain a healthy diet.
- Speak to your doctor or nurse about whether drinking alcohol is safe with your treatment.
- If you have any concerns about recent weight loss or weight gain or questions about your diet, ask to speak to a dietitian.

Fertility

- Some cancer treatments can reduce your fertility. This can make it difficult or impossible to get pregnant or father a child.
- Talk to your doctor or nurse before you start any treatment. Depending on your situation there may be fertility sparing options available to you and/or your partner, discuss these with your doctor or nurse.

Pregnancy and breastfeeding

- Some cancer treatments can be dangerous to unborn babies. Talk to your doctor or nurse if you think there is any chance that you could be pregnant.
- Do not try to get pregnant or father a child during this treatment. Contraception should be used during treatment and after stopping treatment. Ask your doctor or nurse about what type of contraception you should use.
- If you are planning pregnancy/fatherhood after completing this treatment, talk to your doctor. Some doctors advise waiting between 6 months and 2 years after treatment.
- · Do not breastfeed if you are on this treatment, as anti-cancer medications can also pass into breast milk.

Sex life and sexuality

- The desire to have sex may decrease as a result of this treatment or its side effects.
- Your emotions and the way you feel about yourself may also be affected by this treatment.
- It may help to discuss your concerns with your partner and doctor or nurse.

Risk of developing a second cancer

Some anticancer treatments can increase your chance of developing a second cancer, this is rare. Your doctor will discuss with
you the specific risks of your treatment.

Quitting smoking

- It is never too late to guit smoking. Quitting smoking is one of the best things you can do to help your treatment work better.
- There are many effective tools to improve your chances of guitting.
- Talk to your treating team for more information and referral to a smoking cessation support service.

Staying active

- · Research shows that exercise, no matter how small, has many benefits for people during and after cancer treatment.
- Talk to your doctor before starting an exercise program. Your doctor can advise whether you need a modified exercise program.

For more information about cancer treatment, side effects and side effect management see our Patient and carers section.

Where to get more information

Telephone support

• Call Cancer Council on 13 11 20 for cancer information and support

Stomach and oesophageal cancer information

• Pancare Foundation - pancare.org.au

General cancer information and support

- Australian Rare Cancer (ARC) Portal arcportal.org.au/
- Beyondblue beyondblue.org.au
- Cancer Australia canceraustralia.gov.au
- Cancer Council Australia cancer.org.au
- Cancer Voices Australia cancervoicesaustralia.org
- CanTeen canteen.org.au
- Carers Australia carersaustralia.com.au
- · CHILL Cancer related hair loss scalpcooling.org
- eviQ Cancer Treatments Online eviQ.org.au
- LGBTQI+ People and Cancer cancercouncil.com.au/cancer-information/lgbtqi
- Look Good Feel Better lgfb.org.au
- Patient Information patients.cancer.nsw.gov.au
- Radiation Oncology Targeting Cancer targetingcancer.com.au
- Redkite redkite.org.au
- Return Unwanted Medicines returnmed.com.au
- Staying active during cancer treatment patients.cancer.nsw.gov.au/coping-with-cancer/physical-wellbeing/staying-active

Quit smoking information and support

Quitting smoking is helpful even after you have been diagnosed with cancer. The following resources provide useful information and support to help you quit smoking. Talk to your treating team about any other questions you may have.

- Call Quitline on 13 QUIT (13 78 48)
- iCanQuit iCanQuit.com.au
- Patient Information patients.cancer.nsw.gov.au/coping-with-cancer/physical-wellbeing/quitting-smoking
- Quitnow quitnow.gov.au

Additional notes:			

This document is a guide only and cannot cover every possible situation. The health professionals caring for you should always consider your individual situation when making decisions about your care. Contact your cancer clinic staff or doctor if you have any questions or concerns about your treatment, or you are having problems coping with side effects. While eviQ endeavours to link to reliable sources that provide accurate information, eviQ and the Cancer Institute NSW do not endorse or accept responsibility for the accuracy, currency, reliability or correctness of the content of linked external information sources. Use of this document is subject to eviQ's disclaimer available at www.eviQ.org.au

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