



ID: 4114 v.1 Endorsed Essential Medicine List

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



2022

#### Related pages:

- Breast neoadjuvant AC (DOXOrubicin and CYCLOPHOSPHamide) three weekly followed by DOCEtaxel three weekly overview
- Anti-cancer therapy before breast cancer surgery (neoadjuvant therapy)

#### **Treatment schedule - Overview**

#### Cycle 1 to 6

Drug	Dose	Route	Day
DOXOrubicin	50 mg/m <sup>2</sup>	IV	1
DOCEtaxel	75 mg/m <sup>2</sup>	IV infusion	1
CYCLOPHOSPHamide	500 mg/m <sup>2</sup>	IV infusion	1
Pegfilgrastim	6 mg	Subcut	2

Frequency: 21 days

**Cycles:** 6 (up to 8 cycles for neoadjuvant treatment)

#### Notes:

Clinical trials have used prophylactic ciprofloxacin 500 mg twice daily on days 5 to 14. An update on pegfilgrastim +/-ciprofloxacin for primary prophylaxis with TAC is available.<sup>1</sup>

Drug status: Doxorubicin, docetaxel and cyclophosphamide are on the PBS general schedule

Pegfilgrastim is PBS authority

Cost: ~ \$210 per cycle

#### 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 to 6

Day before chemotherapy			
Dexamethasone	8 mg (PO)	TWICE a day with or after food	
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	8 mg (PO)	60 minutes before chemotherapy	
DOXOrubicin	50 mg/m <sup>2</sup> (IV)	over 5 to 15 minutes	
DOCEtaxel	75 mg/m <sup>2</sup> (IV infusion)	in 250 mL to 500 mL sodium chloride 0.9% over 60 minutes	
CYCLOPHOSPHamide	500 mg/m <sup>2</sup> (IV infusion)	in 500 mL sodium chloride 0.9% over 30 to 60 minutes	
Day 2			
Pegfilgrastim	6 mg (Subcut)	inject subcutaneously at least 24 hours post chemotherapy	
Dexamethasone	8 mg (PO)	ONCE a day (or in divided doses) with or after food.	
Day 3 and 4			
Dexamethasone	8 mg (PO)	ONCE a day (or in divided doses) with or after food.  Note: dexamethasone doses on Day 3 and 4 may not be required and may be reduced or omitted at the clinicians discretion*	

<sup>\*</sup> Link to ID 7 Prevention of anti-cancer therapy induced nausea and vomiting

Frequency: 21 days

**Cycles:** 6 (up to 8 cycles for neoadjuvant treatment)

#### Indications and patient population - Adjuvant

• Node-positive or high risk node-negative, HER-2 negative breast cancer.

#### Indications and patient population - Neoadjuvant

• Neoadjuvant treatment for HER-2 negative breast cancer.

#### **Clinical information**

Venous access required IV cannula (IVC) or central venous access device (CVAD) is required to administer this treatment.

Read more about central venous access device line selection

Hypersensitivity/infusion related reaction	High risk with docetaxel
Premedication	The product information states that premedication is required for this treatment.  Please refer to the treatment schedule for the suggested premedication regimen.  This may be substituted to reflect institutional policy.  Read more about premedication for prophylaxis of taxane hypersensitivity reactions
Emetogenicity HIGH	Suggested default antiemetics have been added to the treatment schedule, and may be substituted to reflect institutional policy.  A steroid has been included both as an antiemetic and premedication for hypersensitivity in this
	protocol.
	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
Cumulative lifetime dose of anthracyclines	Cumulative doses should take into account all previous anthracyclines received during a patient's lifetime (i.e. daunorubicin, doxorubicin, epirubicin, idarubicin and mitoxantrone).
	Criteria for reducing the total anthracycline cumulative lifetime dose include:  • patient is elderly
	<ul> <li>prior mediastinal radiation</li> <li>hypertensive cardiomegaly</li> </ul>
	<ul> <li>concurrent therapy with high dose cyclophosphamide and some other cytotoxic drugs (e.g. bleomycin, dacarbazine, dactinomycin, etoposide, melphalan, mitomycin and vincristine).</li> </ul>
	Baseline clinical assessments include echocardiogram (ECHO) or gated heart pool scan (GHPS) and electrocardiogram (ECG) evaluation.
	Patients with normal baseline cardiac function (left ventricular ejection fraction (LVEF) > 50%) and low risk patients require LVEF monitoring when greater than 70% of the anthracycline threshold is reached or if the patient displays symptoms of cardiac impairment. Post-treatment cardiac monitoring is recommended for patients who have received high levels of total cumulative doses of anthracyclines at the clinician's discretion.
	Read more about cardiac toxicity associated with anthracyclines
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
Biosimilar drug	Read more about biosimilar drugs on the Biosimilar Awareness Initiative page
Growth factor support	G-CSF (short or long-acting) is available on the PBS for chemotherapy induced neutropenia depending on clinical indication and/or febrile neutropenia risk.  Access the PBS website
Blood tests	FBC, EUC and LFTs at baseline and prior to each cycle.
Hepatitis B screening and	Routine screening for HBsAg and anti-HBc is recommended prior to initiation of treatment.
prophylaxis	Prophylaxis should be determined according to individual institutional policy.
	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.

#### 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.

Read more about the effect of cancer treatment on fertility

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 <sup>9</sup> /L (pre-treatment blood test)		
0.5 to less than 1.0	Delay treatment until recovery	
less than 0.5	Delay treatment until recovery and consider reducing docetaxel, doxorubicin and cyclophosphamide by 25% for subsequent cycles	
Febrile neutropenia	Delay treatment until recovery and consider reducing docetaxel, doxorubicin and cyclophosphamide by 25% for subsequent cycles	
Platelets x 10 <sup>9</sup> /L (pre-treatm	ent blood test)	
75 to less than 100	Refer to local institutional guidelines; it is the view of the expert clinicians that treatment should continue if patient is clinically well	
50 to less than 75	Delay treatment until recovery	
less than 50	Delay treatment until recovery and consider reducing docetaxel, doxorubicin and cyclophosphamide by 25% for subsequent cycles	

Renal impairment	
Creatinine clearance (mL/min)	
30 to 50	Reduce cyclophosphamide by 25%
less than 30	Reduce cyclophosphamide by 50%

Hepatic impairment	
Hepatic dysfunction	
Minimal	Reduce docetaxel by 25%
Mild	Reduce doxorubicin by 25% and docetaxel by 50%
Moderate	Reduce doxorubicin by 50% and omit docetaxel
Severe	Omit doxorubicin and docetaxel

Peripheral neuropathy	
Grade 2, which is present at the start of the next cycle	Reduce docetaxel by 25%; If persistent, reduce docetaxel by 50%
Grade 3 or Grade 4	Omit docetaxel

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 docetaxel, doxorubicin and cyclophosphamide by 25%  3rd occurrence: Reduce docetaxel, doxorubicin and cyclophosphamide by 50%  4th occurrence: Withhold chemotherapy	
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 docetaxel, doxorubicin and cyclophosphamide by 50%  2nd occurrence: Withhold chemotherapy	

<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 docetaxel, doxorubicin and cyclophosphamide by 25%  3rd occurrence: Reduce docetaxel, doxorubicin and cyclophosphamide by 50%  4th occurrence: Withhold chemotherapy
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 docetaxel, doxorubicin and cyclophosphamide by 50%  2nd occurrence: Withhold chemotherapy

#### **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

Cyclophosphamide		
	Interaction	Clinical management
CYP3A4 inducers (e.g. carbamazepine, phenytoin, phenobarbitone, rifampicin, St John's wort etc.)	Increased toxicity of cyclophosphamide possible due to increased conversion to active (and inactive) metabolites	Avoid combination or monitor for cyclophosphamide toxicity
CYP3A4 inhibitors (e.g. aprepitant, azole antifungals, clarithromycin, erythromycin, grapefruit juice, ritonavir etc.)	Reduced efficacy of cyclophosphamide possible due to decreased conversion to active (and inactive) metabolites	Avoid combination or monitor for decreased clinical response to cyclophosphamide
Amiodarone	Possible additive pulmonary toxicity with high-dose cyclophosphamide (i.e. doses used prior to stem cell transplant; 60 mg/kg daily or 120 to 270 mg/kg over a few days)	Avoid combination or monitor closely for pulmonary toxicity
Allopurinol, hydrochlorothiazide, indapamide	Delayed effect. Increased risk of bone marrow depression; probably due to reduced clearance of active metabolites of cyclophosphamide	Avoid combination, consider alternative antihypertensive therapy or monitor for myelosuppression
Ciclosporin	Reduced efficacy of ciclosporin due to reduced serum concentration	Monitor ciclosporin levels; adjust dosage as appropriate; monitor response to ciclosporin
Suxamethonium	Prolonged apnoea due to marked and persistent inhibition of cholinesterase by cyclophosphamide	Alert the anaesthetist if a patient has been treated with cyclophosphamide within ten days of planned general anaesthesia

Docetaxel		
	Interaction	Clinical management
CYP3A4 and P-gp inhibitors (e.g. amiodarone, aprepitant, azole-antifungals, ritonavir, lapatinib, nilotinib, sorafenib, macrolides, ciclosporin, grapefruit juice etc.)	Increased toxicity of docetaxel possible due to reduced clearance	Avoid combination or monitor for docetaxel toxicity
CYP3A4 inducers (e.g. carbamazepine, phenytoin, phenobarbitone, rifampicin, St John's wort etc.)	Reduced efficacy of docetaxel possible due to increased clearance	Avoid combination or monitor for decreased clinical response to docetaxel

Doxorubicin		
	Interaction	Clinical management
Cardiotoxic drugs (eg. bevacizumab, calcium channel blockers, propranolol, trastuzumab)	Increased risk of doxorubicin-induced cardiotoxicity	Avoid combination or monitor closely for cardiotoxicity
Cyclophosphamide	Sensitises the heart to the cardiotoxic effects of doxorubicin; also, doxorubicin may exacerbate cyclophosphamide induced cystitis	Monitor closely for cardiotoxicity and ensure adequate prophylaxis for haemorrhagic cystitis when combination is used
Glucosamine	Reduced efficacy of doxorubicin (due to induction of glucose-regulated stress proteins resulting in decreased expression of topoisomerase II <i>in vitro</i> )	The clinical effect of glucosamine taken orally is unknown. Avoid combination or monitor for decreased clinical response to doxorubicin
CYP2D6 inhibitors (e.g. SSRIs (esp. paroxetine), perhexiline, cinacalcet, doxepin, flecainide, quinine, terbinafine)	Increased toxicity of doxorubicin possible due to reduced clearance	Monitor for doxorubicin toxicity
CYP3A4 inhibitors (e.g. aprepitant, azole antifungals, clarithromycin, erythromycin, grapefruit juice, ritonavir etc.)	Increased toxicity of doxorubicin possible due to reduced clearance	Monitor for doxorubicin toxicity
CYP3A4 inducers (e.g. carbamazepine, phenytoin, phenobarbitone, rifampicin, St John's wort etc.)	Reduced efficacy of doxorubicin possible due to increased clearance	Monitor for decreased clinical response to doxorubicin

NK-1 antagonist e.g. aprepitant, fosaprepitant, netupitant		
	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**

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: 3 hours

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).

Insert IV cannula or access TIVAD or CVAD.

#### Pre treatment medication

Verify taxane premedication taken or administer as prescribed.

Verify antiemetics taken or administer as prescribed.

#### Ochemotherapy - Time out

#### **Doxorubicin**

#### Administer doxorubicin (vesicant):

- over 5 to 15 minutes
  - via a minibag OR
  - by IV bolus via a side port of a freely flowing IV infusion
- ensure vein is patent and monitor for signs of extravasation throughout administration
- flush with ~150 mL of sodium chloride 0.9%
- potential for flare reaction during administration of doxorubicin (facial flushing and red streaking along the vein) stop infusion and exclude extravasation before continuing at a slower rate of infusion.

Although rare, cardiac arrhythmias may occur during or immediately after doxorubicin administration. If sudden onset of dyspnoea, palpitations or irregular pulse occurs, stop administration immediately and obtain urgent medical officer review.

#### **Docetaxel**

#### Prior to administration:

- · assess patient for fluid retention or weight gain prior to each cycle
  - o notify medical officer of any signs of fluid retention or unexplained weight gain.

The medicines information reference publications stipulate the use of non-PVC containing bags and administration sets. However, this is not consistently recommended in the product information, therefore the decision should be at the discretion of the administering unit.

#### Administer docetaxel (irritant with vesicant properties):

- via IV infusion over 60 minutes
- observe for hypersensitivity reactions
- flush with ~ 100 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.
- for severe reactions seek medical assistance immediately and do not restart infusion.

#### Cyclophosphamide

#### Administer cyclophosphamide:

- via IV infusion over 30 to 60 minutes
- flush with ~ 50 mL of sodium chloride 0.9%
- rapid infusion can cause dizziness, rhinitis, nausea and perioral numbness. If symptoms develop, slow infusion rate.

Remove IV cannula and/or deaccess TIVAD or CVAD.

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

#### **Discharge information**

#### **Antiemetics**

· Antiemetics as prescribed.

#### **Dexamethasone tablets**

• Dexamethasone tablets with written instructions on how to take them.

#### **Growth factor support**

• Arrangements for administration if prescribed.

#### **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 taxanes.  Read more about premedication for prophylaxis of taxane hypersensitivity reactions	
Extravasation, tissue or vein injury	The unintentional instillation or leakage of a drug or substance out of a blood vessel into surrounding tissue. This has the potential to cause damage to affected tissue.  Read more about extravasation management	
Flare reaction	Anthracycline flare reaction is caused by a localised allergic reaction. It is characterised by erythematous vein streaking, urticaria and pruritus which may occur during drug administration and is often associated with too rapid an infusion. Extravasation must be ruled out if flare occurs.	
Nausea and vomiting	Read more about prevention of treatment induced nausea and vomiting	
Red-orange discolouration of urine	Pink/red/orange discolouration of the urine. This can last for up to 48 hours after some anthracycline drugs.	
Taste and smell alteration	Read more about taste and smell changes	
Bone pain	Bone pain, usually in the lower back or pelvis, associated with G-CSF.	

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.  Read more about thrombocytopenia
Arthralgia and myalgia	Generalised joint pain or and/or stiffness and muscle aches, often worse upon waking or after long periods of inactivity. Can improve with movement. May be mild or severe, intermittent or constant and accompanied by inflammation.  Read more about arthralgia and myalgia
Fatigue	Read more about fatigue
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
Diarrhoea	Read more about treatment induced diarrhoea
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
Ocular changes	Symptoms may include eye pain, blurred vision, blepharitis, uveitis, optic neuritis, tear duct stenosis, conjunctivitis, hyperlacrimation, watery or dry eyes and photophobia.
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
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
Fluid retention syndrome	Fluid retention, including peripheral oedema and weight gain, may occur with docetaxel treatment. The main risk factor for development is cumulative docetaxel dose. Pre-medication with dexamethasone may be used. Fluid retention will slowly resolve after cessation of treatment.  Read more about fluid retention syndrome associated with docetaxel
Radiation recall	Erythematous or inflammatory skin reaction resembling severe sunburn at sites previously treated with radiation therapy can occur with certain anti-cancer drugs. Symptoms include vesiculation, desquamation and ulceration of the skin.  Read more about radiation recall

Late (onset weeks to months)		
Anaemia	Abnormally low levels of red blood cells (RBCs) or haemoglobin in the blood.  Read more about anaemia	
Alopecia	Hair loss may occur from all parts of the body. 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	
Cognitive changes (chemo fog)	Changes in cognition characterised by memory loss, forgetfulness and feeling vague. This is also referred to as 'chemo brain' or 'chemo fog'.  Read more about cognitive changes (chemo fog)	
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	

Delayed (onset months to years)		
Menopausal symptoms	Irregular or absent periods, hot flushes, mood swings, sleep disturbance, night sweats, vaginal dryness, decreased libido and dyspareunia. This is caused by ovarian failure and may be temporary or permanent.	
Cardiotoxicity	Anthracyclines are the most frequently implicated anti-cancer drugs associated with cardiotoxicity, which typically manifests as a reduction in left ventricular ejection fraction (LVEF), cardiomyopathy, or symptomatic CHF. Anthracycline induced cardiotoxicity has been categorised into acute, early-onset chronic progressive and late-onset chronic progressive and is usually not reversible. The risk of clinical cardiotoxicity increases with a number of risk factors including higher total cumulative doses.  Read more about cardiac toxicity associated with anthracyclines	

#### Evidence - Adjuvant

The evidence supporting this protocol is provided by a randomised phase III trial (TAX 316) by Martin et al<sup>2</sup> involving 1491 patients with axillary node-positive breast cancer who were randomised to 6 cycles of TAC (docetaxel, doxorubicin, cyclophosphamide) or FAC (fluorouracil, doxorubicin, cyclophosphamide) as adjuvant treatment post surgery.

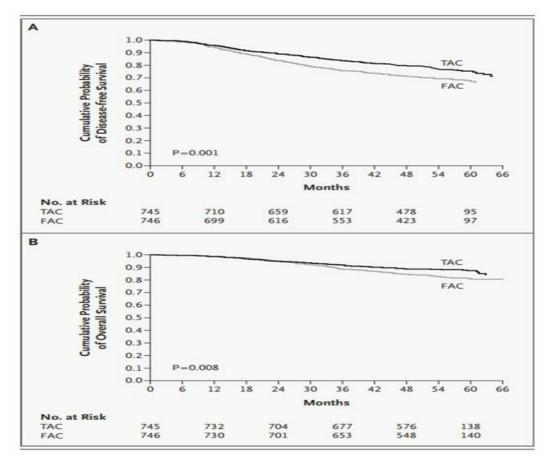
The study enrolled patients between 1997 and 1999 from 20 countries. In total 1480 patients (744 TAC and 736 FAC) received treatment and were included in the safety data. The primary end point was disease free survival (DFS) and the median follow up was 55 months.

TAC patients received ciprofloxacin as supportive care on days 5 to 14 of each treatment cycle. 70% of patients enrolled in this study were ER positive. Randomisation was stratified according to institution and the number of axillary lymph nodes (1 to 3 vs 4 or greater).

#### **Efficacy**

The TAX 316 trial showed there was a statistically significant difference in both DFS (HR 0.72, 95% CI 0.59 to 0.88, p=0.001) and OS (HR 0.70, 95% CI 0.53 to 0.91, p=0.008) favouring TAC over FAC. The estimated rates of DFS at five years were 75% with TAC and 68% with FAC and the overall survival estimates at 5 years were 87% with TAC and 81% with FAC.

Survival rates in the two study groups<sup>2</sup>



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#### **Toxicity**

TAC is more toxic than FAC chemotherapy. Of the TAC group 36% (268/744) of patients developed serious treatment related adverse events compared with 9.2% (67/736) in the FAC arm. The incidence of grade 3/4 adverse events was also higher with TAC (10.2% vs 4.9%). Overall 91.3% of patients completed 6 cycles of TAC and 96.6% completed FAC. TAC was associated with more frequent febrile neutropenia, infection, long-term amenorrhoea, alopecia, peripheral oedema, neurosensory effects and cardiac failure. No deaths occurred as a result of infection. 3 patients developed secondary haematologic cancers (2 in TAC group and 1 in FAC group).<sup>2</sup>

Adverse events* 2	TAC (%)	FAC (%)	<i>p</i> -value
Anaemia	4.3	1.6	0.003
Neutropenia	65.5	49.3	<0.001
Thrombocytopenia	2.0	1.2	0.23
Febrile neutropenia	24.7	2.5	<0.001
Nausea	5.1	9.5	0.001
Stomatitis	7.1	2.0	<0.001

\*Grade 3 or 4 adverse events (except febrile neutropenia - all grades)

#### Evidence - Neoadjuvant

The evidence supporting this protocol is provided by a phase III multicentre international randomised trial (GEPARTRIO) involving 2072 patients with early breast cancer. Between September 2002 and August 2005, 2012 patients were included in the study (intent to treat population).<sup>3</sup>

All patients commenced treatment with two cycles of docetaxel 75 mg/m $^2$ , doxorubicin 50 mg/m $^2$  and cyclophosphamide 500 mg/m $^2$  (TAC), on day 1, every 3 weeks. The early responders (n= 1390) were randomised to received a further four or six cycles of TAC. The non responders (n=622) were randomised to a further four cycles of TAC or switch to a combination of vinorelbine 25

mg/m<sup>2</sup> on days 1 and 8 plus capecitabine 1000 mg/m<sup>2</sup> BD on days 1 to 14, every 3 weeks (NX). No patient received trastuzumab during neo- or adjuvant treatment. Within 21 days after completing chemotherapy and after assessment of the final clinical response patients underwent surgery.<sup>3</sup>

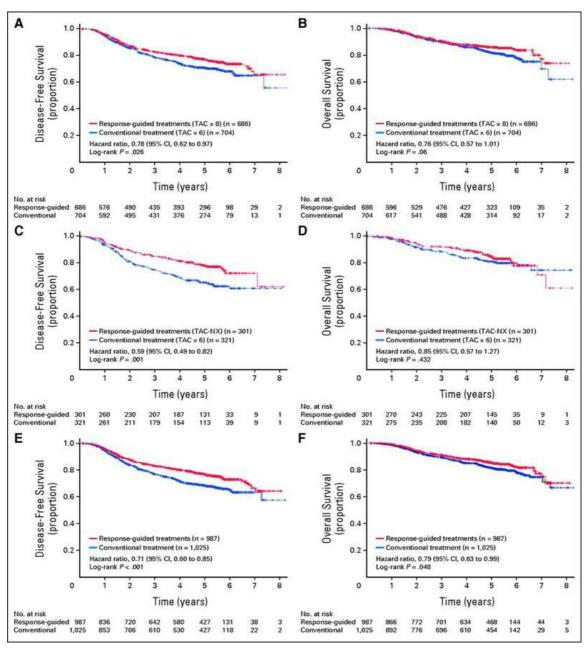
The primary endpoint was to compare the pCR rates in responders and the clinical response rate in non-responders between the randomised arms. The secondary endpoints were disease free survival (DFS) and overall survival (OS).

#### **Efficacy**

At a median follow up of 62 months, DFS was significantly longer in early responders treated with TAC x 8 cycles compared with TAC x 6 cycles (HR, 0.78; 95% CI, 0.62 to 0.97; p=0.026). DFS was also significantly longer in early non-responders treated with TAC-NX than in those receiving TAC x 6 cycles (HR, 0.59; 95% CI, 0.49 to 0.82; p=0.001). There was no significant difference in OS observed in these two groups, though there was a trend toward improved OS in responders receiving TAC x 8 cycles compared with TAC x 6 cycles (HR, 0.76; 95% CI, 0.57 to 1.01; p=0.060). As the sample size was calculated to provide adequate power for the primary end point, the study is underpowered for the endpoints of DFS and OS. In addition, although the groups were reasonably balanced in terms of baseline characteristics, there were fewer patients with grade 3 tumours in the non-responder group treated with TAC-NX (25%) compared with TAC x 6 cycles (36%), and this difference was statistically significant.

Rather than comparing between treatment strategies in the randomised responder vs non-responder groups, the two investigational response-guided arms (TAC x 8 and TAC-NX) were grouped and compared with the conventional therapy arms (TAC x 6). Though this grouping is somewhat counterintuitive, as the intent of the response-guided therapy in responders and non-responders is very different as is the underlying biology, the results suggest that among the hormone receptor-positive patients, DFS was comparable in patients with or without a pCR, but response-guided chemotherapy significantly prolonged DFS compared with conventional chemotherapy. For patients with hormone receptor-negative disease achieving a pCR was associated with longer DFS, but response-guided therapy did not prolong DFS.<sup>3</sup>

DFS and OS in (A, B) responding, (C, D) non-responding, and (E, F) all patients comparing response-guided with conventional chemotherapy<sup>3</sup>



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#### **Toxicity**

The toxicity data for this protocol has been extrapolated from the adjuvant setting.

Grade 3/4 adverse events <sup>2</sup>	TAC (%)
Anaemia	4.3
Neutropenia	65.5
Thrombocytopenia	2.0
Febrile neutropenia	24.7
Nausea	5.1
Stomatitis	7.1

#### References

von Minckwitz, G., S. Kummel, A. du Bois, et al. 2008. "Pegfilgrastim +/- ciprofloxacin for primary prophylaxis with TAC (docetaxel/doxorubicin/cyclophosphamide) chemotherapy for breast cancer. Results from the GEPARTRIO study." Ann

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#### History

#### **Version 1**

Date	Summary of changes
25/08/2022	New multi-indication protocol approved electronically by Medical Oncology reference committee.
30/08/2022	Protocol published on eviQ. Next review in 1 year.
26/10/2022	Treatment schedule updated with missing 'day -1' dexamethasone premedication.

As ID 4114 Breast adjuvant/neoadjuvant TAC (DOCEtaxel DOXOrubicin CYCLOPHOSPHamide) replaces two existing approved protocols, their individual History sections are included below for consistency in documentation.

ID 13 Breast a	adjuvant TAC (DOCEtaxel DOXOrubicin CYCLOPHOSPHamide) version 8
Date	Summary of changes
4/4/2007	PBS listing updated.
1/8/2007	Patient information updated.
29/7/2009	Reviewed and transferred to eviQ.
28/06/2010	Haematological dose modifications updated (20% changed to 25% dose reduction; cut-off for platelets for dose reduction changed from $10 \times 10^9/L$ to $50 \times 10^9/L$ ).
10/12/2010	New format to allow for export of protocol information.  Protocol version number changed to <i>V.2</i> .  Antiemetics and premedications added to the treatment schedule.  Additional Clinical Information, Key Prescribing table and Key Administration table combined into new section titled Clinical Considerations.  Drug specific information placed behind the drug name link.
27/04/2012	Protocol reviewed at Medical Oncology Reference Committee meeting. No change and next review in 2 years.
20/02/2014	Side effects updated, hyperlacrimation and palmar-plantar erythrodysesthesia added.
09/05/2014	Protocol reviewed by email survey. No change and next review in 2 years. PHC view removed.
16/02/2016	Discussion with Medical Oncology Reference Committee Chairs and protocol to be reviewed every 5 years. Next review due in 3 years.
31/05/2017	Transferred to new eviQ website. Protocol version number changed to V.3.  Antiemetic change: Netupitant/palonosetron combination has replaced aprepitant and a 5HT <sub>3</sub> receptor antagonist in combination with dexamethasone for all highly emetogenic regimens.
10/05/2018	Haematological dose modifications updated as per consensus of the expert clinician group. Version number changed to V.4.
24/01/2019	Treatment schedule updated: dexamethasone dose reduced to daily on day 1 and 2 due to drug interaction with netupitant, note added indicating day 3 and 4 dose may not be required as per Medical Oncology Reference Committee consensus. Version number changed to V.5
08/10/2019	Protocol reviewed at Medical Oncology Reference Committee meeting on 30/08/2019. Clinical information updated with PBS expanded indications for GCSF. Next review in 5 years.

03/02/2020	Order of administration for the drugs in TAC note removed.
04/05/2020	Clinical information updated- FBC nadir cycle 1 removed from blood tests, wording changed from 'each treatment' to 'each cycle'. Version number changed to V.7
04/09/2020	Biosimilar drug added to clinical information. Version number changed to V.8.

ID 1295 Breast neoadjuvant TAC (DOCEtaxel DOXOrubicin CYCLOPHOSPHamide) version 7		
Date	Summary of changes	
08/04/2016	New protocol taken to Medical Oncology Reference Committee meeting	
07/07/2016	Protocol approved and published on eviQ	
31/05/2017	Transferred to new eviQ website. Protocol 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.	
10/05/2018	Haematological dose modifications updated as per consensus of the expert clinician group. Version number changed to V.3.	
30/01/2019	Protocol reviewed electronically by Medical Oncology Reference Committee. Treatment schedule updated: dexamethasone dose reduced to daily on day 1 and 2 due to drug interaction with netupitant, note added indicating day 3 and 4 dose may not be required as per Medical Oncology Reference Committee consensus. Version number changed to V.4. Next review in 5 years.	
08/10/2019	Clinical information updated with PBS expanded indications for GCSF.	
03/02/2020	Order of administration for the drugs in TAC note removed.	
04/05/2020	Clinical information updated- FBC nadir cycle 1 removed from blood tests, wording changed from 'each treatment' to 'each cycle'. Approximate treatment time changed to 2.5 hours. Version number changed to V.6.	
04/09/2020	Biosimilar drug added to clinical information. Version number changed to V.7.	

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

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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/4114 08 Jun 2023

# Patient information - Breast cancer adjuvant/neoadjuvant - TAC (docetaxel, doxorubicin, cyclophosphamide)



Patient's name:

#### Your treatment

This treatment can be given either before or after surgery. The aim of neoadjuvant (before surgery) treatment is to shrink the tumour to make it easier to remove. Your doctor will advise which treatment plan is recommended for you.

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

#### TAC (docetaxel, doxorubicin, cyclophosphamide)

This treatment cycle is repeated every 21 days. You will have 6 cycles (or up to 8 cycles for neoadjuvant treatment). Your doctor will discuss your treatment plan with you.

Day	Treatment	How it is given	How long it takes
1	Doxorubicin (dox-oh-roo-bi-sin)	By a drip into a vein	About 3 hours
	Docetaxel (dox-e-tax-elle)		
	Cyclophosphamide (SYE-klo-FOS-fa-mide)		
2	Pegfilgrastim (peg-fil-GRA-stim)	By injection under the skin	About 5 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
<ul> <li>a temperature of 38°C or higher</li> <li>chills, sweats, shivers or shakes</li> <li>shortness of breath</li> <li>uncontrolled vomiting or diarrhoea</li> <li>pain, tingling or discomfort in your chest or arms</li> <li>you become unwell.</li> </ul>	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.

#### Treatment with cyclophosphamide

You should drink at least 8 to 10 glasses of fluid (unless you are fluid restricted) for 2 days after treatment with cyclophosphamide. You should also empty your bladder often.

#### Treatment before surgery (neoadjuvant therapy)

For more information see the eviQ patient information sheet on Anti-cancer therapy before breast cancer surgery (neoadjuvant therapy).

#### 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.
- Antibiotics: you may be given antibiotics to help prevent an infection or to treat an infection quickly if it happens. Your doctor or nurse will tell you how and when to take the antibiotics.
- **G-CSF**: you will be given injection(s) of a drug called G-CSF (also called filgrastim, lipegfilgrastim or pegfilgrastim) under your skin. This helps to boost your white blood cell count. Your white blood cells help to fight infection. Lipegfilgrastim and pegfilgrastim are given once. Filgrastim is given for several days until your white blood cells recover.
- **Docetaxel premedication:** before your treatment with docetaxel you may need to take a tablet called a premedication to help prevent you from having a reaction to docetaxel. A steroid tablet called dexamethasone may be used and should be taken with or after food as directed. The following table may be used to remind you when to take your premedication. Ask your doctor, nurse or pharmacist to fill it out for you.

Tablet	Dose	When to take

Tell your doctor or nurse if you have not taken your premedications before you have your treatment.

#### 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 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.** • This treatment can cause serious injury if it leaks from the area where it is going into the Pain or swelling at injection site (extravasation) This can cause pain, stinging, swelling or redness at or near the site where the drug enters the vein. If not treated correctly, you may get blistering and ulceration. . Tell your doctor or nurse immediately if you get any of the symptoms listed above during or after treatment. • You may get redness and itching along the vein where your chemotherapy is being infused. Redness and itching along This will usually go away within 30 minutes of stopping the injection. vein Tell your doctor or nurse as soon as possible if you get any of the symptoms listed above. Your nurse will check to make sure the drug has not leaked out of the vein. • 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. Your urine will turn an orange or red colour. Urine turning orange or red • This is not harmful and should only last for up to 48 hours after treatment. 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. • You may have discomfort or a dull ache in your pelvis, back, arms or legs. Bone pain after G-CSF • To reduce the pain, take paracetamol before each injection. injection • Tell your doctor or nurse as soon as possible if your pain is not controlled.

#### 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:
  - o a temperature of 38°C or higher
  - o chills, shivers, sweats or shakes
  - o a sore throat or cough
  - uncontrolled diarrhoea
  - shortness of breath
  - o 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.

## Joint and muscle pain and stiffness

- You may get muscle, joint or general body pain and stiffness.
- Applying a heat pack to affected areas may help.
- Talk to your doctor or nurse about other ways to manage these symptoms. You may need medication to help with any pain.

## Tiredness and lack of energy (fatigue)

- You may feel very tired, have no energy, sleep a lot, and not be able to do normal activities or things you enjoy.
- 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 notice a change in the sensations in your hands and feet, including: Nerve damage (peripheral o tingling or pins and needles neuropathy) numbness or loss of feeling 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 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 have: Mouth pain and soreness o bleeding gums (mucositis) o 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 • Tell your doctor or nurse if you get any of the symptoms listed above. · You may get: Eye problems eye pain o red, sore or swollen eyes blurred vision o watery or gritty eyes changes in your eyesight 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.

# Skin that is more sensitive to the sun (photosensitivity)

- After being out in the sun you may develop a rash like a bad sunburn.
- · Your skin may become red, swollen and blistered.
- · 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.

#### Skin rash

- You may get a red, bumpy rash and dry, itchy skin.
- 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.

#### Hand-foot syndrome (palmar-plantar erythrodysaesthesia)

- The palms of your hands and soles of your feet may become:
  - o red and hot
  - swollen
  - 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.

## Extra fluid in the body (fluid retention)

- You may gain weight over a short amount of time.
- Your hands and feet may become swollen, appear red or feel hot and uncomfortable.
- These symptoms are caused by the drug docetaxel.
- Wear loose clothing and shoes that are not too tight.
- Try not to stand up or walk around too much at one time.
- If your ankles or legs get swollen, try raising them.
- Make sure that any cuts or areas of broken skin are treated as soon as possible.
- Tell your doctor or nurse as soon as possible 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 become short of breath.

## Skin reaction in an area previously treated with radiation therapy (radiation recall)

- In the area that was treated with radiation therapy, your skin may become:
  - dry, red and itchy
  - tender and swollen
- It may also:
  - o peel or blister
  - form ulcers
- This usually happens weeks or months after chemotherapy treatment.
- · Avoid wearing tight clothing.
- Avoid direct sunlight and very hot or cold temperatures.
- 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.

Late (onset weeks to mont	hs)
Low red blood cells (anaemia)	<ul> <li>You may feel dizzy, light-headed, tired and appear more pale than usual.</li> <li>Tell your doctor or nurse if you have any of these signs or symptoms. You might need a blood transfusion.</li> <li>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.</li> </ul>
Hair loss (alopecia)	<ul> <li>Your hair may start to fall out from your head and body.</li> <li>Hair loss usually starts 2 to 3 weeks after your first treatment.</li> <li>You may become completely bald and your scalp might feel tender.</li> <li>Use a gentle shampoo and a soft brush.</li> <li>Take care with hair products like hairspray, hair dye, bleaches and perms.</li> <li>Protect your scalp from the cold with a hat, scarf or wig.</li> <li>Protect your scalp from the sun with a hat or sunscreen of SPF 50 or higher.</li> <li>Moisturise your scalp to prevent itching.</li> <li>Ask your doctor or nurse about the Look Good Feel Better program</li> </ul>
Chemo brain (chemotherapy-related cognitive impairment)	<ul> <li>You may notice that you are unable to concentrate, feel unusually disorganised or tired (lethargic) and have trouble with your memory.</li> <li>These symptoms usually improve once treatment is completed.</li> <li>Ask your doctor or nurse for eviQ patient information – Memory changes and chemotherapy (chemo brain).</li> <li>Tell your doctor or nurse if you get any of the symptoms listed above.</li> </ul>
Nail changes	<ul> <li>Your nails may:</li> <li>grow more slowly</li> <li>become darker</li> <li>develop ridges or white lines</li> <li>become brittle and flaky</li> <li>In some cases, you may lose your nails completely.</li> <li>Keep your nails clean and short.</li> <li>Avoid things like biting your fingernails, getting a manicure, pedicure or false nails.</li> <li>Wear gloves when you wash the dishes, work in the garden, or clean the house.</li> </ul>

## Delayed (onset months to years)

#### Menopausal symptoms

- You may get:
  - hot flushes or night sweats
  - mood changes
  - o vaginal dryness
  - o irregular or no periods.
- You may also:
  - have trouble sleeping
  - o find sex painful or lose interest in sex
- These symptoms may go away after treatment, or the menopause may be permanent.
- If you have sex you should use contraception as there is still a risk of pregnancy. Talk to your doctor about what form of contraception is right for you.
- Talk to your doctor or nurse about ways to manage these symptoms.

#### **Heart problems**

- · You may get:
  - chest pain or tightness
  - o 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.
- Grapefruit and grapefruit juice can interact with your medication and should be avoided while you are on this treatment.
- 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 quit 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 quitting.
- 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.

#### **Breast cancer information**

- Australasian Lymphology Association lymphoedema.org.au
- Australasian Menopause Society menopause.org.au
- Breast Cancer Network Australia bcna.org.au
- National Breast Cancer Foundation nbcf.org.au
- YWCA Encore breast cancer exercise program ywcaencore.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 Igfb.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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