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COVID-19 Antibody Testing in Pregnancy

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1 Clinical Perspective

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1 Clinical Perspective

- **COVID-19 Antibody Testing in Pregnancy** 2 Fabrizio Zullo MD,¹ Daniele Di Mascio MD,² Gabriele Saccone MD.¹ 3 ¹Department of Neuroscience, Reproductive Sciences and Dentistry, School of Medicine, 4 University of Naples Federico II, Naples, Italy 5 ²Department of Maternal and Child Health and Urological Sciences, Sapienza University of Rome, 6 7 Rome, Italy Disclosure: The authors report no conflict of interest 8 Financial Support: No financial support was received for this study 9 10 Correspondence: Gabriele Saccone, MD. Department of Neuroscience, Reproductive Sciences and 11 Dentistry, School of Medicine, University of Naples Federico II, Naples, Italy 12 Email: gabriele.saccone.1990@gmail.com 13 Running head: Antibody Response to COVID-19 14 Key words: SARS-COV-2, COVID-19, coronavirus, 2019-nCoV, influenza, vaccine, coronavirus 15 Disclosure of interests: None declared 16 Contribution to authorship: Sole author 17 Details of ethics approval: Not applicable 18 Funding: None 19
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The novel coronavirus 2019, or Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2), is the virus responsible for COVID-19 infection, which can has been associated with maternal and perinatal morbidity and mortality.^{1,2} Almost all patients with COVID-19 infection test positive for antiviral immunoglobulin-G (IgG) within about 10-20 days after symptom onset (Figure 1), but the clinical value of antibody testing has not yet been completely elucidated, either in non-pregnant or even more pregnant patients.³

There are different ways to test for antibody against SARS-CoV-2. The three most common methods currently are IgM and IgG titer measured by either chemiluminescence immunoassay analysis, or enzyme-linked immunosorbent assay (ELISA); and a rapid (results within 15 minutes) IgM-IgG combined antibody test.^{.3} Their sensitivities and specificities are still being studied and vary, but have been reported to be about 48%, 89%, and 89%, respectively; and 100%, 91%, and 91%, respectively.³⁻⁵

Testing pregnant women for antibody response to COVID-19 may have different advantages, including identifying: 1. Possibly 'healed' women (e.g. IgG positive) never tested with real-time reverse-transcriptase-polymerase-chain-reaction (RT-PCR) assay of nasopharyngeal (NP) swab specimens; 2. Women still at risk for COVID-19 infection (e.g. IgM and IgG negative).

Women who do know their infectious status represent a potential threat to others, including healthcare workers (HCWs) and other patients. Indeed, some governments have suggested that the detection of antibodies to SARS-CoV-2 could serve as the basis for an "immunity passport" or "risk-free certificate" (digital or physical documents that certify an individual has been infected and is purportedly immune to SARS-CoV-2) that would enable individuals to, for example, return to work or travel assuming that they are protected against re-infection.⁶

The use of the point-of-care rapid combined antibody test can be of paramount importance inobstetric healthcare settings and may be particularly helpful in testing women before outpatient

45 (Figure 2A) and inpatient (Figure 2B) visits. After the rapid test, the following results can be46 reported:

• IgG negative, IgM negative.

IgG positive, IgM negative. There are limited data on when IgM disappears.^{3,4} Therefore,
even if there is a high chance that the tested person is not contagious anymore, a NP swab should be
offered.

IgG positive, IgM positive or IgG negative, IgM positive. The presence of IgM increases the
 chance that the tested person is still contagious.⁴

We suggest that, if rapid antibody testing and personnel are available, algorithms for care before 53 54 outpatient and inpatient care of pregnant women be implemented as suggested in Figure 2. At our institution, the Department of Obstetrics and Gynecology at University of Naples Federico II 55 (Naples, Italy), all pregnant women are tested with the rapid combined antibody test before hospital 56 admission. If the rapid antibody test is positive to either SARS-CoV-2 IgM and/or IgG, we do offer 57 the NP swab and consider these women COVID-19 positive until the result of the NP swab is 58 available. Patients positive to the rapid combined antibody test are isolated, and inpatient admission 59 is postponed waiting the results of the NP swab, if feasible (e.g. planned cesarean delivery, 60 induction of labor, surgical procedure). If it is not feasible to postpone the admission (e.g. laboring 61 or bleeding pregnant women), the patient is admitted to the COVID unit, and managed as COVID-62 19 positive (Figure 2A). For example, in our Department, recently two pregnant women with 63 positive IgG and IgM had admission to the hospital postponed of 24 hours while awaiting the NP 64 swab test result. In another patient, who tested positive for IgM, postponed hospital admission was 65 not feasible due to heavy bleeding in a first trimester spontaneous abortion. The patient received NP 66 swab and was admitted to the COVID unit where she received dilation and curettage. Result of the 67 RT-PCR assay, available the day after, showed positivity to SARS-CoV-2. 68

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We are now also testing women scheduled for outpatient visits. Those tested positive to either IgM or IgG at the rapid combined antibody test, have NP swab offered, and the outpatient appointment is postponed, as shown in Figure 2B. Women with prior infection, and 'certified' recovered because of two negative NP swabs 24 hours apart, are not tested for antibody response to COVID-19.

It would also be helpful, if available, to test visitors, and HCWs. In our Department, we have
mandatory rapid antibody testing for all HCWs every 7 days. HCWs positive to either IgM or IgG
self-isolate at home, waiting for the results of the NP swab.

In summary, we recommend testing for antibody response to SARS-CoV-2 for pregnant women before receiving care in both the inpatient and outpatient setting, as feasible (Figure 2). Those testing positive to either IgM or IgG rapid immunoassay should receive NP swab, and admission or appointment should be postponed, if feasible, until NP swab test results is available, and considered COVID-19 positive in the meanwhile (Figure 2).

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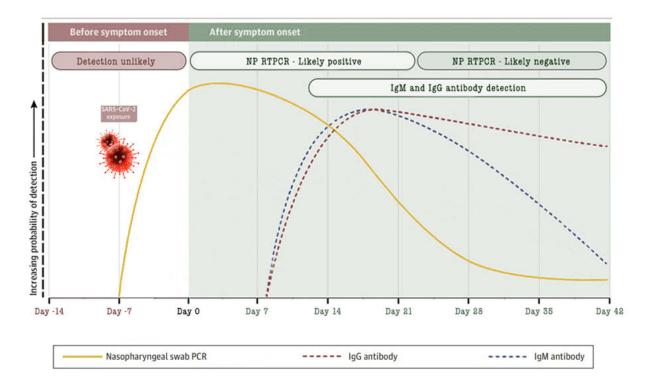
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133 FIGURES

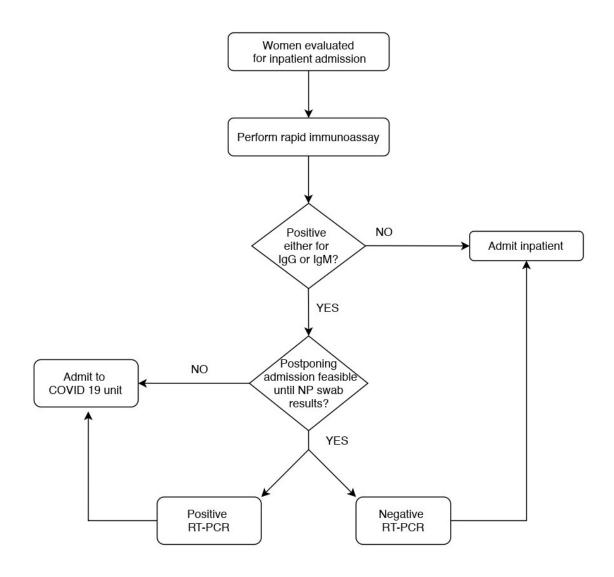
- Figure 1. Antibody response against SARS-COV-2 based on data from several published
 reports. SARS-CoV-2 indicates severe acute respiratory syndrome coronavirus 2; PCR,
 polymerase chain reaction. *Modified from Sethuraman N et al. Interpreting Diagnostic Tests for SARS-CoV-2. JAMA. 2020 May 6. doi: 10.1001/jama.2020.8259*Figure 2. Algorithm for rapid combined antibody test used at University of Naples Federico
- 139 II (Naples, Italy). A) For women before admission to inpatient monitoring; B) For women
- 140 scheduled for outpatient appointment. NP: nasopharyngeal; GP: general practitioner; RT-

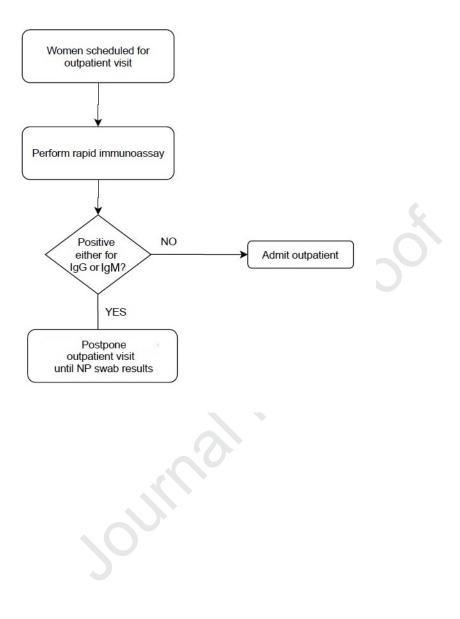
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141 PCR: real-time reverse-transcriptase-polymerase-chain-reaction



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I am an author on this submission, have adhered to all editorial policies for submission as described in the Information for Authors, attest to having met all authorship criteria, and all potential conflicts of interest / financial disclosures appears on the title page of the submission.

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