

Non-Hodgkin lymphoma oBINUTUZumab maintenance

ID: 3588 v.3 Endorsed

Patients with lymphoma should be considered for inclusion into clinical trials. Link to ALLG website, ANZCTR website and Lymphoma Australia website.

Please note: dexamethasone supportive medication has been added to the treatment schedule as a default. Patients should be individually monitored and dexamethasone reduced or omitted as clinically appropriate, to avoid overmedication.

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 eviQ Estimated Glomerular Filtration Rate (eGFR) calculator.

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

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Click here

2022

Related pages:

- Non-Hodgkin lymphoma bendamustine and oBINUTUZumab
- Non-Hodgkin lymphoma O-CHOP21 (oBINUTUZumab CYCLOPHOSPHamide DOXOrubicin vinCRISTine prednisolone)
- Non-Hodgkin lymphoma O-CVP (oBINUTUZumab CYCLOPHOSPHamide vinCRISTine prednisolone)

Treatment schedule - Overview

Cycle 1 to 12

Drug	Dose	Route	Day
oBINUTUZumab	1,000 mg	IV infusion	1

Frequency: 56 days (2 monthly)

Cycles: 12 doses, or up to 2 years duration of treatment.

Drug status: Obinutuzumab: (PBS authority)

Cost: ~ \$4,610 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 12

Day 1		
Paracetamol	1,000 mg (PO)	60 minutes before treatment

Day 1		
Loratadine	10 mg (PO)	60 minutes before treatment. * May be omitted - please see below for criteria.
Dexamethasone	20 mg (IV)	60 minutes before treatment. * May be omitted - please see below for criteria.
oBINUTUZumab	1,000 mg (IV infusion)	in 250 mL sodium chloride 0.9%. Start at 100 mg/hr. Rate can be increased by 100 mg/hr every 30 minutes, to a maximum of 400 mg/hr. **

* Premedication requirements:

- Antihistamine premedication may be omitted if no infusion related reactions (IRR) occurred with the previous infusion.
- Intravenous corticosteroid premedication may be omitted infusions if no grade 1 or 2 IRR occurred with the previous infusion. If a grade 3 IRR occurred OR lymphocyte count > 25 x10⁹/L prior to next treatment, intravenous corticosteroid premedication should be continued.

Frequency: 56 days (2 monthly)

Cycles: 12 doses, or up to 2 years duration of treatment.

Indications and patient population

- Previously untreated advanced CD20 positive follicular B-cell non-Hodgkin lymphoma following immunochemotherapy
- CD20 positive follicular B-cell non-Hodgkin lymphoma, refractory to treatment with rituximab, following bendamustine containing immunochemotherapy regimen

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 obinutuzumab. Hypotension may occur during obinutuzumab infusion. Evaluate individual patient benefits and risks and consider withholding antihypertensive treatments for 12 hours prior to and during, and for one hour after each infusion. Read more about Hypersensitivity reaction

^{**} If the patient experienced a grade 2 or higher infusion related reaction during the previous administration, commence at 50 mg/hr. The rate of infusion should be titrated up at 50 mg/hr increments every 30 minutes to a maximum of 400 mg/hr.

Premedication	The product information states that the premedication for obinutuzumab should consist of an analgesic/antipyretic, an antihistamine and an intravenous corticosteroid for the first initial dose.
	 For subsequent infusions: Antihistamine premedication may be omitted for subsequent infusions if no infusion related reactions (IRR) occurred with the previous infusion.
	 Intravenous corticosteroid* premedication may be omitted for subsequent infusions if no grade 1 or 2 infusion related reactions (IRR) occurred with the previous infusion. If a grade 3 IRR occurred OR lymphocyte counts > 25 x10⁹/L prior to next treatment, intravenous corticosteroid premedication should be continued. Analgesic/antipyretic premedication is given before all infusions.
	Analgesic/antipyretic premedication is given before all linusions.
	A suggested default premedication has been added to the treatment schedule, and may be substituted to reflect institutional policy.
	Note: hydrocortisone is not recommended as it has not been effective in reducing the rate of
	infusion reactions. * IV corticosteroid (may be substituted by oral corticosteroids if contained within the chemotherapy regimen)
Emetogenicity MINIMAL	No antiemetics should be routinely administered before treatment in patients without a history
	of nausea and vomiting. If patients experience nausea and/or vomiting, consider using the low antiemetic prophylaxis regimen.
	Read more about preventing anti-cancer therapy induced nausea and vomiting
Progressive multifocal leukoencephalopathy	Use of monoclonal antibodies may be associated with an increased risk of progressive multifocal leukoencephalopathy (PML), a rare but potentially fatal opportunistic viral infection of the brain. Patients must be monitored for any new or worsening neurological symptoms.
	Read more about progressive multifocal leukoencephalopathy and the Therapeutic Goods Administration Medicines Safety update on progressive multifocal leukoencephalopathy from the Australian Government, Department of Health.
Cardiac toxicity	Arrhythmias (such as atrial fibrillation and tachyarrhythmia), angina pectoris, acute coronary syndrome, myocardial infarction and heart failure may occur with obinutuzumab, particularly in patients with underlying cardiac disease. These cardiac events may occur as part of an infusion-related reaction and can be fatal.
	Monitor patients for signs and symptoms of cardiac toxicity or fluid overload and refer to cardiologist if suspected.
	Read more about cardiac toxicity associated with anti-cancer drugs
Thrombocytopenia	Severe and life-threatening thrombocytopenia, including acute thrombocytopenia (occurring within 24 hours after the infusion), has been observed during treatment with obinutuzumab. Fatal haemorrhagic events have also been reported in cycle one of treatment.
	Patients should be closely monitored for thrombocytopenia throughout treatment, especially during the first cycle, and use of concomitant medications that may worsen haemorrhagic risk (e.g. antiplatelets, anticoagulants) should be taken into consideration. Correlating the timing of thrombocytopenia after obinutuzumab can assist in differentiating from other causes. In clinical trials, thrombocytopenia was reported in 10.4% of patients treated with obinutuzumab.
Obinutuzumab short	Administration of obinutuzumab by short duration infusion is not in line with the product
duration infusion (SDI)	monograph, however published literature indicates that it can be completed safely in patients who meet the appropriate criteria.
	Read more about obinutuzumab short duration infusion
Antiviral prophylaxis	Read more about antiviral prophylaxis drugs and doses
Pneumocystis jirovecii pneumonia (PJP) prophylaxis	Read more about prophylaxis of pneumocystis jiroveci (carinii) in cancer patients
Blood tests	FBC, EUC, eGFR, LFTs and LDH at baseline, and prior to each cycle and as clinically indicated.

Hepatitis B screening and prophylaxis	Routine screening for HBsAg and anti-HBc is recommended prior to initiation of treatment. 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. 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).

Haematological toxicity	
Platelets x 10 ⁹ /L	
Less than 25	Delay treatment until recovery. If occurring within 24 hours of obinutuzumab administration, consider discontinuing treatment, or alternatively, resume same dose and monitor with supportive management (i.e. transfusions, intravenous immunoglobulins). ¹ ,

Renal impairment	
Creatinine clearance (mL/min)	
30 to 50	No dose adjustments necessary, however patients are at greater risk of neutropenia.
less than 30	No data available for obinutuzumab

Hepatic impairment

Hepatic impairment

No data available for obinutuzumab

Infusion-related reactions (IRR)		
Grade 1 or 2	Reduce infusion rate of obinutuzumab and manage symptoms.	
Grade 3	First occurrence: interrupt obinutuzumab infusion and manage symptoms. Once resolved, restart infusion at no more than half the rate of the infusion rate when IRR occurred. Second occurrence: immediately interrupt obinutuzumab infusion and discontinue	
	treatment.	
Grade 4	Immediately interrupt the obinutuzumab infusion and discontinue treatment.	

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

Obinutuzumab		
	Interaction	Clinical management
Antihypertensives	Additive hypotensive effect	Consider withholding antihypertensive medications 12 hours prior to, throughout and 1 hour after the obinutuzumab infusion

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

Handling of monoclonal antibodies and waste management

Safe administration

General patient assessment prior to each day of treatment.

Prime IV line(s) with sodium chloride 0.9%.

Insert IV cannula or access TIVAD or CVAD.

② Treatment - Time out

Obinutuzumab

Prior to administration:

- · check baseline observations
- check for previous adverse events with drug infusions
- verify premedication has been taken. If not, administer 60 minutes prior to obinutuzumab administration:
- · paracetamol 1000 mg orally AND
- loratadine 10 mg orally (or similar antihistamine premedication may be omitted if no infusion related reactions (IRR) occurred with the previous infusion)
- IV corticosteroid if a grade 3 IRR occurred with the previous infusion OR lymphocyte counts > 25 x 10⁹ /L prior to next treatment.

Infusion:

If no IRR or a Grade 1 IRR occurred with the previous infusion:

- commence obinutuzumab infusion at 100 mg/hr
- · repeat observations prior to each rate increase
- increase rate by 100 mg/hr every 30 minutes to a maximum of 400 mg/hr if observations are stable
- flush with ~ 100 mL of sodium chloride 0.9%

If a Grade 2 IRR or higher occurred with the previous infusion:

- commence obinutuzumab infusion at 50 mg/hr
- repeat observations prior to each rate increase
- increase rate by 50 mg/hr every 30 minutes to a maximum of 400 mg/hr if observations are stable
- flush with ~100 mL of sodium chloride 0.9%

Hypotension may occur during obinutuzumab infusion. Consider withholding antihypertensive medication for 12 hours before, during, and for one hour after each infusion (if appropriate).

Remove IV cannula and/or deaccess TIVAD or CVAD.

Continue safe handling precautions (reproductive risk only) for 7 days after completion of drug(s).

Discharge information

Patient information

· Ensure patient receives patient information sheet.

Prophylaxis medications

• Prophylaxis medications (if prescribed) i.e. PJP prophylaxis, antivirals.

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
Flu-like symptoms	
Headache	
Hypotension	Low blood pressure can occur with this treatment.

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	
Atrial fibrillation		
Fatigue	Read more about fatigue	
Insomnia		
Nausea and vomiting	Read more about prevention of treatment induced nausea and vomiting	
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	

Late (onset weeks to months)		
Anaemia	Abnormally low levels of red blood cells (RBCs) or haemoglobin in the blood. Read more about anaemia	
Progressive multifocal leukoencephalopathy (PML)	A rare opportunistic viral infection of the brain, usually leading to death or severe disability, can occur with monoclonal antibodies (e.g. rituximab, obinutuzumab, ofatumumab, brentuximab vedotin) and other targeted therapies (e.g. ibrutinib, ruxolitinib, idelalisib). Onset may occur up to months after the final dose. Read more about progressive multifocal leukoencephalopathy (PML)	

Evidence

Obinutuzumab was studied as maintenance therapy following an obinutuzumab containing induction regimen for either upfront or refractory lymphoma. Maintenance was offered to patients who achieved either CR or PR following induction therapy. It was administered every two months for 2 years (total of 12 doses) unless progression occurred prior.

The efficacy and toxicity data are addressed in the "parent" eviQ regimens. The eviQ reference committee directs you to the following links that detail the relevant evidence and toxicity sections:

- ID 3554 Non-Hodgkin lymphoma bendamustine and obinutuzumab
- ID 3573 Non-Hodgkin lymphoma O-CHOP21 (obinutuzumab CYCLOPHOSPHamide DOXOrubicin vinCRISTine prednisolone)
- ID 3595 Non-Hodgkin lymphoma O-CVP (obinutuzumab CYCLOPHOSPHamide vinCRISTine prednisolone)

References

1 Haage, T. R., A. Surov, D. Mougiakakos, et al. 2022. "Successful Use of Intravenous Immunoglobulins in an Obinutuzumab-

related Acute Thrombocytopenia." Hemasphere 6(8):e751.

2 Fujiwara, Y., T. Urata, D. Niiya, et al. 2022. "Higher incidence of thrombocytopenia during obinutuzumab plus bendamustine therapy for untreated follicular lymphoma: a retrospective analysis by the Okayama Hematology Study Group." Int J Hematol 115(6):811-815.

Bibliography

Cheson, B. D., N. Chua, J. Mayer, et al. 2018. "Overall Survival Benefit in Patients With Rituximab-Refractory Indolent Non-Hodgkin Lymphoma Who Received Obinutuzumab Plus Bendamustine Induction and Obinutuzumab Maintenance in the GADOLIN Study." J Clin Oncol 36(22):2259-2266.

Hiddemann, W., A. M. Barbui, M. A. Canales, et al. 2018. "Immunochemotherapy With Obinutuzumab or Rituximab for Previously Untreated Follicular Lymphoma in the GALLIUM Study: Influence of Chemotherapy on Efficacy and Safety." J Clin Oncol 36(23):2395-2404.

Marcus, R., A. Davies, K. Ando, et al. 2017. "Obinutuzumab for the First-Line Treatment of Follicular Lymphoma." N Engl J Med 377(14):1331-1344.

Sehn, L. H., N. Chua, J. Mayer, et al. 2016. "Obinutuzumab plus bendamustine versus bendamustine monotherapy in patients with rituximab-refractory indolent non-Hodgkin lymphoma (GADOLIN): a randomised, controlled, open-label, multicentre, phase 3 trial." Lancet Oncol 17(8):1081-1093.

Radford, J., A. Davies, G. Cartron, et al. 2013. "Obinutuzumab (GA101) plus CHOP or FC in relapsed/refractory follicular lymphoma: results of the GAUDI study (B021000)." Blood 122(7):1137-1143.

Trotman, J., S.F. Barrington, D. Belada, et al. 2018. "Prognostic value of end-of-induction PET response after first-line immunochemotherapy for follicular lymphoma (GALLIUM): secondary analysis of a randomised, phase 3 trial." Lancet Oncol 19(11):1530-1542.

History

Version 3

Date	Summary of changes	
22/06/2023	Protocol updated, increase to v.3. Updates include:	
	 clinical information - updated hypersensitivity/ infusion related reaction, thrombocytopenia due to obinutuzumab, cardiac toxicity added 	
	dose modifications - management of infusion-related reactions, management for thrombocytopenia	
	obinutuzumab administration	
	side effects	

Version 2

Date	Summary of changes	
16/05/2022	"Obinutuzumab short duration infusion" block added to clinical information. Version number changed to v.2	
11/11/2022	Protocol reviewed electronically by the Haematology Reference Committee, nil changes. Review in 2 years	

Version 1

Date	Summary of changes
11/03/2019	New protocol developed out of session, discussed by Haematology reference committee (discussed electronically via email).
19/03/2019	Protocol approved and published on eviQ V.1. Review in 1 year.
15/04/2020	Nil changes. Review in 2 years.

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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26 Nov 2023

Patient information - Non-Hodgkin lymphoma (NHL) - Obinutuzumab maintenance



Patient's name:

Your treatment

The treatment schedule below explains how the drug for this treatment is given.

Obinutuzumab maintenance			
This treatment cycle is repeated every 2 months for two years. You will have 12 treatments in total.			
Day	Treatment	How it is given	How long it takes
1	Obinutuzumab (OH-bi-nue-TOOZ-ue-mab)	By a drip into a vein	About 3 hours

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 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

Treatment delays

There may 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. Your doctor will explain if you need any 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.

Central venous access devices (CVADs)

This treatment may involve having chemotherapy through 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 CVADs.

Medications for blood pressure

Obinutuzumab may lower your blood pressure. Tell your doctor if you are taking any blood pressure medications. Your doctor may advise you to temporarily stop your blood pressure medications before your obinutuzumab infusions.

Other medications given during this treatment

• **Obinutuzumab premedication:** before your treatment with obinutuzumab you will need to take some tablets called a premedication to help prevent you from having a reaction to the obinutuzumab.

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.** You may get: Flu-like symptoms a fever chills or sweats muscle and joint pain a cough o 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 can take paracetamol if you have a headache. Headache • Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you get a very bad headache that is not helped by pain medication. • You may get low blood pressure from this treatment. Low blood pressure • You may feel dizzy or light-headed. (hypotension) • Tell your doctor if you are taking blood pressure medication. • Your doctor will monitor your blood pressure regularly while you are on this treatment. • Drink plenty of fluids (unless you are fluid restricted). When you want to get up from a sitting or lying down position, get up slowly to let your body adjust to the new position. • Do not drive or operate machinery if you feel dizzy or light-headed.

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.

• Tell your doctor or nurse if you get any of the signs or symptoms listed above.

- · 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
 - a sore throat or cough
 - uncontrolled diarrhoea
 - shortness of breath
 - a fast heartbeat
 - become unwell even without a temperature.

• This treatment lowers the amount of platelets in your blood. Platelets help your blood to clot. Low platelets When they are low, you are at an increased risk of bleeding and bruising. (thrombocytopenia) • 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. · You may get: **Heart palpitations** o chest pain a pounding or fluttering heart (palpitations) o shortness of breath o dizzy or light-headed confused o more tired than usual. • Tell your doctor if you have any heart problems or are on any heart medications. • Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you get any of the symptoms listed above. • 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 have trouble falling or staying asleep. Difficulty sleeping • Try some gentle exercise daily. (insomnia) • Avoid coffee, tea and other caffeinated drinks around bedtime. • Try something to relax before bed, like a bath or meditation. • If you can't sleep get up and do something quietly, such as reading, until you feel tired. • Tell your doctor or nurse if you have difficulty sleeping. • You may feel sick (nausea) or be sick (vomit). Nausea and vomiting 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. • Anti-sickness medication is usually not needed but may help in some people. · Ask your doctor or nurse for eviQ patient information - Nausea and vomiting during cancer

 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.

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.

Late (onset weeks to months)		
Low red blood cells (anaemia)	 You may feel dizzy, light-headed, tired and appear more pale than usual. Tell your doctor or nurse if you have any of these signs or symptoms. You might need a 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. 	
Changes in the way your brain works [progressive multifocal leukoencephalopathy (PML)]	 This treatment can affect your central nervous system. This can be very serious. Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you get any of the following symptoms: trouble with your speech or vision confusion or memory loss changes in your personality weakness in your arms and legs poor balance or coordination fits (seizures). 	

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 and food safety

- 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.
- There are some foods that may cause infection in high risk individuals and should be avoided. For more information on foods to avoid and food hygiene please ask for a copy of the Listeria and food brochure.

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.

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
- Call the Leukaemia Foundation on 1800 620 420 (Mon to Fri 9am 5pm)
- Call the Lymphoma Nurse Support Line on 1800 953 081 (Mon to Fri 9am 5pm)
- Call the Myeloma Australia Support Line on 1800 693 566 (Mon to Fri 9am 5pm)

Haematology, transplant and cellular therapy information

- Arrow bone marrow transplant foundation arrow.org.au
- Australasian Menopause Society menopause.org.au
- Chris O'Brien Lifehouse Total Body Irradiation mylifehouse.org.au/departments/radiation-oncology/total-body-irradiation/
- Healthy Male Andrology Australia healthymale.org.au/
- International Myeloma Foundation myeloma.org
- Leukaemia Foundation leukaemia.org.au
- Lymphoma Australia lymphoma.org.au
- Myeloma Australia myeloma.org.au
- NSW Agency for Clinical Innovation, Blood & Marrow Transplant Network https://aci.health.nsw.gov.au/networks/bmtct
- NSW Agency for Clinical Innovation aci.health.nsw.gov.au/projects/immune-effector-cell-service

- NCCN Guidelines for Patients Immunotherapy Side Effects: CAR T-Cell Therapy nccn.org/patientresources/patient-resources/guidelines-for-patients
- Talk Blood Cancer cmlsupport.org.uk/organisation-type/social-media-groups

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
- Carer Help carerhelp.com.au
- eviQ Cancer Treatments Online eviQ.org.au
- Food Standards Australia New Zealand: Listeria & Food Safety foodstandards.gov.au/publications/pages/listeriabrochuretext.aspx
- 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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