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## Letter to the Editor Regarding "Pulmonary Embolism Prophylaxis in Patients With COVID-19: an Emerging Issue", Heart Lung Circ. 2021;30(10):1435–41.



## To the Editor,

I read the article by Sanidas and colleagues [1] with great interest and congratulate the authors on their excellent work. However, I would like to draw attention to several points. Recent findings suggest that SARS-CoV-2 controls the expression of host regulatory molecules such as miRNAs and transcription factors (TFs), often regulating gene expression at transcriptional and post-transcriptional levels and forming highly complex network interactions [2]. Research has identified four miRNAs (hsa-mir-9-5p, hsa-mir-324-3p, hsa-mir-1827, and hsa-mir-1277-5p) and TFs (EP300, MYC, E2F1, and SP1) that co-regulate important target genes responsible for SARS-CoV-2 host invasion [2]. Studies have found evidence supporting common molecular trajectories of SARS-CoV-2 with clotting, pulmonary embolism, pulmonary oedema, and systematic inflammation, showing both fibrinolysis as a major pathway leading to D-dimer increase and cytokines TGF- $\beta$  and TNF- $\alpha$  as major regulators of fibrinolysis controlling proteins PAI-1 and plasminogen activators also highlighting the role of hsa-mir-9-5p, hsa-mir-324-3p, EP300, and SP1 on pulmonary embolism [2]. It has also been reported that in the circulating exosomes of high D-dimer COVID-19 patients (cut-off 3 µg/mL) there is significant miR-424 upregulation and miR-103a, miR-145, and miR-885 downregulation. Exosomal miR-424 is an independent thromboembolic event predictor in COVID-19 patients [3,4], whereas miR-103a independently regulates D-dimer levels [4]. Furthermore, miR-885 targets von Willebrand factor and miR-145 targets tissue factor [3,4], showing negative correlations with their respective targets [3]. The findings of Sanidas and colleagues [1] add significant information to previously published data; however, also evaluating these aspects would be useful for better understanding of the interplay between pulmonary embolism in patients with COVID-19 and its complex regulatory network.

## **Conflict of Interest**

None declared.

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