



Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.

# A case of gross hematuria and IgA nephropathy flare-up following SARS-CoV-2 vaccination



**To the editor:** We read with great interest the report of Negrea and Rovin of 2 cases of IgA nephropathy with gross hematuria following the Moderna vaccine for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2).<sup>1</sup> We also cared for a 52-year-old Asian female with prior biopsy-proven IgA nephropathy who developed gross hematuria within 24 hours of receiving a second dose of the Pfizer vaccine. Table 1 summarizes clinical data. Her workup was notable for proteinuria of 4.2 g/g of creatinine with serum creatinine at baseline. Of note, SARS-CoV-2 antibody testing prior to vaccination was negative, and she developed no symptoms after the first vaccine dose. Repeated testing within 1 week demonstrated resolution of hematuria and improving proteinuria. Interestingly, she developed gross hematuria following the first shot of the Shingrix vaccine 2 years prior but no symptoms following annual influenza vaccinations. The IgA nephropathy flare in our patient following the second SARS-CoV-2 vaccine dose without known prior exposure to SARS-CoV-2 suggests it was mediated by a delayed-type hypersensitivity reaction. Vasculitis flare-ups following vaccinations have been reported in the past.<sup>2,3</sup>

Our patient's symptoms improved within a week without any intervention aside from continued renin-angiotensin-aldosterone system blockade. It has been reported that severe coronavirus disease 2019 (COVID-19) illnesses can trigger an IgA response in the bronchial mucosa.<sup>4</sup> However, it is unclear how a nonmucosal vaccine triggers this response. We suggest that nephrologists closely follow their patients after COVID-19 vaccination to evaluate for varying degrees of flares, particularly after the second dose of an mRNA vaccine without prior exposure to SARS-CoV-2.

1. Negrea L, Rovin BH. Gross hematuria following vaccination for severe acute respiratory syndrome coronavirus 2 in 2 patients with IgA nephropathy. *Kidney Int.* 2021;99:1487.
2. Lambert EM, Liebling A, Glusac E, Antaya RJ. Henoch-Schönlein purpura following a meningococcal vaccine. *Pediatrics.* 2003;112:e491.
3. McNally A, McGregor D, Searle M, et al. Henoch-Schönlein purpura in a renal transplant recipient with prior IgA nephropathy following influenza vaccination. *Clin Kidney J.* 2013;6:313–315.
4. Hasan Ali O, Bomze D, Risch L, et al. Severe coronavirus disease 2019 (COVID-19) is associated with elevated serum immunoglobulin (Ig) A and antiphospholipid IgA antibodies [e-pub ahead of print]. *Clin Infect Dis.* <https://doi.org/10.1093/cid/ciaa1496>. Accessed September 20, 2020.

Shab E Gul Rahim<sup>1</sup>, Jonathan T. Lin<sup>1,2</sup> and John C. Wang<sup>1,2</sup>

<sup>1</sup>Division of Nephrology and Hypertension, Weill Cornell Medicine, New York, New York, USA; and <sup>2</sup>Rogosin Institute, New York, New York, USA

**Correspondence:** Shab E Gul Rahim, Division of Nephrology and Hypertension, Weill Cornell Medicine, 525 East 68th Street, New York, New York 10065, USA. E-mail: [SHABEGUL1@gmail.com](mailto:SHABEGUL1@gmail.com)

*Kidney International* (2021) **100**, 238; <https://doi.org/10.1016/j.kint.2021.04.024>

Copyright © 2021, International Society of Nephrology. Published by Elsevier Inc. All rights reserved.

# Acute rejection after anti-SARS-CoV-2 mRNA vaccination in a patient who underwent a kidney transplant

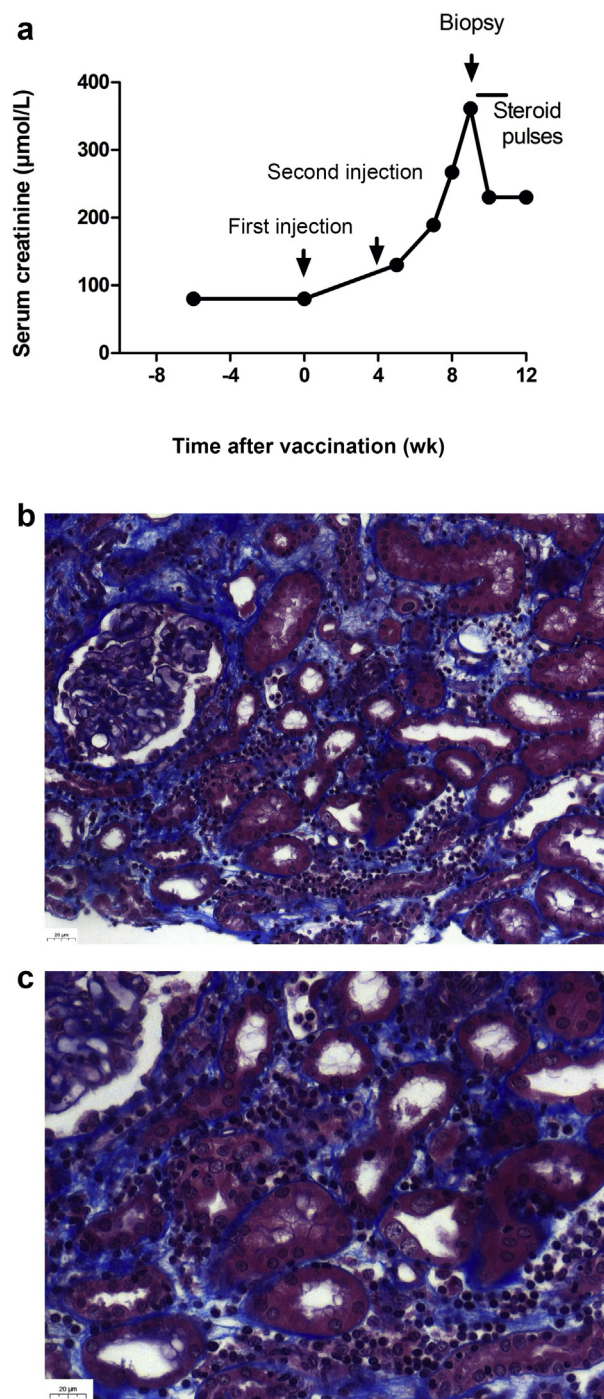


**To the editor:** Anti-severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) vaccination is recommended in patients who underwent a transplant because of an increased risk of developing severe coronavirus disease 2019 (COVID-19), and mortality.<sup>1</sup> Because of a weak immunogenicity of mRNA 2-dose vaccines in transplant patients, the French

**Table 1 | Patient symptoms and details of workup**

Patient characteristic	Data
Year of IgAN diagnosis	2017
Exacerbations since diagnosis	1. April 2019 following URI 2. June 2019 following shingles vaccine
Current treatment	Lisinopril
Baseline Cre	0.7–0.8 g/dl
Last urine microalbumin/Cre before exacerbation (2020)	633.1 mg/g Baseline always <1000 mg/g, except exacerbations
Urine microalbumin/Cre 48 h after Pfizer second dose	2411.3 mg/g
Gross hematuria/RBCs in urine	Yes/yes
Other symptoms	Fever, myalgias, body aches, lower back pain bilaterally
Urine microalbumin/Cre 5 d after Pfizer second dose	1441 mg/g
Hematuria 5 d after Pfizer second dose	Resolved

Cre, creatinine; IgAN, IgA nephropathy; RBC, red blood cell; URI, upper respiratory tract infection.



**Figure 1 | (a) Outcome of kidney function before and after transplantation and (b,c) kidney pathology.** Trichrome Masson staining exhibited inflammatory infiltration, tubulitis, edema, and peritubular capillaritis (original magnification  $\times 20$  [b] and  $\times 40$  [c]). Kidney biopsy was scored as follows, according to the Banff 2019 classification<sup>4</sup>: i2, t2, v0, g0, ptc1, ti1, i-IFTA0, C4d0, cg0, mm0, ah1, cv0, ci0, ct0. To optimize viewing of this image, please see the online version of this article at [www.kidney-international.org](http://www.kidney-international.org).

Health Authority recommended to offer a third dose to immunosuppressed patients to boost the immune response.<sup>2,3</sup> However, no biological monitoring before and after

vaccination is recommended. We report on the case of a 23-year-old non-human leukocyte antigen-sensitized patients who underwent a kidney transplant who presented an acute rejection after the second dose of the BNT162b2 mRNA COVID-19 Vaccine (Pfizer-BioNTech). She had undergone a deceased donor kidney transplantation for nephronophthisis 18 months earlier. The post-transplant period was uneventful. Her maintenance therapy was based on tacrolimus (target trough level, 5 ng/ml and 7 ng/ml), mycophenolic acid, and low-dose steroid. Fifteen days before the first dose, her serum creatinine level was at 80  $\mu\text{mol/L}$  and anti-SARS-CoV-2 serology was negative. Eight days after the second dose, systematic blood tests revealed impaired kidney function at 130  $\mu\text{mol/L}$ , which then raised to 360  $\mu\text{mol/L}$  (Figure 1). A kidney biopsy revealed a cellular acute rejection. Donor-specific anti-human leukocyte antigen antibodies became detectable with a weak intensity, targeting donor human leukocyte antigen class II antigens. Anti-SARS-CoV-2 spike protein antibodies became positive. Tacrolimus trough level was unchanged at 5 ng/ml. At 10 days, after steroid pulses (500 mg/d for 3 days), the patient's serum creatinine level had decreased to 230  $\mu\text{mol/L}$ . Another kidney biopsy is planned to discuss the use of polyclonal antibodies. Hence, this report suggests that kidney function should be carefully monitored in kidney transplantation undergoing anti-SARS-CoV-2 vaccination, especially if a third boost dose is performed.

1. Caillard S, Anglicheau D, Matignon M, et al. An initial report from the French SOT COVID Registry suggests high mortality due to COVID-19 in recipients of kidney transplants. *Kidney Int.* 2020;98:1549–1558.
2. Boyarsky BJ, Werbel WA, Avery RK, et al. Immunogenicity of a single dose of SARS-CoV-2 messenger RNA vaccine in solid organ transplant recipients. *JAMA.* 2021;325:1784–1786.
3. Centre opérationnel de régulation et de réponse aux urgences sanitaires et sociales. Vaccins contre la Covid-19: modalités d'administration des rappels. Available at: [https://www.mesvaccins.net/textes/dgs\\_urgent\\_n43\\_vaccination\\_modalites\\_d\\_administration\\_des\\_rappels.pdf](https://www.mesvaccins.net/textes/dgs_urgent_n43_vaccination_modalites_d_administration_des_rappels.pdf). Accessed April 11, 2021.
4. Mengel M, Loupy A, Haas M, et al. Banff 2019 Meeting Report: molecular diagnostics in solid organ transplantation-consensus for the Banff Human Organ Transplant (B-HOT) gene panel and open source multicenter validation. *Am J Transplant.* 2020;20:2305–2317.

Arnaud Del Bello<sup>1,2</sup>, Olivier Marion<sup>1,2,3</sup>, Audrey Delas<sup>4</sup>, Nicolas Congy-Jolivet<sup>3,5</sup>, Magali Colombat<sup>3,4</sup> and Nassim Kamar<sup>1,2,3</sup>

<sup>1</sup>Department of Nephrology and Organ Transplantation, Toulouse Rangueil University Hospital, Toulouse, France; <sup>2</sup>UMR1043, Center for Pathophysiology of Toulouse Purpan, Toulouse, France; <sup>3</sup>Paul Sabatier University, Toulouse, France; <sup>4</sup>Department of Pathology, Toulouse University Hospital, Toulouse, France; and <sup>5</sup>Department of Immunology, Toulouse University Hospital, Toulouse, France

**Correspondence:** Nassim Kamar, Department of Nephrology and Organ Transplantation, CHU Toulouse Rangueil, TSA 50032, 31059 Toulouse Cedex 9, France. E-mail: [kamar.n@chu-toulouse.fr](mailto:kamar.n@chu-toulouse.fr)

*Kidney International* (2021) **100**, 238–239; <https://doi.org/10.1016/j.kint.2021.04.025>

Copyright © 2021, International Society of Nephrology. Published by Elsevier Inc. All rights reserved.