Perspective article

2 COVID-19 through the One Health lens: adding a missing perspective

3

1

4 Christian Selbach^{1*}, Maarten P. M. Vanhove², Kim N. Mouritsen¹

5

- 6 Department of Biology, Aquatic Biology, Aarhus University, Aarhus, Denmark
- 7 Research Group Zoology: Biodiversity and Toxicology, Centre for Environmental Sciences,
- 8 Hasselt University, Diepenbeek, Belgium

9

* Corresponding author: christian.selbach@bios.au.dk

11

12

Abstract

- 13 The One Health concept offers an integrative approach to disease and health at the human-
- animal-environment interface. It has often been suggested to view the COVID-19 outbreak
- within this framework to better understand and mitigate this global crisis. Here, we discuss how
- the evolutionary ecology of host-pathogen systems can add a valuable additional perspective to
- the debate around SARS-CoV-2 and its implications for public health awareness and policy-
- 18 making. In this context, it is especially important to highlight that changes in nature, such as
- 20 zoonotic spillover events, are often irreversible, and that humans, while deeply embedded in
- 20 ecosystems, are intricate ecosystems themselves. A better recognition of the complex biology
- 21 and evolution of human-parasite interactions will assist our understanding of such zoonoses.

22

23 **Keywords:** One Health, COVID-19, spillover, host-parasite coevolution

24

25

Main text

Undoubtedly, the current COVID-19 pandemic is one of the greatest health crises humanity has faced. To better understand and mitigate this global crisis, and to be better prepared for similar epidemics in the future, it has been suggested to view the COVID-19 outbreak within the One Health (OH) framework (e.g. [1]). The OH concept aims at achieving optimal health for people, non-human organisms, and the environment via inter- and transdisciplinary collaborations across health and environmental sciences [2].

Given the zoonotic origin of SARS-CoV-2 that spilled over to humans from wildlife, such an approach is certainly warranted. And without doubt, linking human, animal and ecosystem health within the OH framework is a powerful approach in predicting, tackling and preventing disease outbreaks [3]. However, even though OH and related concepts such as EcoHealth have expanded beyond an initial narrow focus on human and veterinary health and now include wider ecological perspectives [2], we find that the framing of the COVID-19 pandemic within OH misses a crucial perspective: the evolutionary ecology of this host-parasite system. Rather, in the literature, the stated importance of OH in the COVID-19 context often seems limited to recognizing and detecting zoonotic origins, understanding the human-animal-environment interface, and avoiding the circumstances of transmission. Hence, integrative approaches appear especially useful in preventing future zoonotic epidemics, while less attention is given to their potential for dealing with the ongoing pandemic.

Although vaccination efforts will likely be a great success in tackling the pandemic [4], the virus and COVID-19 are expected to persist, potentially with regular seasonal outbreaks occurring during the coming years [5]. SARS-CoV-2 has therefore become, and will in the foreseeable future remain, part of the human pathobiome [6]. From the viruses' point of view, the human host represents a resource-rich ecosystem that the virus has spilled over to and inhabits, and in which it thrives, replicates and evolves [7]. The emergence of new SARS-CoV-2 variants carrying different mutations highlights the pathogen's rapid evolution within its

human host system [8]. This should caution against any hopes of returning to the *status quo* ante and 'back to normal', once this crisis has been overcome. In evolutionary processes, there is no going back, and a solution that aims at restoring a pre-outbreak situation simply does not exist. Accordingly, the 'new normal' [9] in social contexts comes along with a new biological normality in a long-term host-parasite association with continuous coevolution.

Moreover, after the initial wildlife-to-human transmission of SARS-CoV-2, the virus has continued to spill over to other mammals, most notably American mink *Neovison vison* in Danish and Dutch fur farms, and it has the potential to infect a wide range of domesticated and wild animals [10]. Having been co-introduced around the world by its cosmopolitan human host, there is little reason to assume that no further cross-species transmission events will occur in the future. Such spillovers are largely driven by increased exposure events and the acquisition of genetic variations that allow host switching [11], both of which are common features of the current outbreak. It is therefore well likely that the human host system can serve as a 'stepping stone' for the virus to find its way into yet other host species, which might be the starting point of an ever-changing host range of SARS-CoV-2 throughout its evolutionary history. To fully understand and predict the risks this entails, input from ecological and evolutionary parasitology that focuses on host-pathogen interaction will be crucial.

Considering human, animal and ecosystem health together under the OH umbrella can greatly benefit our understanding of zoonotic diseases, but will require investigating them as novel and potentially persistent host-parasite systems including an eco-evolutionary perspective. In this context, it is especially important to point out that changes in nature, such as zoonotic spillover events, are often irreversible, and that humans, while deeply embedded in ecosystems, are complex ecosystems themselves. Human health, rather than a state of well-being with the mere absence of disease, encompasses the ability to adapt and self-manage in the face of physical, social, and emotional challenges [12]. A better recognition and public awareness of the complex biology and evolution of human-parasite interactions could help our

78	understanding of this pandemic, assist the public in framing often poorly known phenomena
79	like viral mutations, zoonotic spillovers or biological invasions [13], and lead to more
80	autonomous and responsible behaviour in light of current and future health challenges.
81	
82	
83	Acknowledgements
84	This work received funding from the European Union's Horizon 2020 Research and Innovation
85	Programme under the Marie Skłodowska-Curie grant agreement No. 839635 TPOINT (C.S.).
86	M.P.M.V. is supported by the Special Research Fund of Hasselt University (BOF20TT06).
87	
88	Author contributions
89	C.S. conceived the paper and led the writing with primary inputs from M.P.M.V. and K.N.M.
90	
91	Corresponding author
92	Correspondence to christian.selbach@bios.au.dk
93	
94	Competing interests
95	The authors declare no competing interests.
96	
97	References
98	1. Amuasi JH, Walzer C, Heymann D, Carabin H, Huong LT, Haines A, et al. Calling for a
99	COVID-19 One Health Research Coalition. Lancet. 2020;395(10236):1543-4.
100	2. Lerner H, Berg C. A comparison of three holistic approaches to health: One health, ecohealth,
101	and planetary health. Front Vet Sci. 2017;4:1-7.

- 3. Antoine-Moussiaux N, Janssens de Bisthoven L, Leyens S, Assmuth T, Keune H, Jakob Z,
- et al. The good, the bad and the ugly: framing debates on nature in a One Health
- 104 community. Sustain Sci. 2019;14(6):1729–38.
- 4. Scudellari M. How the pandemic might play out in 2021 and beyond. Nature.
- 106 2020;584(7819):22–5.
- 5. Kissler SM, Tedijanto C, Goldstein E, Grad YH, Lipsitch M. Projecting the transmission
- dynamics of SARS-CoV-2 through the postpandemic period. Science.
- 109 2020;368(6493):860–8.
- 6. Bass D, Stentiford GD, Wang H-C, Koskella B, Tyler CR. The Pathobiome in Animal and
- Plant Diseases. Trends Ecol Evol. 2019;34(11):996–1008.
- 7. Rynkiewicz EC, Pedersen AB, Fenton A. An ecosystem approach to understanding and
- managing within-host parasite community dynamics. Trends Parasitol. 2015;31(5):212–
- 114 21.
- 8. Greaney AJ, Loes AN, Crawford KHD, Starr TN, Malone KD, Chu HY, et al. PREPRINT:
- 116 Comprehensive mapping of mutations to the SARS-CoV-2 receptor-binding domain that
- affect recognition by polyclonal human serum antibodies. bioRxiv.
- 2021;2020.12.31.425021. DOI: https://doi.org/10.1101/2020.12.31.425021
- 9. World Health Organization [Internet]. COVID-19: 'new normal'. [cited 2021 Feb 18].
- 120 Available from: https://www.who.int/westernpacific/emergencies/covid-
- 121 <u>19/information/covid-19-new-normal</u>.
- 10. Gryseels S, De Bruyn L, Gyselings R, Calvignac-Spencer S, Leendertz FH, Leirs H. Risk
- of human-to-wildlife transmission of SARS-CoV-2. Mamm Rev. 2020;mam.12225.
- 11. Parrish CR, Holmes EC, Morens DM, Park E-C, Burke DS, Calisher CH, et al. Cross-
- Species Virus Transmission and the Emergence of New Epidemic Diseases. Microbiol
- 126 Mol Biol Rev. 2008;72(3):457–70.

- 12. Huber M, Knottnerus JA, Green L, Horst H v. d., Jadad AR, Kromhout D, et al. How should
- we define health? BMJ. 2011;343(2):d4163–d4163.
- 13. Nuñez MA, Pauchard A, Ricciardi A. Invasion Science and the Global Spread of SARS-
- 130 CoV-2. Trends Ecol Evol. 2020;35(8):642–5.