

An Acad Bras Cienc (2020) 92(4): e20201080 DOI 10.1590/0001-3765202020201080 Anais da Academia Brasileira de Ciências | Annals of the Brazilian Academy of Sciences Printed ISSN 0001-3765 I Online ISSN 1678-2690 www.scielo.br/aabc | www.fb.com/aabcjournal

LETTER TO THE EDITOR

Cardiac glycosides and COVID-19: would it be a promising therapeutic approach?

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Abstract: Cardiac glycosides have been found to have an anti-viral effect. This was noted in the past during various epidemics including MERS and SARS. It is due to their inhibitory effect on the Na, K-ATPase membrane pump. Furthermore, they exhibit anti-inflammatory properties. These preclinical observations may prove useful in further clinical utility of these well-known compounds in the current COVID-19 pandemic.

Key words: antiviral effect, cardiac glycosides, COVID-19.

In their recent article Trindade et al. (2020) propose some therapeutic approaches to COVID-19 pandemic, related to the SARS-CoV-2 coronavirus.

Cardiac glycosides (CGS) digoxin and digitoxin merit a privileged position among the antiviral compounds. CGS inhibit the Na, K-ATPase membrane pump, a potent signal transduction pathway and ion-exchanger (Riganti et al. 2011). RNA viruses use the Na, K-ATPase pump, in order to attach to the host cell membrane and proceed in endocytosis and replication (Amarelle & Lecuona 2018). CGS were found in the course of previous viral epidemics including MERS and SARS, to inhibit the signaling properties of the pump, and provoke a host intracellular depletion of potassium (Burkard et al. 2015, Grosso et al. 2017). By these mechanisms, CGS inhibited viral transmembrane internalization and viral RNA and pre-RNA splicing necessary for the synthesis of viral proteins, interrupting the replication process (Amarelle et al. 2019). Furthermore, CGS possess significant anti-inflammatory properties, in vitro and in vivo, when administered in the course of inflammatory diseases. CGS downregulate various cytokines and activators, such as TNFα, TGFβ and NF-KB ((Yang et al. 2005, Prassas & Diamandis 2008).

Given that COVID-19 is characterized by catastrophic proinflammatory cytokine storm in the target organs, an anti-inflammatory action of CGS provokes further investigation. CGS constitute a historical and emblematic treatment in heart failure and supraventricular arrhythmias. Should preclinical observations regarding the antiviral properties of CGS be translated to clinical validation, a new and promising COVID-19 therapeutic approach would emerge.

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How to cite

SINIORAKIS E, ARVANITAKIS S & ELKOURIS M. 2020. Cardiac glycosides and COVID-19: would it be a promising therapeutic approach?. An Acad Bras Cienc 92: e20201080. DOI 10.1590/0001-3765202020201080.

Manuscript received on July 8, 2020; accepted for publication on September 10, 2020

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Eftychios Siniorakis contributed to the conception, data acquisition and discussion. Spyridon Arvanitakis contributed to the critical manuscript review. Maximilianos Elkouris contributed to data acquisition and discussion. All authors gave their final approval and agree with all aspects of the work.

