



# Case Report SARS-CoV-2-Associated Obliterative Arteritis Causing Massive Testicular Infarction

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**Abstract:** A 26-year-old man with symptomatic SARS-CoV-2 infection developed a sudden-onset acute testicular pain. The echo-doppler images showed massive testicular infarction, so orchiectomy was performed. On gross examination, the surgical specimen showed complete testicular necrosis and diffuse thickening of the testicular coverings. Under the microscope, a severe obliterative arteritis was evidenced. SARS-CoV-2 spike antibody was detected by immunohistochemistry in the arterial endothelium. Electron microscopy displayed intracytoplasmic spiky viral particles in endothelial cells. The patient was treated with corticoids and was asymptomatic at last contact.

Keywords: SARS-CoV-2; COVID-19; arteritis; pathology; testicular infarction



Citation: Parrón, D.; Gartzia, A.; Iturregui, A.M.; Imaz, I.; Manini, C.; García-Olaverri, J.; López, J.I. SARS-CoV-2-Associated Obliterative Arteritis Causing Massive Testicular Infarction. *Clin. Pract.* **2021**, *11*, 246–249. https://doi.org/10.3390/ clinpract11020037

Received: 19 March 2021 Accepted: 21 April 2021 Published: 6 May 2021

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## 1. Introduction

Individual host reactivity against SARS-CoV-2 is largely unpredictable and responsible for most symptoms and sequelae of this infection. Vasculitis is a systemic response to the virus with variable qualitative and quantitative clinical manifestations. Here, we present an extreme case of massive unilateral testicular necrosis occurring in the recovery phase of SARS-CoV-2 infection in a young man, a manifestation which represents a previously non reported clinical complication of this infection.

# 2. Case Report

A 26-year-old man with symptomatic SARS-CoV-2 infection, including mild dyspnea and fever, developed a left testicular pain two days after the acute symptomatic infection was vanished (PCR negative, IgG positive). No antecedents of any systemic disease were recorded in the anamnesis and analytical data, other than those specifically related to the SARS-CoV-2, were within the normal limits. No signs of coagulopathy were detected. The echo-doppler study of the testis at that time showed mild hypoperfusion. Testicular pain was then thought to be secondary to orchitis and symptomatic therapy with empiric antibiotic coverage was started. Pain, however, did not recover and 3 days after a second echo-doppler showed a complete testicular infarction (Figure 1A). Left orchiectomy was then performed (Figure 1B). No signs or symptoms of vasculopathy were detected elsewhere. The patient was treated with corticoids and was asymptomatic at last contact. Right testis is not affected.

Grossly, the testicle (length of the surgical specimen: 8 cm, testicular size: 3 cm, surgical specimen weight: 73 g) showed a massive infarction and diffuse thickening of the spermatic cord and testicular coverings (Figure 1C). A segmentary obliterative endotheliitis affecting large, medium, and small-sized arteries was observed in the spermatic cord. Vascular injury was heterogeneous in intensity and distribution. The more severely damaged vessels, including the spermatic artery, displayed occlusive endothelial proliferation (Figure 2A) admixed with neutrophils, eosinophils, and lymphocytes. Less damaged vessels showed endothelial tumefaction and vacuolization with associated inflammation (Figure 2B–D). Additionally, there were well-preserved arteries and veins without any apparent morphological alteration. Vascular lumina showed only very occasional thrombotic phenomena.



**Figure 1.** Echo-doppler of the left testicle shows diffuse ischemic infarction (**A**); left orchiectomy specimen (**B**) showing complete testicular necrosis and thickening of the spermatic cord and peritesticular coverings (**C**).



**Figure 2.** Spermatic cord arteries show temporal heterogeneity of the lesions, from severe luminal occlusion ((**A**), original magnification,  $\times$ 40) to mild endotheliitis with luminal polymorphonuclear elements ((**B**),  $\times$ 100) and endothelial tumefaction ((**C**),  $\times$ 640). Moderate lesions showed endothelial thickening with mixed inflammatory infiltrates ((**D**),  $\times$ 240). Positive immunostaining with SARS-CoV-2 spike antibody was detected in endothelial cells of arteries (((**E**),  $\times$ 400) with a negative control in a vein of the same paraffin block ((**F**),  $\times$ 400)).

The intravascular lymphocyte population associated with endothelial damage included both CD4 and CD8 cells. Interstitial and perivascular infiltrates were composed of a mixture of macrophages, and T and B cells. Immunohistochemistry against the SARS-CoV-2 spike protein (Gene Tex, clone 1A9, dilution 1:500) showed positive endothelial reaction (Figure 2E, negative control in Figure 2F). Moreover, viral spiked particles were detected in endothelial cells (Figure 3). Testicular parenchyma was diffusely necrotic.



Figure 3. Spiked viral particles were identified within the cytoplasm of endothelial cells.

### 3. Discussion

COVID-19-associated vasculopathy has been well documented in almost any organ [1] being responsible for most of the symptoms and sequelae of this infection. Direct endothelial damage induced by the virus alters the in situ glycocalyx thus favoring hypercoagulability [2]. Diffuse alveolar damage, interstitial, and organizing pneumonia have been observed in the lungs of several autopsy studies [3]. However, almost every organ can be affected. Some other similar examples of ischemic changes secondary to virus-related vasculopathy, i.e., ischemic vascular stroke [4], intestinal ischemia [5], and gallbladder ischemic necrosis [6], some of them occurring in young people, have been occasionally reported. Orchitis and impaired spermatogenesis have been previously associated with COVID-19 infection [7,8]. However, massive testicular ischemic infarction in the context of COVID-19 infection has not been reported so far.

The present case recalls the attention of general practitioners and urologists when facing testicular pain in the context of SARS-CoV-2 infection.

**Author Contributions:** D.P. and A.G.: Sampling selection and histological analysis; A.M.I. and J.G.-O.: surgery, patient care and follow-up; I.I., C.M., J.I.L.: histological and electron microscopy analysis; J.I.L.: histopathological diagnosis and manuscript writing. All authors have read and agreed to the published version of the manuscript.

Funding: This research received no external funding.

Institutional Review Board Statement: Not applicable.

Informed Consent Statement: Informed consent was obtained from the patient.

Conflicts of Interest: The authors declare no conflict of interest.

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