## LETTER TO THE EDITOR

## Post-COVID-19 reactive arthritis

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Dear Editor,

We read with interest the published manuscript by Bekaryssova et al. [1] summarizing cases of post-COVID-19 reactive arthritis (ReA). In their manuscript, the authors started their review by discussing the definition of ReA and its progression over time. Then, they summarized cases of COVID-19 ReA and those of COVID-19 vaccinationinduced arthritis. They described clinical features, therapeutic management of ReA, and outcomes of these patients.

Twenty-two cases of COVID-19 ReA were summarized [1]. Several studies showed that COVID-19 infection is not only responsible for ReA but also viral arthritis. Thus, it is necessary to specify whether articular manifestations are due to ReA or viral arthritis.

Viral arthritis is characterized by an early onset and is commonly associated with symptoms of disseminated viral infection. This diagnosis remains possible in cases of early onset of articular symptoms. In this review, two cases of early arthritis associated with signs of disseminated viral infection were considered ReA [2, 3]. Indeed, Houshmand et al. described the case of a 10-year-old boy who presented with a concomitant fever, urticaria, and arthritis. Urticarial lesions tended to appear simultaneously with systemic viral symptoms. They have been reported in 11 to 19% of patients with COVID-19 infections [4]. These findings support the diagnosis of viral arthritis rather than post-COVID-19 ReA in this patient. The post-COVID-19 ReA diagnosis can be

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established based on the occurrence of rheumatological manifestations with a delay between the onset of these manifestations and the confirmed diagnosis of COVID-19 and nasopharyngeal real-time polymerase chain reaction (RT-PCR) negativation at the time of articular signs onset [5]. It is important to emphasize that RT-PCR positivity can persist in patients who have already been in remission [6].

In this review, the authors did not specify the delay between the COVID-19 infection diagnosis and articular manifestations onset [1].

Besides, the authors did not describe the treatments received during the COVID-19 infection [1]. Indeed, articular manifestations can be triggered by several drugs prescribed during COVID-19 infection. Favipiravir may induce hyperuricemia and even acute gouty arthritis [7]. Likewise, interferon  $\beta$ 1 treatment may be responsible for inflammatory rheumatic disease occurrence [8, 9].

There is no agreement on the ReA criteria. Therefore, we believe that ruling out differential diagnoses such as viral arthritis related to COVID-19, drug-induced arthritis, crystal-induced flares, or a recent onset of another inflammatory rheumatic disease is necessary before making the ReA diagnosis [10].

The authors included seven cases of ReA after COVID-19 vaccination with corona VAC, Sinovac, and SPUTNIK-V.

While some authors consider joint manifestations induced by anti-COVID vaccination a form of ReA, others consider it an autoimmune or inflammatory syndrome induced by adjuvants. Adjuvants are components added to subunit and inactivated vaccines to stimulate innate and adaptive immune responses. This stimulation increases the effectiveness of subunit and inactivated vaccines, but it can be responsible for autoimmune/inflammatory syndrome induced by adjuvant (ASIA) [11]. Diagnosis criteria have been suggested for ASIA. The first major criterion is exposure to external stimuli such as a vaccine, and the second is the appearance of typical clinical manifestations such as arthritis [11]. These manifestations typically occur 2 to 5 days after COVID-19 vaccination [12]. The detection of autoantibodies or antiadjuvant antibodies is possible and considered a minor



diagnosis criterion [12]. It is not clear whether articular manifestations occurring after COVID-19 vaccination correspond to ReA or ASIA [11].

ReA post-COVID-19 infection or post-COVID-19-vaccination is a matter of concern that must support increased research into the pathophysiology of ReA.

## Declarations

Disclosures None.

## References

- Bekaryssova D, Yessirkepov M, Zimba O, Gasparyan AY, Ahmed S (2022) Reactive arthritis before and after the onset of the COVID-19 pandemic. Clin Rheumatol 41(6):1641–1652
- Talarico R, Stagnaro C, Ferro F, Carli L, Mosca M (2022) Symmetric peripheral polyarthritis developed during SARS-CoV-2 infection. Lancet Rheumatol 2(9):e518–e519
- 3. Houshmand H, Abounoori M, Ghaemi R, Bayat S, Houshmand G (2020) Ten-year-old boy with atypical COVID-19 symptom presentation: a case report. Clin Case Rep 9(1):304–308
- 4. Genovese G, Moltrasio C, Berti E, Marzano AV (2020) Skin manifestations associated with COVID-19: current knowledge and future perspectives. Dermatol Basel Switz 237(1):1–12
- 5. Wendling D, Verhoeven F, Chouk M, Prati C (2021) Can SARS-CoV-2 trigger reactive arthritis? Joint Bone Spine 88(1):105086

- Ruiz-Galiana J, De Lucas RP, García-Botella A, García-Lledó A, Gómez-Pavón J, González Del Castillo J et al (2022) Persistence and viability of SARS-CoV-2 in primary infection and reinfections. Rev Espanola Quimioter Publicacion Of Soc Espanola Quimioter 35(1):1–6
- Hase R, Kurata R, Ishida K, Kurita T, Muranaka E, Mito H (2020) Acute gouty arthritis during favipiravir treatment for coronavirus disease 2019. Intern Med Tokyo Jpn 59(18):2327–2329
- 8. Hojjati SMM, Heidari B, Babaei M (2016) Development of rheumatoid arthritis during treatment of multiple sclerosis with interferon beta 1-a. Coincidence of two conditions or a complication of treatment: a case report. J Adv Res 7(5):611–613
- Toussirot E, Béreau M, Bossert M, Malkoun I, Lohse A (2014) Occurrence of psoriatic arthritis during interferon beta 1a treatment for multiple sclerosis. Case Rep Rheumatol 2014:949317
- Slouma M, Abbes M, Mehmli T, Dhahri R, Metoui L, Gharsallah I, Louzir B (2022) Reactive arthritis occurring after COVID-19 infection: a narrative review. Infection. https://doi.org/10.1007/ s15010-022-01858-z
- Perricone C, Colafrancesco S, Mazor RD, Soriano A, Agmon-Levin N, Shoenfeld Y (2013) Autoimmune/inflammatory syndrome induced by adjuvants (ASIA) 2013: unveiling the pathogenic, clinical and diagnostic aspects. J Autoimmun 47:1–16
- Jara LJ, Vera-Lastra O, Mahroum N, Pineda C, Shoenfeld Y (2022) Autoimmune post-COVID vaccine syndromes: does the spectrum of autoimmune/inflammatory syndrome expand? Clin Rheumatol 41(5):1603–1609

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