

PATIENT INFORMATION ON

RITUXIMAB

(Brand name: Mabthera)

This information sheet has been produced by the Australian Rheumatology Association to help you understand the medicine that has been prescribed for you. It includes important information about:

- **how you should take your medicine**
- **what are the possible side effects**
- **what tests you will have to monitor your condition**
- **other precautions you should take while you are taking rituximab,**

Please read it carefully and discuss it with your doctor.

Important things to remember

- While taking rituximab you must see your rheumatologist regularly to ensure the treatment is working and minimise any possible side effects.
- If you stop rituximab for any reason you must contact your doctor. Failure to do so may mean that your continued treatment will no longer be subsidised.
- If you are worried about any side effects you should contact your rheumatologist as soon as possible.
- It is important to tell your doctor if you have had cancer or if you develop cancer while you are taking rituximab.
- If you are taking rituximab and plan to become pregnant you must discuss the timing with your doctor.

For more information about RHEUMATOID ARTHRITIS see Arthritis Australia's website: www.arthritisaustralia.com.au

What is rituximab?

Rituximab (brand name Mabthera) belongs to a new class of medicines called **biological disease modifying antirheumatic drugs (biological DMARDs or bDMARDs)**.

bDMARDs have now been given to over a million people worldwide since their initial use in the late 1990s.

B-cells are white blood cells, which normally produce 'antibodies'. Antibodies help protect the body from infections. In rheumatoid arthritis however, some B-cells produce harmful 'autoantibodies', which cause inflammation in the joints. These result in pain, joint swelling and stiffness, and can lead to joint damage.

By temporarily removing the harmful B-cells, rituximab reduces inflammation, lessens the symptoms and helps stop further joint damage. Rituximab also removes some 'good' B-cells, but these return some months after treatment.

Because of its effects on harmful B-cells, rituximab has been used for many years to treat lymphoma, a cancer of the B-cells in lymph nodes.

What benefit can you expect from your treatment?

The improvement in your arthritis from rituximab may take a number of weeks. Benefits will usually be seen by 3 months.

Stopping rituximab

Continue with your treatment unless advised by your doctor or unless side effects develop (see *Side effects*).

If you stop rituximab for any reason you **must** contact your doctor. Failure to do so may mean that your continued treatment may no longer be subsidised.

How will your condition be monitored?

In view of the current prescribing restrictions for all bDMARDs:

- Rituximab will only be given if your disease is active and if standard treatments have been unsuccessful.
- It will not be continued unless it helps your condition. This will be assessed at least 12 weeks after the start of treatment.
- Blood tests will be required during your treatment to monitor your condition and to determine the effectiveness of treatment.
- The frequency of blood tests will depend on what other medicines you are taking and what other illnesses you might have. Your rheumatologist will determine the frequency of tests required.
- You may require blood tests to check your antibody and B-cell levels before you commence a course of treatment and before repeated treatment.

How is rituximab given?

Rituximab is given as a drip (infusion) into the vein. The infusion normally takes 2 to 4 hours. You will need to stay for at least an hour after the infusion to make sure you don't have any side effects.

Rituximab is given in combination with the DMARD methotrexate.

Sometimes a boost of a steroid may be used as part of a premedication to reduce side effects (see *Side effects*).

What is the dosage?

A course of treatment consists of 2 doses given two weeks apart. The dosage is 1000mg for each of the infusions.

Can other medicines be taken with rituximab?

Rituximab may be used with other arthritis medicines including:

- other DMARDs such as methotrexate
- steroid medicines such as prednisolone or cortisone injections into the joint
- anti-inflammatory medicines (NSAIDs) such as naproxen (Naprosyn) or ibuprofen (Brufen, Nurofen)
- simple pain medicines such as paracetamol.

There are separate information sheets for the medicines mentioned above.

Rituximab cannot be used with other bDMARDs. It should also not be given within 4 weeks of treatment with etanercept (Enbrel), within 8 weeks of receiving infliximab (Remicade), adalimumab (Humira), golimumab (Simponi), certolizumab (Cimzia) or abatacept (Orencia).

How long is the treatment continued?

A course of treatment is usually repeated every six months for Rheumatoid Arthritis. Your response will be monitored by your rheumatologist, with blood tests and examination, about 3 to 4 months after your last infusion.

Are there any side effects?

You might experience side effects with your treatment. Contact your doctor if you have any concerns about possible side effects. Many side effects disappear when rituximab treatment is stopped.

Most common possible side effects

- *Blood pressure:* Because rituximab may cause a drop in your blood pressure your doctor may advise you to stop taking your blood pressure medicine temporarily before your treatment.
- *During infusion:* Side effects may include fever, chills, shaking, fatigue, tongue swelling, itch, flushing, fast heart beat, chest pain, shortness of breath or muscle and joint pain.
These effects can usually be reduced by giving corticosteroids (e.g. prednisone or

cortisone), antihistamines and paracetamol before the treatment.

- *Headaches, cough and stomach/bowel discomfort* may also occur.
- *Infections:* Infections (e.g. colds and sinusitis) may occur more frequently than usual.
- *Allergies:* If you have received previous treatment with other biological medicines you may experience an allergic reaction with rituximab.

Less common or rare possible side effects

- *Serious infections:* A rare virus infection of the brain (progressive multifocal leukoencephalopathy, or PML) is found more commonly in patients with immune diseases such as systemic lupus erythematosus (SLE) and rheumatoid arthritis than in the general population. It is thought to be more common in patients with SLE than rheumatoid arthritis but it is still very rare (less than 1 in 10 000 people). This may be increased further in patients with SLE or rheumatoid arthritis who are given rituximab.
- *Diarrhoea.*
- *Muscle stiffness, pins and needles, or numbness in the skin.*
- *Nervousness, feeling anxious or agitated or inability to sleep.*
- *Sweating or night sweats.*
- It is still unclear from research if there is an increased risk of cancer due to rituximab treatment (see *Precautions*).

What precautions are necessary?

Infections

- If you have an active infection of any kind treatment with rituximab will not be given until the infection is treated successfully.
- Rituximab will not be given if you have active untreated HIV (AIDS) infection as it is likely to make these conditions worse.
- Hepatitis B or C infection may not necessarily exclude treatment.

- Because of the risks associated with infection the following tests may be conducted before commencing treatment with rituximab:
 - blood tests for hepatitis B and C
 - chest x-ray
 - HIV tests are required for those who are at risk of this infection.
- If you have an active infection of any kind, it should be treated quickly. See your doctor if you think you have an infection. Treatment with rituximab may be withheld until the infection is treated successfully.

Use with other medicines

- Rituximab can interact with other medicines. You should tell your doctor (including your general practitioner, rheumatologist and others) about all medicines you are taking or plan to take. This includes over the counter or herbal/naturopathic medicines.
- You should also mention your treatment when you see other health professionals.
- Rituximab does not increase the risk of side effects from low dose aspirin (taken for prevention of heart attack and strokes).

Vaccines

- If you are on rituximab it is recommended you should not be immunised with 'live' vaccines such as MMR (measles, mumps and rubella), OPV (oral polio virus), BCG (Bacillus Calmette Guerin) or yellow fever. Talk with your rheumatologist before receiving any vaccines.
- Pneumovax and the combined yearly seasonal flu/swine flu vaccinations are safe and recommended to reduce your risk of those infections.
- Because rituximab removes antibody-forming B-cells, vaccinations are less effective for several months after a course of treatment. You should plan vaccinations before a course of rituximab or between courses. You should discuss this with your rheumatologist or general practitioner.

Surgery

- Rituximab treatment may be delayed if surgery is being planned. Infusions can be recommenced as long as wound healing has taken place and there are no signs of infection.

Cancer risk

- Lymphoma, a cancer of lymph glands, is found more commonly in patients with severe active rheumatoid arthritis than in the general population. To date there is no evidence to suggest that rituximab increases lymphoma.
- If cancer has been previously treated and cured it may be possible for rituximab to be used safely.
- For general cancer prevention, stopping smoking and taking skin cancer prevention measures are recommended. It is important to use sunscreen and avoid prolonged sun exposure. A yearly skin check is recommended.
- Talk to your doctor if you have any concerns about issues relating to cancer risk.

Use with alcohol

- You may drink alcohol while taking rituximab. However, if you are also taking methotrexate you should be cautious about your alcohol intake. It is not known precisely what level of drinking is safe when on methotrexate, however there is general agreement that 1 to 2 standard drinks taken once or twice a week is unlikely to cause a problem.
- Drinking more than 4 standard drinks on one occasion, even if infrequently, is strongly discouraged.

Use in pregnancy and when breastfeeding

- Not enough is known about the possible side effects of rituximab on the unborn baby. If you plan to become pregnant it is important to discuss this with your doctor as each case is different.
- You should not breastfeed when taking rituximab. It is not known whether rituximab passes into breast milk.

Questions?

If you have any questions or concerns write them down and discuss them with your doctor.

Your doctor's contact details

If you are taking rituximab you should see your rheumatologist regularly to make sure the treatment is working and to minimise any possible side effects.

How to help us help you

Sign up to the ARAD project now!

The Australian Rheumatology Association collects information on how well these drugs work and how often they cause problems.

The best way to get this information is from you!

Contact us in any of the following ways:

Email: ARAD@monash.edu

Telephone: Sydney 02 9463 1889 or
Melbourne 03 9508 3424

Fax: 1-800-022-730

Visit our website: www.ARAD.org.au

The information in this sheet has been obtained from various sources and has been reviewed by the Australian Rheumatology Association. It is intended as an educational aid and does not cover all possible uses, actions, precautions, side effects, or interactions of the medicines mentioned. This information is not intended as medical advice for individual problems nor for making an individual assessment of the risks and benefits of taking a particular medicine. It can be reproduced in its entirety but cannot be altered without permission from the ARA. The NHMRC publication: *How to present the evidence for consumers: preparation of consumer publications* (2000) was used as a guide in developing this publication.