I have never taken medication for rheumatoid arthritis before. Should I take methotrexate (Rheumatrex®) alone or with other disease-modifying anti-rheumatic drugs for rheumatoid arthritis?

A Cochrane decision aid to discuss options with your doctor

This decision aid is for you if:
- You are 16 or older.
- Your doctor has told you that you have active rheumatoid arthritis (RA).
- You have never taken methotrexate or any disease-modifying anti-rheumatic drug before such as azathioprine (Imuran®), cyclosporine (Neoral®), gold therapy or sodium aurothiomalate (Myochrysine®), hydroxychloroquine (Plaquenil®), leflunomide (Arava®), penicillamine (Cuprimine®) sulfasalazine (Salazopyrin®, generics), tacrolimus (Prograf®) or others.

What is Rheumatoid Arthritis (RA)?
When you have rheumatoid arthritis, your immune system, which normally fights infection, attacks the lining of your joints making them inflamed. This inflammation causes your joints to be hot, swollen, stiff, and painful. The small joints of your hands and feet are usually affected first. If the inflammation goes on without treatment, it can lead to damaged joints. Once the joint is damaged it cannot be repaired, so treating rheumatoid arthritis early is important.

What is Methotrexate?
Methotrexate is a disease-modifying antirheumatic drug (DMARD). It is the most commonly used DMARD in rheumatoid arthritis. It works to control inflammation in your joints. Methotrexate is taken once per week and it comes in pill form or as an injection. Methotrexate only works while you are taking it. It will take 6 to 8 weeks for the methotrexate to work. It is important for you to keep taking the medicine. Your doctor may start you on a low dose and gradually increase your dose. Pay close attention to your doctor’s directions and to keep your medical appointments to catch any complications early.

What are your options?
- Methotrexate alone
- Methotrexate (MTX) plus other Disease modifying anti-rheumatic drugs (DMARDs)
  - I will discuss the other treatment and prevention options with my doctor.

What other health factors may affect your choice?
Check ✓ any that apply and discuss your concerns with your doctor.
- I have had a previous reaction to methotrexate.
- I am pregnant, planning on becoming pregnant, or am breast feeding.
- I am a man who is planning a pregnancy with my partner.
- I have an active infection.
- I am taking antibiotics containing sulfa such as (Bactrim or Septra).
- I have problems with my liver, kidney, or a blood disorder.
- Other factors: ____________________________

Checklist developed by Andy Thompson MD FRCPC (Revised November, 2007), ©2007 Available at www.RheumInfo.com

Working through the 4 steps of this decision aid may help you decide.
What does the research show?
Here is a ‘best estimate’ of what happens to 100 people who have never tried methotrexate before. There is no way of knowing in advance if you will be affected.

Benefits ++

19 more people who took MTX with other DMARDs had a 50% improvement in the number of tender or swollen joints and other outcomes such as pain and disability (ACR 50) after 1-2 years. This may have happened by chance.

Option A: Take MTX alone
71 Do not improve
29 Improve

Option B: Take MTX with other DMARDs
49 Do not improve
51 Improve

Side Effects +++

2 more people who took MTX alone stopped treatment due to side effects. This may have happened by chance.

We do not have precise information about side effects and complications. This is particularly true for rare but serious side effects. Possible side effects include nausea, headache and tiredness. Rare complications may include lung problems. It is important to keep medical appointments to catch any serious adverse events early.

83 avoid side effects
17 stop treatment

81 avoid side effects
19 stop treatment

Note: The options include ‘estimates’ of what happens to groups of people based on the available research. The quality of these estimates is summarized using the GRADE system as follows:

++++ High quality – further research is very unlikely to change the estimate.
+++ Moderate quality – further research may change the estimate.
++ Low quality – further research is likely to change the estimate.
+ Very low quality – further research is very likely to change the estimate.
**Step 2: Which reasons to choose each option matter most to you?**

Common reasons to choose each option are listed below. Check ☑ how much each reason matters to you on a scale from 0 to 5. ‘0’ means it is not important to you. ‘5’ means it is very important to you.

<table>
<thead>
<tr>
<th>How important is it to you...</th>
<th>not important</th>
<th>very important</th>
</tr>
</thead>
<tbody>
<tr>
<td>…to have 50% improvement in the number of tender or swollen joints and other outcomes such as pain and disability in your RA?</td>
<td>MTX alone 0 1 2 3 4 5</td>
<td>MTX with other DMARDs</td>
</tr>
<tr>
<td>…to avoid adverse events or side effects?</td>
<td>MTX alone 0 1 2 3 4 5</td>
<td>MTX with other DMARDs</td>
</tr>
<tr>
<td>…to avoid taking several different medications?</td>
<td>MTX with other DMARDs 0 1 2 3 4 5</td>
<td>MTX alone</td>
</tr>
</tbody>
</table>

List other reasons that are important:

- ________________________________
- ________________________________
- ________________________________

Now, think about which option has the reasons that are most important to you.

**Which option do you prefer?** Check ☑ one.

- [ ] Methotrexate alone
- [ ] Methotrexate plus other disease modifying anti-rheumatic drugs
- [ ] I will discuss the other treatment and prevention options with my doctor
Step 3: What else do you need to prepare for decision making?

Knowledge
Find out how well this decision aid helped you learn the key facts.
Check the best answer.
1. Which option has the **highest** chance of improving symptoms of rheumatoid arthritis?
   - [ ] MTX
   - [ ] MTX with DMARDs
2. Methotrexate can be taken as a pill or an injection
   - [ ] True
   - [ ] False
3. If 100 people take methotrexate with other DMARDs, how many **more** people had a 50% improvement in the number of tender or swollen joints and other outcomes such as pain and disability compared to people taking methotrexate alone.
   - [ ] 10
   - [ ] 19
   - [ ] 29
   - [ ] Don’t know
4. If 100 people take methotrexate alone, how many **more** people will have side effects compared to people taking methotrexate with other DMARDs?
   - [ ] 2
   - [ ] 15
   - [ ] 40
   - [ ] Don’t know

Check your answers at the bottom of the page.

Do you know enough about the benefits and side effects of taking methotrexate with other DMARDS?
- [ ] Yes
- [ ] No

Values
Are you clear about which benefits and side effects matter most to you?
- [ ]

Support
Do you have enough support and advice from others to make a choice?
- [ ]

Uncertainty
Do you feel sure about the best choice for you?
- [ ]

Step 4: What are the next steps?

- [ ] I will take Methotrexate alone
- [ ] I will take Methotrexate with other disease modifying anti-rheumatic drugs
- [ ] I need more information and I will discuss the options with my doctor

This information is not intended to replace the advice of a health care provider.

Answers for key facts: 1. MTX with DMARDs 2. True 3. 19 4. 2

This decision aid was developed by Rader, T. Maxwell, L. Ghogomu, E. Tugwell, P. Welch, V. Conflict of interest available from trader@uottawa.ca. Funded in part by the Canadian Institutes of Health Research. Format based on the Ottawa Personal Decision Guide © 2000, A O’Connor, D Stacey, University of Ottawa, Canada. Last reviewed: February 2011.

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