Levothyroxine treatment in women with thyroid antibodies may not increase live birth rate
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Treating women with thyroid antibodies but a normal thyroid function with a medicine called Levothyroxine does not make them more likely to deliver a live baby, new research led by the University of Birmingham suggests.

The research, which was led by researchers from the Institute of Metabolism and Systems Research, Birmingham Clinical Trials Unit, the Institute of Applied Health Research, and Tommy's Centre for Miscarriage Research at the University of Birmingham, was published today (March 23rd) in New England Journal of Medicine. Funded by the MRC and the National Institute for Health Research (NIHR), the research was also presented today at ENDO 2019, the Endocrine Society's annual meeting in New Orleans, USA.

Miscarriage occurs in one in five women who conceive, making it one of the most common complications of pregnancy. Previous research has found a strong association between the presence of thyroid peroxidase antibodies and miscarriage. However, it was unclear from previous evidence whether treating women with normal thyroid function and with thyroid peroxidase antibodies with Levothyroxine would improve live-birth rates.

The University of Birmingham has now led the largest study of its kind to investigate whether treatment with Levothyroxine would increase the rates of live births at or beyond 34 weeks gestation among women who have thyroid antibodies and also a history of miscarriage or infertility, does not improve the chances of live birth.

"We were surprised by the results of our study as previous small studies suggested there could be a benefit of Levothyroxine treatment in women with thyroid antibodies."

The authors conducted a study of 940 women at 49 UK hospitals with normal thyroid function aged between 16 and 41 who were positive for thyroid peroxidase antibodies; had a history of miscarriage or infertility, and were trying to conceive naturally or with assisted conception.

Outcomes in both groups were similar: 266 of the 470 (56.6 per cent) women who received Levothyroxine, and 274 of 470 (58.3 per cent) women who received a placebo, became pregnant; 176 women (37.4 per cent) taking Levothyroxine, and 178 (37.9 per cent) taking a placebo, had live births.

Dr. Rima Dhillon-Smith added: "Thyroid peroxidase antibodies are found in the blood in approximately one in 10 women who have normal thyroid function, and they have been linked to increased risk of miscarriage and preterm birth.

"International guidelines currently recommend the consideration of Levothyroxine treatment for women with thyroid antibodies, as there is thought to be minimal chance of harm and a potential to help increase the chance of having a live birth.
"As our study was large and of high quality, we can now be confident that Levothyroxine does not improve pregnancy success for women with thyroid antibodies and normal thyroid function and therefore should not be recommended or used in clinical practice. This will mean no longer providing unnecessary medication to women who do not need it."

Dr. Kristien Boelaert, also of the University of Birmingham, said: "Our trial has definitively answered an important clinical question.

"In a recent survey we carried out of UK fertility clinicians almost 40 per cent said they routinely use Levothyroxine in women with thyroid peroxidase antibodies to reduce miscarriage and pre-term birth.

"We now pose the question of whether testing for thyroid peroxidase antibodies should be performed at all in women with infertility or previous miscarriages.

"While thyroid peroxidase antibodies testing may inform about future risk of progression to thyroid disease, our research has shown its treatment with Levothyroxine does not improve pregnancy outcomes and may only generate patient anxiety and unnecessary healthcare costs.

"We hope that national and international guidelines are updated to remove current recommendations which advise consideration of the use of Levothyroxine in these women.

"We also hope that the current practice of routine testing of thyroid peroxidase antibodies in high risk populations such as women with miscarriage and reduced fertility is re-considered."

Provided by University of Birmingham