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## Outcomes of Pregnancy in Women With Idiopathic Inflammatory Myopathies in Africa



Dear Editor,

Idiopathic inflammatory myopathies (IIMs) are associated with adverse pregnancy outcomes in Caucasian and Asian populations,<sup>1,2</sup> but this issue is unclear in Africa. For the purpose of this paper, we conducted a systematic review of the literature to identify studies on IIMs and pregnancy in Africa from electronic and hand searches up to March 4, 2021, using key search terms referring to IIMs, pregnancy and African countries as per the United Nations Classification.<sup>3</sup>

Of 118 records retrieved from PubMed, Embase, Africa Journals Online and hand searches, we included 4 relevant case reports and 2 case-series<sup>4–9</sup> from Gabon, Mali, Morocco, Senegal and Tunisia. The search strategy in PubMed and Embase as well as the study selection process are summarized in the [Supplementary Table](#) and the [Supplementary Figure](#). Included records report a total of 18 singleton post-IIM pregnancies and 10 singleton pre-IIM pregnancies in 12 women aged 26–42 years at conception. Among women with ethnicity data, 6 were Black Africans, 1 Black Caribbean and 1 North African. Specified IIM subtypes were overlap myositis ( $n=4$ ), dermatomyositis ( $n=4$ ) and immune-mediated necrotizing myopathy ( $n=2$ ). Regarding pre-IIM pregnancies, there were only 2 adverse pregnancy outcomes: medical termination of a pregnancy (for unspecified cause) and one stillbirth. In women with post-IIM pregnancy data, 8 of 18 pregnancies were successful. Adverse maternal outcomes recorded in post-IIM pregnancies were premature delivery ( $n=4$ ), cesarean section ( $n=3$ ), medical termination for unspecified causes ( $n=3$ ) and pulmonary infection ( $n=1$ ). Adverse fetal/neonatal outcomes were pre-term birth ( $n=4$ ), neonatal death ( $n=2$ ), small for gestational age ( $n=2$ ), stillbirth ( $n=1$ ) and neonatal lupus ( $n=1$ ) ([Table 1](#)).

Maternal and offspring outcomes of pre- and post-IIM pregnancies are poorly characterized in Africa. It remains unknown whether the observed adverse outcomes were coincidental or connected with IIMs, although this small pooled sample likely suggests together with studies from other regions<sup>1,2</sup> that, increased rates of adverse outcomes may be observed in women (and their infants) with IIMs in Africa as well. There is a need for a prospective multicenter African registry to better assess the link between IIMs and adverse pregnancy outcomes, as well as the impact of pregnancy on IIM activity in Africa.

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## Conflicts of interest

The authors declare no conflict of interest.

## Appendix A. Supplementary data

Supplementary data associated with this article can be found, in the online version, at [doi:10.1016/j.reuma.2021.04.005](https://doi.org/10.1016/j.reuma.2021.04.005).

**Table 1**

Characteristics and outcomes of pregnancy in women with idiopathic inflammatory myopathies in Africa.

| Study             | Period of recruitment | Country | Patient number | Ethnicity/race                 | Age, years           | Number of pre-IIM pregnancy(ies) | Outcomes of pre-IIM pregnancy             | Year of diagnosis of IIM | Subtype of IIM                | Post-IIM pregnancy | Treatment before pregnancy | Disease activity at conception and throughout pregnancy | treatment of IIMs through pregnancy | Outcomes of post-IIM pregnancy         | Post-partum outcome of IIMs   |                      |
|-------------------|-----------------------|---------|----------------|--------------------------------|----------------------|----------------------------------|---|--------------------------|-------------------------------|--------------------|----------------------------|---|-------------------------------------|--|---|----------------------|
|                   |                       |         |                |                                |                      |                                  |   |                          |                               | Number             | Year                       |   |                                     |  |   |                      |
| Iba-Ba, 2009 [4]  | 2004–2005             | Gabon   | 1              | Black African                  | 30                   | 0                                | –   | 2004                     | Myositis overlapping with SLE | 1                  | 2005                       | -GCs -AM  | Inactive                            | None                                   | -Premature delivery<br>-Cesarean section at 32 weeks, due to pneumonia<br>-Neonatal death at day 12 of life | NR                   |
| Kaddour, 2009 [5] | 1979–2007             | Tunisia | 2              | NR                             | 21<br>27<br>28       | 0                                | –   | 1993                     | Myositis overlapping with RA  | 0<br>1<br>2        | –<br>1999<br>2000          | -<br>GCs<br>-GCs<br>-MTX                                | Inactive<br>Inactive                | -<br>GCs<br>-GCs<br>-MTX               | -<br>Successful Medical termination for unspecified cause   | NR<br>NR             |
|                   |                       |         | 3              |                                | 30<br>32<br>22<br>26 | 2                                | Successful                                | 2000                     | Myositis overlapping with APS | 3<br>4<br>1        | 2002<br>2004<br>2004       | GCs<br>GCs  | Inactive<br>Inactive                | GCs<br>GCs                             | Successful  | NR<br>NR             |
|                   |                       |         | 4              |                                | 28                   |                                  |   |                          |                               | 2                  | 2006                       | GCs   | Inactive                            | -GCs<br>-LDA<br>-LMWH                  | Stillbirth at 28 weeks gestation<br>-Premature delivery at 32 weeks gestation<br>-Neonatal death at day 3   | NR                   |
|                   |                       |         | 41             |                                | 41                   | 3                                | Successful                                | 2002                     | Myositis overlapping with SLE | 0                  | –                          | –   | –                                   | –                                      | –   | –                    |
|                   |                       |         | 42             |                                | 42                   | 1                                | Medical termination for unspecified cause |                          |                               | 1                  | 2003                       | -GCs -AM  | Inactive                            | -GCs -AM                               | Medical termination for unspecified cause   | NR                   |
|                   |                       |         | 5              |                                | 25<br>28             | 0                                | Stillbirth                                | 2003                     | DM                            | 0<br>1             | –<br>2006                  | -<br>GCs<br>-AM<br>-MTX                                 | Inactive                            | -<br>GCs<br>-MTX<br>-AM<br>-GCs<br>-AM | -<br>Medical termination for unspecified cause  | NR                   |
|                   |                       |         | 29             |                                |                      |                                  |   |                          |                               | 2                  | 2007                       |   | Inactive                            |  | Successful  | NR                   |
| Ousmane, 2016 [6] | 2012                  | Senegal | 6<br>7         | Black Caribbean                | 26<br>26             | 1<br>1                           | Successful<br>Successful                  | 2008<br>2012             | DM<br>IMNM                    | 1<br>0             | 2008                       |   |                                     |  | Successful<br>–   | NR<br>Flare          |
| Awatef, 2016 [7]  | 2016                  | Morocco | 8<br>9         | Black African<br>North African | 35<br>28             | 1<br>0                           | Successful<br>–                           | 2012<br>NR               | IMNM<br>DM                    | 0<br>1             | –<br>NR                    | –<br>NR   | –<br>Active                         | –<br>GCs<br>-AM                        | –<br>Successfull  | Flare<br>Improvement |
| Cisse, 2018 [8]   | NR                    | Mali    | 10             | Black African                  | NR                   | NR                               | NR  | NR                       | DM                            | 1                  | NR                         | NR  | NR                                  | NR                                     | Neonatal cutaneous lupus  | NR                   |
| Iba-Ba, 2019 [9]  | 2008–2018             | Gabon   | 11             | Black Africans                 | 20<br>21             | NR<br>NR                         | NR<br>NR                                  | NR                       | NR                            | 1                  | NR                         | NR  | Inactive                            | NR                                     | -Caesarean section<br>-Term Live birth  | NR                   |
|                   |                       |         | 23             |                                | NR                   |                                  | NR  |                          |                               | 2                  | NR                         | NR  | Inactive                            | NR                                     | -Premature delivery<br>-Caesarean section   | NR                   |
|                   |                       |         | 24             |                                | NR                   |                                  | NR  |                          |                               | 3                  | NR                         | NR  | Inactive                            | NR                                     | -SGA<br>-Premature delivery<br>-Caesarean section   | NR                   |
|                   |                       |         | 12             |                                | 29<br>36.5<br>39.5   | NR<br>NR<br>NR                   | NR<br>NR<br>NR                            | NR                       | NR                            | 0<br>1<br>2        | –<br>NR<br>NR              | –<br>NR<br>NR   | Inactive<br>Inactive                | –<br>NR<br>NR                          | -Successful<br>Successful   | NR<br>NR             |

IIM, idiopathic inflammatory myopathy; SLE, systemic lupus erythematosus; GCs, glucocorticoids; AM, antimalarials; NR, not reported; RA, rheumatoid arthritis; MTX, methotrexate; APS, antiphospholipid syndrome; LDA, low-dose aspirin; LMWH, low molecular weight heparin; DM, dermatomyositis; IMNM, immune-mediated necrotizing myopathy; SGA, small for gestational age.

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## The Value of a Negative Antinuclear Antibody (ANA) Test: An Often Forgotten Result



### *El valor de una prueba de anticuerpos antinucleares (ANA) negativa: un resultado a menudo olvidado*

Dear Editor:

It is quite clear that medicine is biased towards positive results and the same applies to the practice of pathology.<sup>1</sup> One of the ubiquitous tests in autoimmunity, the antinuclear antibody (ANA) suffers from this very same fate. A number of guidelines report on the clinical utility of a positive ANA and dissuade clinicians from requesting this test in the setting of low pre-test probability

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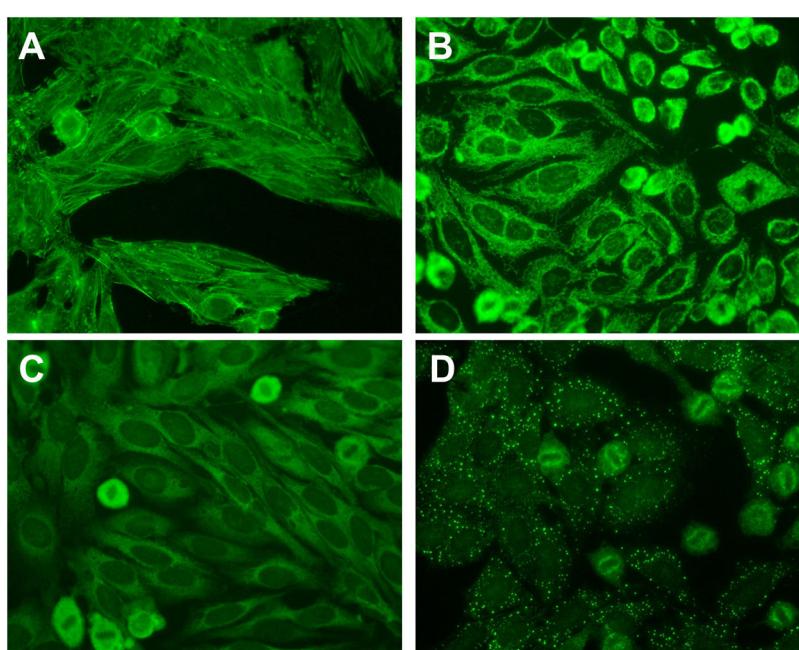
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for an ANA-associated autoimmune disorder (AAD).<sup>2</sup> This is certainly sound advice and prevents unnecessary investigations and healthcare expenditure. Yet, it is important to realise the clinical importance and pitfalls of a negative ANA results which sometimes becomes forgotten.

The internationally-accepted “gold standard” to measure ANA is via indirect immunofluorescence on HEp-2 cells.<sup>3</sup> A negative ANA test on HEp-2 substrate usually means that there is no significant detection of IgG ANA (in the nucleus) at a specified dilution of serum – usually 1:80 to 1:160. There is a move to also classify positive cytoplasmic and mitotic staining of the HEp-2 substrate as ANA positive.<sup>3,4</sup> This may improve the sensitivity of detecting AADs and prompt appropriate further testing and follow-up (Fig. 1).<sup>3</sup>



**Fig. 1.** Example cytoplasmic staining on the HEp-2 substrate. (A) F-actin staining suggesting the presence of smooth muscle antibodies found in autoimmune hepatitis and related disorders. (B) Coarse, granular cytoplasmic staining suggestive of anti-mitochondrial antibodies found in primary biliary cirrhosis. (C) Smooth, homogenous cytoplasmic staining suggestive of anti-ribosomal P antibodies found in systemic lupus erythematosus. (D) Large cytoplasmic dots staining suggestive of anti-GW bodies. All micrographs are taken at a magnification of 400×.