Pregnancy, lactation, and postpartum care

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Pregnancy Planning

Contraception

Genetics of multiple sclerosis

Infertility and multiple sclerosis

When should I start pregnancy discussions with my multiple sclerosis provider?

 At least one year before starting to try to become pregnant



What if I don't want to have kids right now?

 Any form of contraception is safe for patients with multiple sclerosis

> Review > Mult Scler. 2017 May;23(6):757-764. doi: 10.1177/1352458517701314. Epub 2017 Mar 24.

Contraception for women with multiple sclerosis: Guidance for healthcare providers

Maria K Houtchens ¹, Lauren B Zapata ², Kathryn M Curtis ², Maura K Whiteman ²
Affiliations + expand
PMID: 28338393 PMCID: PMC5785786 DOI: 10.1177/1352458517701314
Free PMC article



Will I pass my multiple sclerosis to my children?

- Children of one parent with multiple sclerosis
 - Male: 0.3-2.1%
 - Female: 1.1-3.5%
- Children with both parents having multiple sclerosis
 - Male: 12.0-32.0%
 - Female: 12.5-30.4%

> Ann Neurol. 2000 Dec:48(6):927-31.

Conjugal multiple sclerosis: population-based prevalence and recurrence risks in offspring.
Canadian Collaborative Study Group

G C Ebers 1, I M Yee, A D Sadovnick, P Duquette

Affiliations + expand PMID: 11117550



Is Multiple Sclerosis associated with infertility?

- Evidence is conflicting, no clear evidence there is decreased fertility secondary to multiple sclerosis
 - Recommend referral to fertility treatment if pregnancy is not achieved after 3-6 months of optimal conception attempts







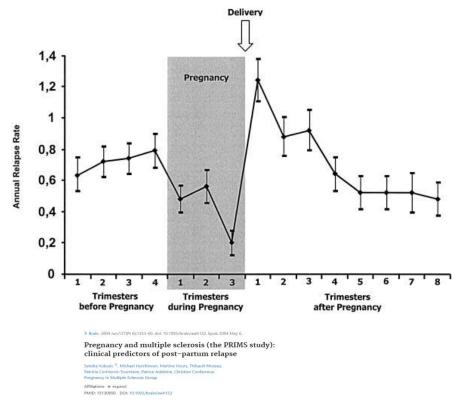
What can we expect in pregnancy

Disease modifying therapy and pregnancy

Symptomatic management during pregnancy

What is the risk of relapse during pregnancy and after delivery?

- Relapses in MS decrease during pregnancy
- Relapse rate increases during the first 3-6 months following delivery
 - Increased rate of both clinical and radiographic relapse





What increases rate of relapse?

- If you had an active MRI brain before becoming pregnant
- If you suddenly discontinued either Tysabri or Gilenya (without treatment planning) you were more likely to experience post partum relapse

 This odds of postpartum relapse decreases with exclusive breastfeeding (chestfeeding) for at least 3 months.



What happens if I have a relapse while pregnant

- MRI imaging safe to undergo while pregnant
 - Avoid contrast dye
- We want to try to avoid steroids in the first trimester
- High dose steroids can be given in the second or third trimester







What can we expect in pregnancy

Disease modifying therapy and pregnancy

Symptomatic management during pregnancy

Disease modifying therapies and pregnancy Self-Injectable therapies

- Interferon beta (Avonex, Rebif, Plegrity etc)
 - This medication can be used right up until positive pregnancy test (discontinue at positive pregnancy test)
 - Extensive data on use during early pregnancy

- Glatiramer acetate (Copaxone)
 - This medication can be used right up until positive pregnancy test, and during pregnancy
 - Extensive safety data



Disease modifying therapies and pregnancy Self-Injectable therapies

Ofatumumab (Kesimpta)

- Short drug half life (16 days), average total clearance 12 weeks
- No adverse effects seen in animals exposed during pregnancy
- As of August 2021, no congenital anomalies or malformations have been reported
- Recommendation is to stop this medication <u>3-6 months</u> before attempting to conceive



Teriflunomide (Aubagio)

- Known to cause embryofetal toxicity
- If you wish to become pregnant, must discontinue the medication and undergo an accelerated drug elimination procedure, check blood levels (must be <0.02mg/L)
- Without accelerated drug elimination it takes between 8 months- 2 years to achieve desired blood level
- This is recommended for both men and women



- Sphingosine-1-phosphate (S1P) receptor modulators (Fingolimod, Siponimod, ozanimod, ponesimod)
 - This medication has evidence for birth defects in animal studies
 - Recommend to discontinue the medication at least 2 months prior to attempting to conceive
 - Possible rebound multiple sclerosis activity after stopping these medications (reported incidence of 4-25%)
 - Recommend to transition to another DMT rather than

sudden stop

Disease Reactivation after Fingolimod Discontinuation in Pregnant Multiple Sclerosis Patients

Assunta Bianco ^{3, 2}, Matteo Lucchini ^{3, 4}, Rocco Totaro ⁵, Roberta Fantozzi ⁶, Giovanna De Luca ⁷ Sonia Oi Lemme ⁶, Giorgia Presicce ³, Luana Evangelista ⁵, Valeria Di Tommasco ⁷, Roberta Pastorino ⁶, Cilviar De Fino ³, Valeria De Arcangelis ³, Diego Centonze ⁶, 9, Massimillano Mirabelista ^{4, 4}

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PMID: 34494237 PMCID: PMC8803993 DOI: 10.1007/s13311-021-01106-6



- Dimethyl Fumarate (Tecfidera and generic)
 - This medication has no known current evidence for human fetotoxicity
 - Recommend discontinuation of medication and attempt to conceive
 - No rebound associated with stopping this medication



Cladribine (Mavenclad)

- Known to cause fetal harm in animal studies
 - Recommend two forms of birth control while on medication!
- Recommend waiting <u>6 months after last dose</u> before attempting to conceive
 - Due to the mechanism of action of this drug, protection against relapse continues after last dose
 - This medication will drop your white blood cell count, with the lowest counts occurring 2-3 months after treatment. Need to make sure these counts recover before attempting pregnancy

Disease modifying therapies and pregnancy Infusion medications

- B cell depleting agents (Ocrevus/Ocrelizumab, Rituxan/Rituximab)
 - Recommendation to wait 2-3 months after infusion date to attempt to conceive
 - B cells still depleted for a total of 6 months (in some individuals this effect may be longer), allowing for continued protection against relapse while attempting to conceive



Disease modifying therapies and pregnancy Infusion medications

Alemtuzumab (Lemtrada)

- Recommend to wait 4 months from last infusion before attempting to conceive
- Due to the mechanism of action of this drug, protection against relapse continues after last infusion
- Development of thyroid related complications are common with this medication
 - Recommend thyroid studies prior to attempting to conceive and continued monitoring during pregnancy



Disease modifying therapies and pregnancy Infusion medications

Natalizumab (Tysabri)

- Recommended to wait to attempt to become pregnant 1-3 months after last infusion.
- High risk for rebound multiple sclerosis activity after stopping infusion
 - Up to 40% of women will experience rebound relapse
 - 10-20% risk of significant long-term disability from rebound activity
- Please make a plan with your multiple sclerosis provider!







What can we expect in pregnancy

Disease modifying therapy and pregnancy

Symptomatic management during pregnancy and post-partum considerations

Fatigue, Multiple Sclerosis, and Pregnancy

- Fatigue is a common symptom of multiple sclerosis, reported by at least 75% of MS patients.
- Some (off label) medications used to treat fatigue include Modafinil, Ritalin, Adderall, Amantadine
 - Modafinil should be discontinued prior to pregnancy due to data suggesting increased risk for congenital malformation
- Cognitive behavioral therapy
 - Shown to be as effective in reducing fatigue as compared with modafinil

Session CE1 - Cutting Edge Developments 1
CE1.2. A Comparative Effectiveness Trial of TelephoneDelivered Cognitive Behavioral Therapy, Modafinil, and
Combination Therapy of Both Interventions for Fatigue in
Multiple Sclerosis: Primary Results From the COMBO-MS
Trial



Postpartum considerations

- Consideration for pelvic floor physical therapy for any worsening urinary symptoms or pain
- Monitor post partum depression and anxiety
- Consideration of physical or occupational therapy needs







Breastfeeding (chestfeeding)

Disease modifying therapy and breastfeeding

Reasons to consider breastfeeding (chestfeeding)

- Breastfeeding (chestfeeding) decreases risk of postpartum relapse
- Decreases risk for type 2 diabetes, postpartum depression, high blood pressure, breast and ovarian cancer for mom
- Breastfed (chestfed) babies have a lower risk of

infections, SIDS, type 2 diabetes

Meta-Analysis > JAMA Neurol. 2020 Mar 1;77(3):327-338. doi: 10.1001/jamaneurol.2019.4173.

Association Between Breastfeeding and Postpartum Multiple Sclerosis Relapses: A Systematic Review and Meta-analysis

Kristen M Krysko ¹, Alice Rutatangwa ¹, Jennifer Graves ^{1 2}, Ann Lazar ³, Emmanuelle Waubant ¹

Affiliations + expand

PMID: 31816024 PMCID: PMC6902174 DOI: 10.1001/jamaneurol.2019.4173



Disease modifying therapies and Breastfeeding (chestfeeding)

- Safe for breastfeeding (data available to confirm safety)
 - Glatiramer acetate (copaxone)
 - Interferon beta (Avonex, Rebif, etc)



Disease modifying therapies and Breastfeeding

- Probably Safe for breastfeeding, (limited data)
 - Rituximab/Rituxan
 - Ocrelizumab/Ocrevus
 - Ofatumumab/Kesimpta
 - Natalizumab/Tysabri



Monoclonal antibody treatment during pregnancy and/or lactation in women with MS or neuromyelitis optica spectrum disorder

