



Frontiers in Virology: An Innovative Platform for Integrative Virus Research

Akio Adachi*

Department of Microbiology, Kansai Medical University, Osaka, Japan

Keywords: virology, biology, naturally occurring viruses, pathogenic viruses, emerging viruses, viral diseases, pandemic

Since the first discovery of a plant virus in 19th century, numerous viruses have been found and identified as admirably unique genetic agents in virtually all unicellular and multicellular living creatures (1–3) (Figure 1). They are parasitic entities that replicate in a remarkably different manner in their specific hosts. Owing to extensive and enthusiastic efforts to date by numbers of scientists in diverse research fields, viruses are now recognized to be the most abundant replicating species in nature. However, naturally occurring viruses are found to be beyond the scope of our expectations. Each viral species has its own unique strategy to interact with its host species for survival and frequently gives unexpected results upon its infection, representing a wide variety of scientifically important biological issues to be addressed and solved (1, 4-26). Furthermore, new dreadful pathogenic viruses with distinct characteristics keep emerging or reemerging in various parts of the world, generating significant menaces to humans (4, 10, 27-34). It is thus increasingly important to systemically explore viruses in detail particularly in this challenging time (29, 32, 34-36) (Figure 1). Viral infectious diseases are certainly one of the most critical issues in medical, environmental, ecological, and health sciences. Extensive efforts need to be made against these pathogenic viruses. Today's virology, at a global level, aims at understanding practical aspects of viruses as well as elucidating relevant biological basic matters (Figure 1).

OPEN ACCESS

Edited and reviewed by: Mario Mago Clerici, University of Milan, Italy

*Correspondence:

Akio Adachi adachi@tokushima-u.ac.jp; adachiak@hirakata.kmu.ac.jp

Received: 08 February 2021 Accepted: 12 February 2021 Published: 25 March 2021

Citation:

Adachi A (2021) Frontiers in Virology: An Innovative Platform for Integrative Virus Research. Front. Virol. 1:665473. doi: 10.3389/fviro.2021.665473

Virology has gradually developed to the present multidisciplinary status from the initial stage 130 years ago. During the period, virology clearly has been acting as strong driving forces to make paradigm shifts for basic biology and molecular biology (1-3), and also for applied sciences of various fields including nanotechnology, virus vectors, and medical therapies (37-39). Viruses are the smallest and most simple organic molecules with highly ordered structures that can replicate as a population under some specific conditions. On one hand, viruses are sometimes pathogenic/fatal for their hosts and sometimes transmittable among different host species. Targeting and taking the advantage of these characteristic viruses, a number of highly memorable investigations on biology and medicine, Nobel prize and Nobel prize-worthy studies, have been performed as wellknown. Invaluable studies on various viruses are currently in progress all over the world. Needless to mention, the development and success of current virology have been critically supported by the innovative next-generation methodologies/technologies recently available. These include the forefront systems, such as bioinformatics, modellings, and sequencings, to sensitively and precisely analyze the concerned issues (3, 18, 40-56) (Figure 1). Given the multi-faceted perspective and inherent thoroughness, results obtained by these methods would readily point out the next projects to be done. We virologists can now predict, albeit to a certain degree, how viruses behave in response to environmental changes and are able to verify the predicted outcomes in an open and demonstrative way by well-designed experiments (1-3). Virology should spread its wings to be more scientifically general and inspirational, and should also continue to be a major academic discipline of the biology.





2

Frontiers in Virology actively reports all the scientifically critical investigations/efforts relevant to all kinds of viruses in nature (Figure 1). It publishes high-quality studies on all species and topics of viruses in one journal in an integrative manner. It surely impacts both academic and public communities by eminent methodologies/technologies used for studies as well as crucial scientific contents reported. On one hand, as repeatedly emphasized and stated above, viruses are beyond the scope of our expectations. We virologists must keep exploring the nature of viruses by in-depth analytical studies. We do not understand the viruses themselves as yet to a sufficient extent for various biological and molecular biological points. In contrast, unfortunately, we certainly know that there are highly invasive and pathogenic viruses in our environments. It is becoming more likely that humans may experience new viral diseases caused by some emerging zoonotic viruses (an awful pandemic in the worst case), if we think about the world's environmental situation today. Also in this regards, it is pivotal for us virologists to stare at these mysterious viruses from diversified standpoints to really grasp their essentials. Toward this end, we have constructed a united cooperative platform for scientific information and interaction in the rapidly growing virology field. In order to effectively share, explore, and discuss about new findings, Frontiers in Virology consists of 10 distinct unique specialties, that is, "Fundamental virology," "Bioinformatic and predictive virology," "Virus and host immunity," "Antivirals and vaccines," "Emerging and reemerging viruses," "Viral diversification and evolution," "Translational virology," "Viral disease investigation," "Systems virology," and "Modeling of viral replication and pathogenesis." Summaries, keywords, and focuses for each section are as follows.

FUNDAMENTAL VIROLOGY

All basic issues related to various viruses, biology and molecular biology, virus replication, structural and functional virology, viral genetics, chemistry, classification.

BIOINFORMATIC AND PREDICTIVE VIROLOGY

Experimental and computational sciences, informatics, *in silico* studies, homology modeling, MD simulation, theoretical virology, predictive virology.

VIRUS AND HOST IMMUNITY

Host immunity against viruses, recognition of viruses by hosts, interaction between host and virus, viral escape mutations from host immunity, viral pathogenesis.

ANTIVIRALS AND VACCINES

Study and development of effective anti-viral drugs and vaccines, effects of the drugs and vaccines on individuals and the populations, anti-viral treatments.

EMERGING AND REEMERGING VIRUSES

All issues (basic and applied) related to emerging and reemerging viruses, adaptation and evolution of the viruses, description of new viruses in detail.

VIRAL DIVERSIFICATION AND EVOLUTION

Viral mutation, viral adaptation, viral evolution, viral diversity with respect to replication property and biological properties such as pathogenicity, population genetics.

TRANSLATIONAL VIROLOGY

Applied medical sciences based on significant results/findings of basic virology, future perspective, description of (new) systems for the translational research in detail.

VIRAL DISEASE INVESTIGATION

All issues related to various viral diseases, pandemics, medical studies, pre-clinical and clinical studies, pathophysiology of viral diseases, persistent infection.

SYSTEMS VIROLOGY

Innovative research systems, novel technologies/methodologies, experimental/demonstrative studies based on innovative research systems/technologies/methodologies.

MODELING OF VIRAL REPLICATION AND PATHOGENESIS

In vitro model studies, animal models for viral replication and pathogenicity, organoid models for virus infections, any model systems for viral replication and viral diseases.

Finally, we encourage all active researchers, who are experts in virology and also who are not experts in virology but are specially or specifically interested in the research area or the particular theme, to submit the best works to the most fitted section in Frontiers in Virology. We are sincerely looking forward to receiving field-moving scientific articles. Please join us contributing to the promotion and development of scientific interplay in virology and to the successful fight against viral diseases.

AUTHOR CONTRIBUTIONS

AA is a sole contributor to this manuscript and approved its submission.

ACKNOWLEDGMENTS

I would like to thank Masako Nomaguchi and Takaaki Koma (Tokushima University, Tokushima, Japan) for

continuous hot and extensive discussions about Virology. I am awfully indebted to them for conceptualizing and drawing of **Figure 1**. I also appreciate Fumie

Nishina (Kansai Medical University, Osaka, Japan) and Kazuko Yoshida (Tokushima University) for excellent editorial assistance.

REFERENCES

- Nomaguchi M, Adachi A. Virology as biosystematics: towards understanding the viral infection biology. *Front Microbiol.* (2010) 1:2. doi: 10.3389/fmicb.2010.00002
- Sanfaçon H. Grand challenge in plant virology: understanding the impact of plant viruses in model plants, in agricultural crops, and in complex ecosystems. *Front Microbiol.* (2017) 8:860. doi: 10.3389/fmicb.2017.00860
- Adachi A. Grand challenge in human/animal virology: unseen, smallest replicative entities shape the whole globe. *Front Microbiol.* (2020) 11:431. doi: 10.3389/fmicb.2020.00431
- Nomaguchi M, Adachi A. Editorial: highly mutable animal RNA viruses: adaptation and evolution. *Front Microbiol.* (2017) 8:1785. doi: 10.3389/fmicb.2017.01785
- Nomaguchi M, Fujita M, Miyazaki Y, Adachi A. Viral tropism. Front Microbiol. (2012) 3:281. doi: 10.3389/fmicb.2012.00281
- Tsunetsugu-Yokota Y, Terahara K. Receptor usage and the pathogenesis in acute and chronic virus infections. *Front Microbiol.* (2012) 3:289. doi: 10.3389/fmicb.2012.00289
- Ueda K. For the future studies of Kaposi's sarcoma-associated herpesvirus. *Front Microbiol.* (2012) 3:237. doi: 10.3389/fmicb.2012.00237
- Flores R, Moreno P, Falk B, Martelli GP, Dawson WO. e-Book on Closteroviridae. Front Microbiol. (2013) 4:411. doi: 10.3389/fmicb.2013.00411
- Koito A, Ishizaka Y. Retroviruses, retroelements and their restrictions. Front Microbiol. (2013) 4:197. doi: 10.3389/fmicb.2013.00197
- Adachi A, Miura T. Animal model studies on viral infections. *Front Microbiol.* (2014) 5:672. doi: 10.3389/fmicb.2014.00672
- Chan S-W. The unfolded protein response in virus infections. *Front Microbiol.* (2014) 5:518. doi: 10.3389/fmicb.2014.00518
- Kimura H, Ryo A. Pathophysiology and epidemiology of virus-induced asthma. Front Microbiol. (2014) 5:562. doi: 10.3389/fmicb.2014.00562
- Laguette N, Benkirane M. Shaping of the host cell by viral accessory proteins. Front Microbiol. (2015) 6:142. doi: 10.3389/fmicb.2015.00142
- Mahony J, van Sinderen D. Gram-positive phage-host interactions. Front Microbiol. (2015) 6:61. doi: 10.3389/fmicb.2015.00061
- Suzuki N, Sasaya T, Choi IR. Editorial: viruses threatening stable production of cereal crops. Front Microbiol. (2015) 6:470. doi: 10.3389/fmicb.2015.00470
- Aherfi S, Colson P, La Scola B, Raoult D. Giant viruses of amoebas: an update. *Front Microbiol.* (2016) 7:349. doi: 10.3389/fmicb.2016.00349
- Atluri VSR. Editorial: HIV and illicit drugs of abuse. Front Microbiol. (2016) 7:221. doi: 10.3389/fmicb.2016.00221
- Rakonjac J, Das B, Derda R. Editorial: filamentous bacteriophage in bio/nano/technology, bacterial pathogenesis and ecology. *Front Microbiol.* (2016) 7:2109. doi: 10.3389/fmicb.2016.02109
- Liu J-Z, Li F, Liu Y. Editorial: plant immunity against viruses. Front Microbiol. (2017) 8:520. doi: 10.3389/fmicb.2017.00520
- Bertazzoni U, Ciminale V, Romanelli MG. Editorial: molecular pathology of HTLV-1. Front Microbiol. (2018) 9:3069. doi: 10.3389/fmicb.2018.03069
- Martinez-Salas E, Francisco-Velilla R, Fernandez-Chamorro J, Embarek AM. Insights into structural and mechanistic features of viral IRES elements. *Front Microbiol.* (2018) 8:2629. doi: 10.3389/fmicb.2017.02629
- Ziv C, Zhao Z, Gao YG, Xia Y. Multifunctional roles of plant cuticle during plant-pathogen interactions. *Front Plant Sci.* (2018) 9:1088. doi: 10.3389/fpls.2018.01088
- Dobnik D, Kogovšek P, Jakomin T, Košir N, Tušek Žnidarič M, Leskovec M, et al. Accurate quantification and characterization of adeno-associated viral vectors. *Front Microbiol.* (2019) 10:1570. doi: 10.3389/fmicb.2019.01570
- Nasir A, Caetano-Anollés G, Claverie JM. Editorial: viruses, genetic exchange, and the tree of life. *Front Microbiol.* (2019) 10:2782. doi: 10.3389/fmicb.2019.02782
- Rooney BV, Crucian BE, Pierson DL, Laudenslager ML, Mehta SK. Herpes virus reactivation in astronauts during spaceflight and its application on earth. *Front Microbiol.* (2019) 10:16. doi: 10.3389/fmicb.2019.00016

- Sumbria D, Berber E, Rouse BT. Factors affecting the tissue damaging consequences of viral infections. *Front Microbiol.* (2019) 10:2314. doi: 10.3389/fmicb.2019.02314
- Nakayama E, Saijo M. Animal models for Ebola and Marburg virus infections. Front Microbiol. (2013) 4:267. doi: 10.3389/fmicb.2013.00267
- Saiz JC, Vázquez-Calvo Á, Blázquez AB, Merino-Ramos T, Escribano-Romero E, Martín-Acebes MA. Zika virus: the latest newcomer. *Front Microbiol.* (2016) 7:496. doi: 10.3389/fmicb.2016.00496
- Bueno-Marí R, Saiz J-C, Salomón OD, Villamil-Jiménez LC, Heukelbach J, Alencar C H, et al. Editorial: Zika virus research. *Front Neurol.* (2018) 9:168. doi: 10.3389/fneur.2018.00168
- 30. Xu S, Li X, Yang J, Wang Z, Jia Y, Han L, et al. Comparative pathogenicity and transmissibility of pandemic H1N1, avian H5N1, and human H7N9 influenza viruses in tree shrews. *Front Microbiol.* (2019) 10:2955. doi: 10.3389/fmicb.2019.02955
- Yong CY, Ong HK, Yeap SK, Ho KL, Tan WS. Recent advances in the vaccine development against Middle East respiratory syndrome-coronavirus. *Front Microbiol.* (2019) 10:1781. doi: 10.3389/fmicb.2019.01781
- Koma T, Adachi S, Doi N, Adachi A, Nomaguchi M. Toward understanding molecular bases for biological diversification of human coronaviruses: present status and future perspectives. *Front Microbiol.* (2020) 11:2016. doi: 10.3389/fmicb.2020.02016
- Larsen JR, Martin MR, Martin JD, Kuhn P, Hicks JB. Modeling the onset of symptoms of COVID-19. *Front Public Health*. (2020) 8:473. doi: 10.3389/fpubh.2020.00473
- Mercatelli D, Giorgi FM. Geographic and genomic distribution of SARS-CoV-2 mutations. Front Microbiol. (2020) 11:1800. doi: 10.3389/fmicb.2020.01800
- Tse LV, Meganck RM, Graham RL, Baric RS. The current and future state of vaccines, antivirals and gene therapies against emerging coronaviruses. *Front Microbiol.* (2020) 11:658. doi: 10.3389/fmicb.2020.00658
- Wang N, Shang J, Jiang S, Du L. Subunit vaccines against emerging pathogenic human coronaviruses. *Front Microbiol.* (2020) 11:298. doi: 10.3389/fmicb.2020.00298
- Szunerits S, Barras A, Khanal M, Pagneux Q, Boukherroub R. Nanostructures for the inhibition of viral infections. *Molecules*. (2015) 20:14051–81. doi: 10.3390/molecules200814051
- Athanasopoulos T, Munye MM, Yáñez-Muñoz RJ. Nonintegrating gene therapy vectors. *Hematol Oncol Clin North Am.* (2017) 31:753–70. doi: 10.1016/j.hoc.2017.06.007
- Graham BS, Gilman MSA, McLellan JS. Structure based vaccine antigen design. Annu Rev Med. (2019) 70:91–104. doi: 10.1146/annurev-med-121217-094234
- Miyazaki Y, Fujita M, Nomaguchi M, Adachi A. Structural biology for virus research. Front Microbiol. (2012) 3:91. doi: 10.3389/fmicb.2012.00091
- Sato H, Yokoyama M, Toh H. Genomics and computational science for virus research. Front Microbiol. (2013) 4:42. doi: 10.3389/fmicb.2013.00042
- Bai XC, McMullan G, Scheres SH. How cryo-EM is revolutionizing structural biology. *Trends Biochem Sci.* (2015) 40:49–57. doi: 10.1016/j.tibs.2014.10.005
- Angermueller C, Pärnamaa T, Parts L, Stegle O. Deep learning for computational biology. *Mol Syst Biol.* (2016) 12:878. doi: 10.15252/msb.20156651
- Chen C, Huang H, Wu C. Protein bioinformatics databases and resources. Methods Mol Biol. (2017) 1558:3–39. doi: 10.1007/978-1-4939-6783-4_1
- Dutilh BE, Reyes A, Hall RJ, Whiteson KL. Editorial: virus discovery by metagenomics: the (im)possibilities. *Front Microbiol.* (2017) 8:1710. doi: 10.3389/fmicb.2017.01710
- Dutta D, Heo I, Clevers H. Disease modeling in stem cellderived 3D organoid systems. *Trends Mol Med.* (2017) 23:393–410. doi: 10.1016/j.molmed.2017.02.007
- Hasin Y, Seldin M, Lusis A. Multi-omics approaches to disease. *Genome Biol.* (2017) 18:83. doi: 10.1186/s13059-017-1215-1
- 48. Yamamoto T, Honda A, Sato T, Ryo A. Editorial: perspectives for the next generation of virus research: spearheading the use of innovative

technologies and methodologies. Front Microbiol. (2017) 8:758. doi: 10.3389/fmicb.2017.00758

- Adli M. The CRISPR tool kit for genome editing and beyond. *Nat Commun.* (2018) 9:1911. doi: 10.1038/s41467-018-04252-2
- Artegiani B, Clevers H. Use and application of 3D-organoid technology. Hum Mol Genet. (2018) 27:R99–R107. doi: 10.1093/hmg/ddy187
- Cheng Y. Membrane protein structural biology in the era of single particle cryo-EM. *Curr Opin Struct Biol.* (2018) 52:58–63. doi: 10.1016/j.sbi.2018.08.008
- Ibrahim B, McMahon DP, Hufsky F, Beer M, Deng L, Mercier PL, et al. A new era of virus bioinformatics. *Virus Res.* (2018) 251:86–90. doi: 10.1016/j.virusres.2018.05.009
- Rossi G, Manfrin A, Lutolf MP. Progress and potential in organoid research. Nat Rev Genet. (2018) 19:671–87. doi: 10.1038/s41576-018-0051-9
- Sbalzarini IF, Greber UF. How computational models enable mechanistic insights into virus infection. *Methods Mol Biol.* (2018) 1836:609–31. doi: 10.1007/978-1-4939-8678-1_30

- van Dijk EL, Jaszczyszyn Y, Naquin D, Thermes C. The third revolution in sequencing technology. *Trends Genet.* (2018) 34:666–81. doi: 10.1016/j.tig.2018.05.008
- Pickar-Oliver A, Gersbach CA. The next generation of CRISPRCas technologies and applications. *Nat Rev Mol Cell Biol.* (2019) 20:490–507. doi: 10.1038/s41580-019-0131-5

Conflict of Interest: The author declares that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Copyright © 2021 Adachi. This is an open-access article distributed under the terms of the Creative Commons Attribution License (CC BY). The use, distribution or reproduction in other forums is permitted, provided the original author(s) and the copyright owner(s) are credited and that the original publication in this journal is cited, in accordance with accepted academic practice. No use, distribution or reproduction is permitted which does not comply with these terms.