

Colorectal metastatic raltitrexed

ID: 606 v.4 Endorsed

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



Treatment schedule - Overview

Cycle 1 and further cycles

Drug	Dose	Route	Day
Raltitrexed	3 mg/m ²	IV infusion	1

Frequency: 21 days

Cycles: Continuous until disease progression or unacceptable toxicity

Notes:

The efficacy of raltitrexed in patients who progress on 5-FU/FA is unknown.

Drug status: Raltitrexed is on the PBS general schedule

Cost: ~ \$870 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 and further cycles

Day 2 and 3

Day 1		
Dexamethasone	8 mg (PO)	60 minutes before chemotherapy
Palonosetron	0.25 mg (IV bolus)	30 minutes before chemotherapy
Raltitrexed	3 mg/m ² (IV infusion)	in 50 mL to 250 mL sodium chloride 0.9% over 15 minutes

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Day 2 and 3		
Dexamethasone	8 mg (P0)	ONCE a day (or in divided doses) with or after food. Note: dexamethasone doses on day 2 and 3 may not be required and may be reduced or omitted at the clinicians discretion. *

^{*} Link to ID 7 Prevention of chemotherapy induced nausea and vomiting

Frequency: 21 days

Cycles: Continuous until disease progression or unacceptable toxicity

Indications and patient population

Indications:

- metastatic colorectal cancer where single agent chemotherapy is considered
 - Consider switching to raltitrexed in patients who experience cardiac toxicity from 5FU or capecitabine.¹

Cautions

• severe adverse reactions and death have been reported in patients with renal impairment prescribed raltitrexed.

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
Emetogenicity MODERATE	Suggested default antiemetics have been added to the treatment schedule, and may be substituted to reflect institutional policy.
	For patients with a prior episode of chemotherapy induced nausea or vomiting, a NK1 receptor antagonist is available on the PBS in combination with a 5HT ₃ antagonist and steroid.
	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
Diarrhoea	Antidiarrhoeals (e.g. loperamide) are usually prescribed with this treatment.
	Read more about treatment induced diarrhoea
Blood tests	FBC, EUC and LFTs at baseline and prior to each treatment.
Hepatitis B screening and prophylaxis	Routine screening for HBsAg and anti-HBc is NOT usually recommended for patients receiving this treatment.
	Read more about hepatitis B screening and prophylaxis in cancer patients requiring cytotoxic and/or immunosuppressive therapy
Vaccinations	Live vaccines are contraindicated in cancer patients receiving immunosuppressive therapy and/or who have poorly controlled malignant disease.
	Refer to the recommended schedule of vaccination for immunocompromised patients, as outlined in the Australian Immunisation Handbook.
	Read more about COVID-19 vaccines and cancer.

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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).

The potential for life threatening toxicity increases if the dose is not reduced or if treatment is not stopped as appropriate. Patients with toxicity necessitating discontinuation of raltitrexed should be managed promptly with standard supportive care measures including intravenous hydration and bone marrow support. Calcium folinate (*Leucovorin*®) 25 mg/m² may be given intravenously every 6 hours until resolution of symptoms.

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

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

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Renal impairment	
Creatinine clearance (mL/min)	
55 to 65	Reduce raltitrexed by 25% and change dosing interval to every 28 days
25 to 54	Reduce raltitrexed by 50% and change dosing interval to every 28 days
less than 25	Omit raltitrexed

Hepatic impairment	
Hepatic dysfunction	
Mild/Moderate	No dose modifications necessary but use with caution
Severe	Omit raltitrexed

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 raltitrexed by 25% 3rd occurrence: Reduce raltitrexed by 50% 4th occurrence: Omit raltitrexed
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 raltitrexed by 50% 2nd occurrence: Omit raltitrexed

<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 raltitrexed 25% 3rd occurrence: Reduce raltitrexed by 50% 4th occurrence: Omit raltitrexed
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 raltitrexed by 50% 2nd occurrence: Omit raltitrexed

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

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Raltitrexed		
	Interaction	Clinical management
Folic acid, calcium folinate (Leucovorin®)	Reduced efficacy of raltitrexed likely due to competition for the same sites of action	Advise patients not to take folic acid supplements (including multivitamins) or Leucovorin around the time of receiving treatment with raltitrexed

		treatment with raltitrexed
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

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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 product information monographs via the TGA website for further information.

Day 1

Approximate treatment time: 30 minutes

Safe handling and waste management

Safe administration

General patient assessment prior to each treatment.

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

Prime IV line(s).

Insert IV cannula or access TIVAD or CVAD.

Pre treatment medication

Verify antiemetics taken or administer as prescribed.

Verify dexamethasone taken or administer as prescribed.

Ochemotherapy - Time out

Raltitrexed

- administer via IV infusion over 15 minutes
- flush with 50 mL of sodium chloride 0.9%.

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.

Antidiarrhoeals

· Antidiarrhoeals as 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.

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Immediate (onset hours to days)	
Nausea and vomiting	Read more about prevention of treatment induced nausea and vomiting
Taste and smell alteration	Read more about taste and smell changes
Flu-like symptoms	

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
Oral mucositis	Erythematous and ulcerative lesions of the gastrointestinal tract (GIT). It commonly develops following chemotherapy, radiation therapy to the head, neck or oesophagus, and high dose chemotherapy followed by a blood and marrow transplant (BMT). Read more about oral mucositis
Diarrhoea	Read more about treatment induced diarrhoea
Fatigue	Read more about fatigue
Anorexia	Loss of appetite accompanied by decreased food intake. Read more about anorexia
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	

Evidence

With newer agents such as oxaliplatin and irinotecan being used as first line therapy for advanced CRC, raltitrexed's place in therapy is limited and should only be used in a select group of patients. It may be a used as an alternative for fluorouracil or capecitabine in patients who are unable to tolerate these drugs. Raltitrexed is currently not approved in the United States and is recommended only in clinical trials in the United Kingdom.²

Four large phase III trials have been conducted comparing raltitrexed with fluorouracil/folinic acid (5FU/FA) in advanced colorectal cancer.^{3, 4, 5, 6} Three of the four trials compared raltitrexed with bolus 5FU/FA (Mayo) regimen and one trial compared raltitrexed with bolus and infusional 5FU/FA (De Gramont) regimen.

The trials were well balanced in both treatment arms and end points included overall survival, progression-free survival, response rates and toxicity.

Efficacy

Overall, median survival and response rates of raltitrexed were comparable to 5FU/FA. However, in two of the four trials, time to progression was significantly shorter in the raltitrexed arm. ^{5,6}

Efficacy	Cunningham ³		Maughan ⁴		Cocconi ⁵		Pazdur ⁶	
	Raltitrexed	5FU/FA	Raltitrexed	5FU/FA	Raltitrexed	5FU/FA	Raltitrexed	5FU/FA*
Median Survival (months)	10.3	10.3	8.8	9.8	10.9	12.3	9.7	12.7
Median PFS (months)	4.7	3.6	4.8	5.7	3.9	6.0	3.1	5.6
Response rates (%)	19.3	16.7	18	23	19	18	14.3	15.2

^{*} De Gramont regimen

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Toxicity

When compared with bolus 5FU/FA regimens, raltitrexed had a significantly lower incidence of both severe leucopenia and mucositis.³ However, when compared to the infusional 5FU/FA De Gramont regimen, raltitrexed was associated with greater toxicity and treatment related deaths (18 versus 1). 3 of the deaths were in patients who did not have appropriate dose adjustments for diarrhoea or thrombocytopenia.⁴

Toxicity³

Table 3. Grade 3 or 4 adverse events irrespective of relationship to study treatment and occurring in $\geq 2\%$ of patients in either treatment group.

Event (WHO category)	Number (% with advers	6) of patients se event	P value	Holm's signifi-
	'Tomudex' (n = 222)			level
Anaemia	20 (9)	5 (2)	< 0.001	0.005*
Leucopenia	32 (14)	63 (30)	< 0.001	0.005
Infection	12 (5)	11 (5)	1.000	0.025
Fever	6 (3)	5 (2)	1.000	0.017
Thrombocytopenia	9 (4)	1 (<1)	0.020	0.006
Haemorrhage	4(2)	6 (3)	0.536	0.010
Oral effects				
(mucositis)	5 (2)	47 (22)	< 0.001	0.004
Diarrhoea	32 (14)	29 (14)	0.890	0.013
Nausea and vomiting	29 (13)	20 (9)	0.288	0.007
Constipation	6 (3)	6 (3)	1.000	0.050
Pain	11 (5)	15 (7)	0.420	0.008
Elevated ALT and/				
or AST	23 (10)	0 (0)	< 0.001	0.004
Asthenia ^b	13 (6)	4(2)	0.046	0.006

Statistically significant according to Holm's significance level.

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Toxicity⁴

	de Gramont*	Lokich*	Raltitrexed*
Toxic effect			
Nausea	8 (2.9%)	14 (5.1%)	26 (9.5%)†
Vomiting	9 (3.3%)	12 (4.4%)	21 (7.7%)
Anorexia	9 (3.3%)	17 (6.2%)	30 (11.0%)†
Alopecia	0	1 (0.4%)	0
Rash	4 (1.5%)	7 (2.6%)	7 (2.6%)
Stomatitis	1 (0.4%)	11 (4.1%)†	4 (1.5%)
Diarrhoea	9 (3.3%)	17 (6.3%)	34 (12.4%)†
Lethargy	21 (7.6%)	26 (9.5%)	53 (19.4%)†
Thrombocytopenia	0	0	9 (3.3%)†
Anaemia	4 (1.5%)	4 (1.5%)	8 (2.9%)
Leucopenia	5 (1.8%)	1 (0.4%)	14 (5.1%)
Neutropenia	7 (2.6%)	0	22 (8.0%)†

^{*}Based on between 267 and 277 patients in the de Gramont group, 267 and 273 in the Lokich group, and 267 and 275 in the raltitrexed group with data available. †p<0.01 compared with the de Gramont group.

Table 3: Grade 3 or 4 toxic effects reported by clinicians up to 12 weeks

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References

1 Ransom, D., K. Wilson, M. Fournier, et al. 2014. "Final results of Australasian Gastrointestinal Trials Group ARCTIC study: an

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b Severe event (not WHO graded).

- audit of raltitrexed for patients with cardiac toxicity induced by fluoropyrimidines." Ann Oncol 25(1):117-121
- 2 NICE. 2005. "Irinotecan, oxaliplatin and raltitrexed for the treatment of advanced colorectal cancer. (National Institute for Health and Clinical Excellence guidance)."
- 3 Cunningham, D., J. R. Zalcberg, U. Rath, et al. 1996. "Final results of a randomised trial comparing 'Tomudex' (raltitrexed) with 5-fluorouracil plus leucovorin in advanced colorectal cancer. "Tomudex" Colorectal Cancer Study Group." Ann Oncol 7(9):961-965.
- 4 Maughan, T. S., R. D. James, D. J. Kerr, et al. 2002. "Comparison of survival, palliation, and quality of life with three chemotherapy regimens in metastatic colorectal cancer: a multicentre randomised trial." Lancet 359(9317):1555-1563.
- 5 Cocconi, G., D. Cunningham, E. Van Cutsem, et al. 1998. "Open, randomized, multicenter trial of raltitrexed versus fluorouracil plus high-dose leucovorin in patients with advanced colorectal cancer. Tomudex Colorectal Cancer Study Group." J Clin Oncol 16(9):2943-2952.
- 6 Pazdur, R and M Vincent. 1997. "Raltitrexed (Tomudex) versus 5-fluorouracil and leucovorin (5-FU+ LV) in patients with advanced colorectal cancer (ACC): results of a randomized, multicenter, North American trial."

History

Version 4

Date	Summary of changes
30/04/2010	New protocol taken to Medical Oncology Reference Committee meeting.
25/06/2010	Approved and published on eviQ.
25/01/2011	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.
26/03/2012	PHC OMIS view updated.
01/05/2012	Palonosetron added as the preferred 5HT ₃ antagonist for moderate emetogenicity.
13/09/2013	Protocol reviewed at Medical Oncology Reference Committee meeting. Indications updated to include patients who experience cardiac toxicity from 5FU or capecitabine. Next review in 2 years.
25/08/2014	PHC view removed.
18/02/2016	Discussion with Medical Oncology Reference Committee Chairs and protocol to be reviewed every 5 years. Next review due in 3 years.
07/11/2016	The following change made post Medical Oncology Reference Committee meeting held on 21 October 2016: link to AGTIG and ANZCTR added.
31/05/2017	Transferred to new eviQ website. Version number change to V.3. Hepatitis screening changed to not required.
16/02/2018	Protocol reviewed electronically by Medical Oncology Reference Committee. No changes. Review in 5 years.
10/05/2018	Haematological dose modifications updated as per consensus of the expert clinician group. Version number changed to V.4.
22/06/2018	Antiemetics updated to be in line with international guidelines. Note to dexamethasone added.
01/04/2022	Drug status updated to PBS general schedule.
20/10/2022	Protocol reviewed electronically by Medical Oncology Reference Committee. No changes. Next review 4 years.

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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/606

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Patient information - Bowel cancer metastatic - Raltitrexed



Patient's name:

Your treatment

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

Raltitrexed					
This treatm	This treatment cycle is repeated every 21 days. Your doctor will advise you of the number of treatments you will have.				
Day	Treatment	How it is given	How long it takes		
1	Raltitrexed (ral-ti-TREX-ed)	By a drip into a vein	About 30 minutes		

Tell you doctor if you are taking any medications or supplements that contain folic acid.

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

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.

Other medications given during this treatment

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

Side effects

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

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

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

Immediate (onset hours to da	ys)
Nausea and vomiting	 You may feel sick (nausea) or be sick (vomit). 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.
Taste and smell changes	 You may find that food loses its taste or tastes different. 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.
Flu-like symptoms	 You may get: a fever chills or sweats muscle and joint pain a cough headaches. Tell your doctor or nurse if you get any of the symptoms listed above. Tell your doctor or nurse immediately, or go to the nearest hospital Emergency Department if you have a temperature of 38°C or higher.

Early (onset days to weeks)

Infection risk (neutropenia)

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

Mouth pain and soreness (mucositis)

- · You may have:
 - bleeding gums
 - 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:
 - o 1/4 teaspoon of salt in 1 cup of warm water, or
 - 1/4 teaspoon of bicarbonate of soda in 1 cup of warm water
- Ask your doctor or nurse for eviQ patient information Mouth problems during cancer treatment
- Tell your doctor or nurse if you get any of the symptoms listed above.

Diarrhoea

- You may get bowel motions (stools, poo) that are more frequent or more liquid.
- 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.

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.
Appetite loss (anorexia)	 You may not feel like eating. Try to avoid drinking fluids at meal times. Try to eat small meals or snacks regularly throughout the day. Try to eat food that is high in protein and calories. If you are worried about how much food you can eat, or if you are losing weight, ask to speak to a dietitian.
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)	
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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.

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

Other medical and dental treatment

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

Diet

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

Fertility

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

Pregnancy and breastfeeding

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

Sex life and sexuality

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

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 guitting.
- Talk to your treating team for more information and referral to a smoking cessation support service.

Staying active

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

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

Where to get more information

Telephone support

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

Bowel cancer information

- Australian Council of Stoma Associations australianstoma.com.au
- Australian Government Bladder and Bowel bladderbowel.gov.au
- Australian Government Department of Health & Ageing Stoma appliance scheme health.gov.au/internet/main/publishing.nsf/Content/Stoma+Appliance+Scheme-1
- Bowel Cancer Australia bowelcanceraustralia.org
- National Public Toilet map toiletmap.gov.au
- Recovering after Pelvic Radiation Therapy: A guide for women https://www.targetingcancer.com.au/useful-resources/recovering-after-pelvic-radiation-therapy-a-guide-for-women/

General cancer information and support

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

Quit smoking information and support

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

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

Additional notes:	

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

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