

Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.





ScienceDirect



Editorial overview: Special issue on antiviral strategies in *Current Opinion in Virology* Richard K Plemper



Current Opinion in Virology 2021, 50:95–96 For complete overview about the section, refer "Engineering for viral resistance"

Available online 20th August 2021

https://doi.org/10.1016/j.coviro.2021.07.008

1879-6257/© 2021 Elsevier B.V. All rights reserved.

Richard K Plemper



Center for Translational Antiviral Research, Georgia State University, Atlanta, GA, USA e-mail: rplemper@gsu.edu

Richard K. Plemper is the Director of the Center for Translational Antiviral Research and Distinguished University Professor at Georgia State University. His research focuses on the biology of respiratory RNA viruses with particular emphasis on developing direct-acting small-molecule therapeutics targeting the entry and replication machineries of viral pathogens of the paramyxovirus, orthomyxovirus, and coronavirus families. He has made numerous contributions to developing novel drug screening technology, isolating druggable targets, and identifying and characterizing novel antiviral candidates directed against respiratory syncytial virus, parainfluenzaviruses, and influenza viruses. His laboratory was instrumental in the development of EIDD-2801/molnupiravir against influenza and the subsequent repurposing of the drug for the treatment of SARS-coV-2 infection.

The COVID-19 pandemic has resulted in approximately 200 million cases and an estimated 4 million deaths worldwide to date. The impact on global health, economies, and politics is unprecedented for a single viral pathogen. Since the first reports of the emergence of a novel betacoronavirus appeared in 2019 and a global pandemic was declared in early 2020, SARS-CoV-2 has created a watershed moment also for the research community in terms of unparalleled need for effective therapeutics and vaccines, the goal of dramatically shortened timelines to advance drug candidates to the clinic, and in meeting public expectations. Exciting new technologies have been developed and innovative collaborative initiatives launched, but also lessons of pitfalls in drug discovery relearned.

This year's annual issue of Antiviral Strategies in *Current Opinion in Virology* focusses exclusively on the SARS-CoV-2 challenge. Opinions on what constitutes the most effective response strategy to the still raging pandemic vary widely and it may take years until the individual impact of different approaches can be fully appreciated. At the current phase from widespread SARS-CoV-2 community transmission to increased vaccination coverage, reopening of large geographical areas, and the emergence of a number of viral variants of concern, this issue has collected the insight of subject experts in an attempt to provide an early retrospective.

Opinion articles herein address three broad areas of COVID-19 response research: i) foundational studies of SARS-CoV-2 biology; ii) drug discovery, development, and repurposing research; and iii) the development of novel vaccine technologies.

- Specifically, contributing authors Lowen [1] and de Swart [2] and their coauthors discuss animal models developed for studying SARS-CoV-2 pathogenesis and transmission, respectively, which are instrumental for preclinical stage drug and vaccine development. Cox *et al.* [3] examine the importance of protein structural information to the identification of druggable targets and structure-guided drug optimization, whereas Gotte [4] evaluate the impact of *in vitro* biochemical assays on drug development by example of the mechanistic characterization of remdesivir. Lee and coauthors (insert LINK) summarize current SARS-CoV-2 entry assay technology and illuminate the importance of standardized assay conditions for reproducible results.
- In two articles directed at drug discovery strategies, Jonsson [5] and Sourimant [6] and colleagues assess the effectiveness of the many open drug discovery and drug repurposing screens that were launched in 2020 in response to the mounting pandemic. Focusing on specific

druggable targets for direct-acting antivirals, Painter [7] and Deval [8] and co-authors recapitulate the development of EIDD-2801/molnupiravir and SARS-CoV-2 protease inhibitors, respectively. Ngo and Gewirtz [9] provide a broader view on the effect of the host microbiome on COVID-19 progression, whereas Glazier and colleagues (insert LINK) discuss the predictive power of *in silico* modeling to appreciate the dynamic interplay between virus and host.

• On COVID-19 prophylaxis, Kurup and Schnell [10] examine vaccine technologies considered to prevent SARS-CoV-2 infection. This overview is complemented by a discussion of the specific strengths and challenges of mRNA nanoparticle vaccines by Igyarto *et al.* [11]. Summarizing the unprecedented expansion of diagnostic capacity during the pandemic, Greninger [12] discusses the impact of newly established infrastructure on surveillance and our ability to gauge therapeutic success.

COVID-19 was certainly not the last pandemic viral threat to human health. Assessing the effectiveness of response research strategies will be important to improve pandemic preparedness. Ideally, the compendium of opinion articles forming this special issue will contribute to identifying approaches that have meaningfully contributed towards this goal.

Acknowledgement

This work was supported, in part, by Public Health Service grant AI141222 (redirected) from the NIH/NIAID. The funders had no role in data collection and interpretation, or the decision to submit the work for publication.

References

- 1. Lee CY, Lowen AC: Animal models for SARS-CoV-2. Curr Opin Virol 2021, 48:73-81.
- de Vries RD, Rockx B, Haagmans BL, Herfst S, Koopmans MP et al.: Animal models of SARS-CoV-2 transmission. Curr Opin Virol 2021, 50:8-16.
- Cox RM, Plemper RK: The impact of high-resolution structural data on stemming the COVID-19 pandemic. Curr Opin Virol 2021, 49:127-138.
- 4. Gotte M: Remdesivir for the treatment of Covid-19: the value of biochemical studies. *Curr Opin Virol* 2021, **49**:81-85.
- Jonsson CB, Golden JE, Meibohm B: Time to' Mind the Gap' in novel small molecule drug discovery for direct-acting antivirals for SARS-CoV-2. Curr Opin Virol 2021, 50:1-7.
- Sourimant J, Aggarwal M, Plemper RK: Progress and pitfalls of a year of drug repurposing screens against COVID-19. Curr Opin Virol 2021, 49:183-193.
- Painter GR, Natchus MG, Cohen O, Holman W, Painter WP: Developing a direct acting, orally available antiviral agent in a pandemic: the evolution of molnupiravir as a potential treatment for COVID-19. Curr Opin Virol 2021, 50:17-22.
- Vandyck K, Deval J: Considerations for the discovery and development of 3-chymotrypsin-like cysteine protease inhibitors targeting SARS-CoV-2 infection. *Curr Opin Virol* 2021, 49:36-40.
- 9. Ngo VL, Gewirtz AT: Microbiota as a potentially-modifiable factor influencing COVID-19. *Curr Opin Virol* 2021, 49:21-26.
- Kurup D, Schnell MJ: SARS-CoV-2 vaccines the biggest medical research project of the 21st century. Curr Opin Virol 2021, 49:52-57.
- Igyarto BZ, Jacobsen S, Ndeupen S: Future considerations for the mRNA-lipid nanoparticle vaccine platform. Curr Opin Virol 2021, 48:65-72.
- 12. Greninger AL: Test it earlier, result it faster, makes us stronger: how rapid viral diagnostics enable therapeutic success. Curr Opin Virol 2021, 49:111-116.