Editorial—Special Issue on Molecular Communications for Diagnostics and Therapeutic Development of Infectious Diseases

THE COVID-19 pandemic in the last year has brought along numerous challenges on all fronts for humanity. Besides the high number of fatalities, economic and societal impact, the virus has also raised a question on the current approaches and methodologies for combatting pandemics. An important factor in this global fight against the pandemic, is the fact that key experts within the fields of biotechnology, virology and immunology have to return to the drawing board to develop new approaches to treat the virus. This has triggered a multi and inter-disciplinary research drive to bring in new concepts, techniques and methodologies that can be used to understand the virus in order to develop novel treatment techniques. The field of molecular communications, which is just over a decade old, is one area of research in the field of communications and networking that can contribute towards understanding of virus and their infection process. This is the aim of this special issue, where we have collected a number of publications from key researchers in the field of molecular communications that can contribute towards understanding the viral properties and behavior, which can be used to support the current as well as future pandemics.

The first publication selected by the guest editors -"Molecular Communications in Viral Infections Research: Modeling, Experimental Data and Future Directions," aims to collect vital data for a number of well known viruses that can be used by molecular communications researchers to model and simulate their propagation behavior. Examples of these viruses that have been analyzed in this paper include SARS-CoV-(1, 2), MERS-CoV, Ebola, Dengue, Zika, and Hepatitis C. The paper discusses a number of different propagation methods within the body, such as through the lungs, circulatory systems, between cells, as well as propagation behavior when respiratory clouds are secreted from people. The objective is to entice a new direction in molecular communications, where the community can use the data set to simulate accurate propagation scenarios in order to provide support for virologists and immunologists in better understandings of the virus.

In the second publication "VIVID: *In Vivo* End-to-End Molecular Communication Model for COVID-19," the authors aim to understand the transmission and propagation dynamics of coronavirus inside the respiratory system. The paper models the transmission and propagation of coronavirus inside the respiratory tract, starting from the nasal area to alveoli

using concepts from molecular communications, and simulated using COMSOL multi-physics modeling tool. The analysis conducted includes the end-to-end communication model of path loss, delay, and gain.

The publication "Molecular Communication Theoretical Modeling and Analysis of SARS-CoV-2 Transmission in Human Respiratory System" models and analyzes the SARS-CoV-2 transmission through the human respiratory tract using concepts from molecular communication. The analysis considers the virus diffusion within the mucus layer, and found that the shape of the respiratory tract does not play a role in the transmission process. The publication also presents a unique impulse response of SARS-CoV-2 and its binding process to the angiotensin-converting enzyme 2 (ACE2) expressing cells to determine the proportion of the virus in different parts of the lungs. The significance of this study is the relationship that has been established between high mucus flow rate that impacts on the virus migration to the lower part of the respiratory tract.

The publication "A Molecular Communications System for the Detection of Inflammatory Levels Related to COVID-19 Disease" analyzes the excessive interleukin (IL)-6 level and its evolution in the blood vessels using molecular communications theory. The publication considers the mechanism for the SARS-CoV-2 to bind to ACE2 expressing cells, which are found in the endothelium lining of the circulatory system, to diffuse through and lead to endothelial inflammation. This then leads to the stimulation of cytokine storms as part of the immune response. The work developed in this publication was analyzed using the BiNS2 simulator, to characterize the flow-based molecular communications in blood vessels, as well as Markov models of the endothelium.

In "A Molecular Communication Perspective on Airborne Pathogen Transmission and Reception via Droplets Generated by Coughing and Sneezing," the spreading mechanism of infectious disease by airborne pathogen transmission is modeled using concepts from molecular communications. The airborne propagation model considers the impact of the gravity, initial velocity and buoyancy for the droplets as it propagates through the air. The results from the study found that a number of factors that impact the infection process, such as the exposure time, and the angle of sneezing, can have an impact on the propagation behavior. The results of this study can contribute to designs of novel mask structures that prevent the maximum quantity of droplets from diffusion into the environment, as well as appropriate social-distance.

The publication "Viral Aerosol Concentration Characterization and Detection in Bounded Environments" proposes a respiratory model for breathing, coughing, and sneezing of aerosol droplets propagation as a molecular communication system that is detected through a biosensor within an environment. The propagation modeling considers the secreted molecules as a cloud and analyzes the spatialtemporal virus concentration propagated within a bounded environment. A contribution from this publication is a closedform expression for the virus concentration. The analysis is based on simulations that analyze the detection efficiency of the biosensor. The contribution of this publication can lead to optimal placements of the biosensors within an environment to maximize the detection of secreted droplets from people.

publication "Duality Between Coronavirus Transmission and Air-Based Macroscopic Molecular Communication," analyzes the airborne aerosol and droplet transmission viral infection process as an air-based molecular communication system. The aerosol and droplet transmission is modeled as a multiuser molecular communication scenario employing respiratory-event-driven molecular variableconcentration shift keying modulation process. Based on molecular communications testbeds that have been previously proposed for characterizing the air-based molecular communication testbed, the study proposes a simulation tool for estimating the transmission of the aerosols within an environment. This publication can lead to accurate techniques of understanding the infection processes as the virus is secreted from the respiratory system, which can lead to more accurate proposals for social distancing as well as the development of masks for people during a pandemic.

We would like to thank Chan-Byoung Chae, the Editor-in-Chief of IEEE TRANSACTIONS ON MOLECULAR, BIOLOGICAL, AND MULTI-SCALE COMMUNICATIONS and Christina Tang-Bernas, the journal manager, for their endless support and guidance to make this special issue a success. The collection of publications from this special issue will hopefully attract a new research direction from the molecular communications community that can contribute towards new understanding in viral pandemics.

SASITHARAN BALASUBRAMANIAM, Guest Editor Department of Computer Science and Engineering University of Nebraska-Lincoln Lincoln, NE 68588 USA

e-mail: sasitharanb@gmail.com

MICHAEL TAYNNAN BARROS, Guest Editor School of Computer Science and Electronic Engineering University of Essex Colchester CO4 3SQ, U.K. e-mail: m.barros@essex.ac.uk

MLADEN VELETIĆ, Guest Editor Intervention Center Oslo University Hospital 0372 Oslo, Norway e-mail: mlavel@rr-research.no

MASAMITSU KANADA, *Guest Editor*Department of Pharmacology and Toxicology
Michigan State University
East Lansing, MI 48824 USA
e-mail: kanadama@msu.edu

MASSIMILIANO PIEROBON, *Guest Editor*Department of Computer Science and Engineering
University of Nebraska–Lincoln
Lincoln, NE 68588 USA
e-mail: maxp@unl.edu

SEPPO VAINIO, Guest Editor Biochemistry and Molecular Medicine University of Oulu 90570 Oulu, Finland e-mail: seppo.vainio@oulu.fi

ILANGKO BALASINGHAM, Guest Editor
Department of Electronic Systems
Norwegian University of Science and Technology
7491 Trondheim, Norway
e-mail: ilangko.balasingham@ous-research.no



Sasitharan Balasubramaniam (Senior Member, IEEE) received the B.E. degree in electrical and electronic engineering from The University of Queensland, Brisbane, QLD, Australia, in 1998, the M.E. degree in computer and communication engineering from the Queensland University of Technology, Brisbane, in 1999, and the Ph.D. degree from The University of Queensland in 2005. He was the Director of the Research with the Telecommunication Software and Systems Group, Waterford Institute of Technology, Waterford, Ireland. He is currently an Associate Professor with the Department of Computer Science and Engineering, University of Nebraska-Lincoln (UNL), Lincoln, NE, USA. He was a Principal Investigator with the Science Foundation Ireland VistaMilk (SFI) Research Centre, and was a Funded Investigator with the SFI CONNECT and the FutureNeuro Research Centres. He has authored or coauthored more than 100 articles in various journals and conferences and actively participates in a number of technical program committees for various conferences. His current research interests include molecular communications. Internet

of (Bio) NanoThings, and terahertz wireless communications. He is on the Steering Board Committee of ACM NANOCOM, which he co-founded. In 2018, he was the IEEE Nanotechnology Council Distinguished Lecturer. He has served as an Associate Editor for the IEEE Internet of Things Journal. He serves as an Associate Editor for the IEEE Transactions on Molecular, Biological and Multi-Scale Communications, the IEEE Transactions on Mobile Computing, Nano Communication Networks (Elsevier), and Digital Communication and Networks (Elsevier).



Michael Taynnan Barros received the Ph.D. degree in telecommunication software from the Waterford Institute of Technology in 2016. He is a Marie Skłodowska Curie Individual Fellow (MSCA-IF) with the BioMediTech Institute, Tampere University, Finland, and an Assistant Professor (Lecturer) with the School of Computer Science and Electronic Engineering, University of Essex, U.K. He has authored or coauthored over 70 research papers in various international flagship journals and conferences in the areas of wireless communications, molecular and nanoscale communications, as well as bionanoscience. His research interests include Internet of BioNanoThings, molecular communications, bionanoscience, and 6G communications. He received the CONNECT Prof. Tom Brazil Excellence in Research Award in 2020.



Mladen Veletić received the B.Sc. and M.Sc. degrees in electronics and telecommunications from the Faculty of Electrical Engineering, University of Banja Luka (UNIBL), Bosnia and Herzegovina, in 2010 and 2012, respectively, and the Ph.D. degree in telecommunications from the Department of Electronic Systems, Norwegian University of Science and Technology, Norway, and the Faculty of Electrical Engineering, UNIBL. From 2011 to 2017, he was a Senior Teaching and a Research Assistant with UNIBL. He is currently a Postdoctoral Research Scientist with the Intervention Center, Oslo University Hospital. He was awarded a Gold Plaque from the UNIBL for his achievements throughout his undergraduate education. His research interests include molecular and nano-neural communications, wireless communications, and positioning in cellular networks.



Masamitsu Kanada studied cell biology and fluorescence live cell imaging at Tsukuba University, Japan. He worked for a pharmaceutical company developing cancer therapies. During his postdoctoral training, he learned intravital microscopy and whole-body preclinical imaging in the laboratories of Susumu Terakawa with Hamamatsu University School of Medicine and Christopher Contag with Stanford University. He is an Assistant Professor with the Department of Pharmacology and Toxicology. His research focuses on addressing questions in cancer biology using multidisciplinary approaches.



Massimiliano Pierobon received the Ph.D. degree in electrical and computer engineering from the Georgia Institute of Technology, Atlanta, GA, USA, in 2013. Since August 2013, he has been an Assistant Professor with the Department of Computer Science and Engineering, University of Nebraska–Lincoln (UNL), Lincoln, NE, USA, where he also holds a courtesy appointment with the Department of Biochemistry. His research interests are in molecular communication theory, nanonetworks, intrabody networks, communication engineering applied to synthetic biology, and the Internet of Bio-Nano Things. His selected honors include the 2011 Georgia Tech BWN Lab Researcher of the Year Award, the 2013 IEEE COMMUNICATIONS LETTERS Exemplary Reviewer Award, the UNL CSE Upper and Graduate Level Teaching Award in 2016 and 2017, the 2017 IEEE INFOCOM Best Paper Runner-Up Award, and the Best In-Session Presentation Award. He is currently the PI of the NSF Project WetComm: Foundations of Wet Communication Theory and a Co-PI of the NSF Project Redox-Enabled Bio-Electronics for Molecular Communication and

Memory (RE-BIONICS), and has been the lead PI of the NSF Project TelePathy: Telecommunication Systems Modeling and Engineering of Cell Communication Pathways. He has been the Co-Editor-in-Chief of *Nano Communication Networks* (Elsevier) since July 2017, and was an Associate Editor of IEEE TRANSACTIONS ON COMMUNICATIONS from 2013 to 2020.



Seppo Vainio is a PI with Biocenter Oulu and Infotech Oulu research organizations. He has published over 130 original research articles and reviews. His Hirsch index is 29 and his work has been cited more than 10 000 times. He has delivered invited presentations to peer-reviewed, internationally established conferences. He also serves as a referee for many international journals, such as *Nature*, *Cell*, *Development*, *Proceedings of the National Academy of Sciences of the United States of America*, *Journal of the American Society of Nephrology*, and Nal. He has also organized international research congresses, such as a symposia on Development, Regeneration & Aging in Oulu 7–9, 2012 and served as the Session Chair as in the interdisciplinary Argumenta Finnish Cultural Foundation funded symposia in Oulu 7–9, 2012; The Gordon Research Conference on Mammary Gland Biology, June 1–6, 2008; Il Ciocco Hotel and Resort Lucca (Barga) and International Symposium for Gonad and Brain Sex Differentiation, Fukuoka, Japan, in 2008; The 1st Kumamoto University Global COE International Symposium on Cell Fate Regulation,

January 16, 2008; Fourth Swedish–Finnish Connective Tissue Meeting, April 13 to April 14, 2007; Lund–Sweden World Congress on Nephrology, April 21–25, 2007, Rio de Janeiro, Brazil; ELSO—the European Life Scientist Organization Meeting in 2007, Dresden, 1–4 September; and the Fourth Course on Genetics From Developmental Genes to Dysmorphology. He had substantial research funding from the Academy of Finland, European Union. He is the PI of the Academy of Finland Centers of Excellence and a Coordinator of the EU Project "Wnt and Stem Cells" (QLK3-CT-2001-01275)(2001–2004). He is a member of the Centre of Excellence of Academy of Finland. He has hosted FiDIpr Susan Quaggin in his lab and has also been a Visiting Distinguished Professor with Ulm University since 2013, Prof. Pool Grant 2012–2013, Biocenter Oulu. His awards include discovery of the year 2011, Physiologist of the Year in Finland (1999), and discovery of the year in Biocenter Oulu (1999, 2004, 2009, and 2010 rank position). His granted patents include "Induction of kidney tubule formation, U.S. patents No 09/937,735, Wnt signalling in reproductive organs, U.S. patent No 00246/232001 and Process for producing active Wnt proteins in prokaryotes. Finnish Patent application FI 20040842."



Hangko Balasingham (Senior Member, IEEE) received the M.Sc. and Ph.D. degrees in signal processing from the Department of Electronic Systems, Norwegian University of Science and Technology (NTNU), Trondheim, Norway, in 1993 and 1998, respectively. He performed his Master's degree thesis with the Department of Electrical and Computer Engineering, University of California at Santa Barbara, USA. From 1998 to 2002, he worked as a Research Engineer developing image and video streaming solutions for mobile handheld devices with Fast Search & Transfer ASA, Oslo, Norway, which is currently part of Microsoft Inc. Since 2002, he has been with the Intervention Center, Oslo University Hospital, Oslo, where he heads the Wireless Sensor Network Research Group. He was appointed a Professor of Medical Signal Processing and Communications with NTNU in 2006. From 2016 to 2017, he was a Professor by courtesy with the Frontier Institute, Nagoya Institute of Technology, Japan. His research interests include super robust short-range communications for both in-body and on-body sensors, body area sensor

network, microwave short-range sensing of vital signs, short-range localization and tracking mobile sensors, and nanoscale communication networks. He has authored or coauthored over 280 journal and conference papers, seven book chapters, 42 abstracts, six patents, and 20 articles in popular press. He has given 21 invited/keynotes at international conferences. In addition, he is active in organizing conferences (the Steering Committee Member of ACM NANOCOM since 2018; the General Chair of the 2019 IEEE International Symposium of Medical ICT and the 2012 Body Area Networks (BODYNETS) conference; and the TPC Chair of the 2015 ACM NANOCOM) and editorial board (an Area Editor of *Nano Communication Networks* (Elsevier) since 2013 and the Specialty Chief Editor of *Frontiers in Communications and Networks* since 2020).