







Institute for Neurological Research and Diagnostic







FUNDATIA JURNALULUI Journal of Medicine

and Life



Seminars

Department of Neurosciences University of Medicine and Pharmacy "Iuliu Hatieganu" Cluj-Napoca | Romania



VIRTUAL MEETING

Welcome Address

It is a pleasure to welcome you to the 88th edition Seminars 13 April, 2022. The seminar is hosted by the Department of Neurosciences, Faculty of Medicine, "Iuliu Hatieganu" University of Medicine and Pharmacy, Cluj-Napoca. This seminar aims to establish itself as a highly useful framework that will enable local specialists to benefit from the expertise of our invited speakers who are part of associated international faculty of our Department of Neurosciences Cluj-Napoca, Romania and RoNeuro Science network. Our scope is to flourish over years and set up an educational vector aiming to meet our junior and senior specialists' needs.

In contrast to large international conferences, the intention behind these seminars is to create an informal and intimate setting, which hopefully will stimulate open discussions.

Due to the uncertainties about the continuing impact of the COVID-19 pandemic, our events will be held in the virtual space, for the time being. As organizers, we would therefore be deeply grateful if you participate and share your time with us.

We are looking forward to your active participation in this educational event!

With consideration,

Prof. Dr. Dafin F. Muresanu,

Chairman Department of Neurosciences, Faculty of Medicine, "Iuliu Hatieganu" University of Medicine and Pharmacy, Cluj-Napoca, Romania

Defiction hureman

Program Coordinator



Dafin F. Mureşanu

President of the European Federation of NeuroRehabilitation Societies (EFNR)

Chairman of EAN Communication and Liaison Committee

Co-Chair EAN Scientific Panel Neurotraumatology

Past President of the Romanian Society of Neurology

Professor of Neurology, Chairman Department of Neurosciences "Iuliu Hatieganu" University of Medicine and Pharmacy, Cluj-Napoca, Romania

International Guest Lecturer



Claudio Bassetti

Dean, Medical Faculty, University Bern

President European Academy of Neurology

Full Professor of Neurology and Chair, Neurology Department, University Hospital, Bern, Switzerland











Institute for Neurological Research and Diagnostic





Academia de Științe Medicale din România







Scientific program

13 **APRIL**, 2022 VIRTUAL MEETING

- 13:00 13:45 Sleep and Neurology Claudio Bassetti /Switzerland
- 13:45 14:20 Sleep and Dementia Claudio Bassetti /Switzerland
- 14:20 15:10 Sleep and Movement Disorders (SRMD) Claudio Bassetti /Switzerland





SPEAKER

Professor of Neurology, Senior Neurologist, Chairman of the Neurosciences Department, Faculty of Medicine, "Iuliu Hatieganu" University of Medicine and Pharmacy Cluj-Napoca, President of the European Federation of Neurorehabilitation Societies (EFNR), Chairman Communication Committee of the European Academy of Neurology (EAN), Past President of the Romanian Society of Neurology, President of the Society for the Study of Neuroprotection and Neuroplasticity (SSNN), Chairman "RoNeuro" Institute for Neurological Research and Diagnostic, Corresponding Member of the Romanian Academy, Member of the Academy of Medical Sciences, Romania and secretary of its Cluj Branch. He is member of 17 scientific international societies (being Member of the American Neurological Association (ANA) - Fellow of ANA (FANA) since 2012) and 10 national ones, being part of the executive board of most of these societies. Professor Dafin F. Muresanu is also a specialist in Leadership and Management of Research and Health Care Systems (specialization in "Management and Leadership, Arthur Anderson Institute, Illinois, USA, 1998"; "MBA - Master of Business Administration - Health Care Systems Management, The Danube University - Krems, Austria, 2003"). He has performed valuable scientific research in high interest fields such as: neurobiology of central nervous system (CNS) lesion mechanisms; neurobiology of neuroprotection and neuroregeneration of CNS; the role of the Blood-brain barrier (BBB) in CNS diseases; developing comorbidities in animal models to be used in testing therapeutic paradigms; nanoparticles neurotoxicity upon CNS; the role of nanoparticles in enhancing the transportation of pharmacological therapeutic agents through the BBB; cerebral vascular diseases; neurodegenerative pathology; traumatic brain injury; neurorehabilitation of the central and peripheral nervous system; clarifying and thoroughgoing study on the classic concepts of Neurotrophicity, Neuroprotection, Neuroplasticity and Neurogenesis by bringing up the Endogenous Defense Activity (EDA) concept, as a continuous nonlinear process, that integrates the four aforementioned concepts, in a biological inseparable manner.

Professor Dafin F. Muresanu is coordinator in international educational programs of European Master (i.e. European Master in Stroke Medicine, University of Krems), organizer and co-organizer of many educational projects: European and international schools and courses (International School of Neurology, European Stroke Organisation Summer School, Danubian Neurological Society Teaching Courses, Seminars - Department of Neurosciences, European Teaching Courses on Neurorehabilitation) and scientific events: congresses, conferences, symposia (International Congresses of the Society for the Study of Neuroprotection and Neuroplasticity (SSNN), International Association of Neurorestoratology (IANR) & Global College for Neuroprotection and Neuroregeneration (GCNN) Conferences, Vascular Dementia Congresses (VaD), World Congresses on Controversies in Neurology (CONy), Danube Society Neurology Congresses, World Academy for Multidisciplinary Neurotraumatology (AMN) Congresses, Congresses of European Society for Clinical Neuropharmacology, European Congresses of Neurorehabilitation). His activity includes involvement in many national and international clinical studies and research projects, over 500 scientific participations as "invited speaker" in national and international scientific events, a significant portfolio of scientific articles (260 papers indexed on Web of Science-ISI, H-index: 25) as well as contributions in monographs and books published by prestigious international publishing houses. Prof. Dr. Dafin F. Muresanu has been honoured with: "Dimitrie Cantemir" Medal of the Academy of The Republic of



Dafin F. Muresanu /Romania



Moldova in 2018, Ana Aslan Award 2018 - "Performance in the study of active aging and neuroscience", for the contribution to the development of Romanian medicine, National Order "Faithful Service" awarded by the President of Romania in 2017; "Iuliu Hatieganu" University of Medicine and Pharmacy Cluj-Napoca, Faculty of Medicine, the "Iuliu Hatieganu Great Award 2016" for the best educational project in the last five years; the Academy of Romanian Scientists, "Carol Davila Award for Medical Sciences / 2011", for the contribution to the Neurosurgery book "Tratat de Neurochirurgie" (vol.2), Editura Medicala, Bucuresti, 2011; the Faculty of Medicine, "Iuliu Hatieganu" University of Medicine and Pharmacy Cluj-Napoca "Octavian Fodor Award" for the best scientific activity of the year 2010 and the 2009 Romanian Academy "Gheorghe Marinescu Award" for advanced contributions in Neuroprotection and Neuroplasticity.



SPEAKER

Claudio Bassetti was born and raised in Ticino, is married and father of three boys. He received his MD degree from the University of Basel in 1984. He trained in neurology in Bern and Lausanne and performed research fellowships in basic neurophysiology (Basel) and sleep medicine (Ann Arbor, USA). In 2000 he was appointed professor of neurology at the University of Zurich. In 2009 he founded the Neurocenter of Southern Switzerland which he directed for 3 years. Since 2012 he is full professor of neurology at the University of Bern and director of the neurology department at Inselspital. Bassetti authored over 400 scientific publications and eight books. He pioneered the research on the bidirectional relationship between, sleep, sleep disorders and stroke using both a human and animal/ experimental approaches. He made also fundamental contributions to our understanding of narcolepsy, including the recent discovery of specific autoreactive-T lymphocytes supporting the hypothesis of an autoimmune etiology. He served as president of the European Neurological Society, European Sleep Research Society and Swiss Neurological Society and was the founder of the Swiss Federation of Clinical Neurosocieties. In 2018 he became an elected member of the Swiss Academy of Medical Sciences. He currently serves as president of the European Academy of Neurology and as the Dean of the Medical Faculty in Bern.



Claudio Bassetti /Switzerland

Representative publications

1) Bassetti C, Aldrich M, Chervin R, Quint D. Sleep apnea in patients with TIA and Stroke. A prospective study of 59 patients. Neurology 1996

- 2) Bassetti C, Aldrich MS. Idiopathic hypersomnia. A series of 42 patients. Brain 1997
- 3) Bassetti C, Vella S, Donati F. SPECT during Sleepwalking. Lancet 2000

4) Khatami R, Maret S, Werth E, Rétey J, Schmid D, Maly F, Tafti M, Bassetti CL. A monozygotic twin pair concordant for narcolepsy-cataplexy without any detectable abnormality in the hypocretin pathway. Lancet 2004

5) Schwartz S, Ponz A, Poryazova R, Werth E, Boesiger P, Khatami R, Bassetti CL. Abnormal activity in hypothalamus and amygdala during humour processing in human narcolepsy with cataplexy. Brain 2008

6) Pace M, Baracchi F, Gao B, Bassetti C. Identification of sleep-modulated pathways involved in neuroprotection from stroke. Sleep 2015

7) Brill AK, Horvath T, Seiler A, Camilo M, Haynes AG, Ott SR, Egger M, Bassetti CL. CPAP as treatment of sleep apnea after stroke- a meta-analysis of randomized trials. Neurology 2018

8) Leemburg S, Gao B, Cam E, Sarnthein J, Bassetti CL. Power spectrum slope is related to motor function after focal cerebral ischemia in the rat. Sleep 2018

9) Latorre D, Kallweit U,....Bassetti C*, Sallusto F*. T cells in patients with narcolepsy target self-antigens of hypocretin neurons. Nature 2018 (*co-shared last authors) 10) Bassetti C.L.A., A. Adamantidis, D. Burdakov, et al. Narcolepsy. Nature Rev Neurol 2019



SLEEP AND NEUROLOGY

INTRODUCTION: the main theories on the function of sleep (neuronal restoration/ integrity^{1,2}; learning/memory

consolidation³⁻⁵; energy saving/allocation6) and the principles of sleep staging/ scoring and sleep-wake regulation are presented.

EPIDEMIOLOGY AND CLINICAL FEATURES: the frequency of sleep-wake disturbances (SWD) in neurological patients and the overall impact of SWD on the course of neurological disorders is discussed. Important causes of "neurogenic" insomnia (e.g. restless legs syndrome, stroke, Creutzfeldt-Jakob disease, frontal lobe lesions, M. Alzheimer) hypersomnia (e.g. narcolepsy, stroke, Parkinsonism) and parasomnia (e.g. Parkinsonism) are presented⁷⁻¹².

DIAGNOSIS/MANAGEMENT: history taking in patients with SWD, when to refer patients to specialized sleep centers and treatment options for neurogenic SWD are presented.

SLEEP-RELATED MOVEMENT DISORDERS (SRMD)

The vignette of a patient with a SRMD is presented at the beginning, and its solution at the end of the lecture.

INTRODUCTION: the physiology of motor control in sleep and the variety of physiological sleep-associated motor activities are briefly discussed^{13,14}. The concept of "state dissociation" as pathophysiological principle of (most) SMD is presented^{15,16.}

EPIDEMIOLOGY, CLINICAL ASPECTS: sleepwalking and REM sleep behavior disorder are the most important complex SRMD¹⁷⁻²¹. The list of other SRMD is shortly discussed^{13,14,22}.

DIAGNOSIS/MANAGEMENT: the diagnostic work-up and treatment options for patients with SRMD are discussed.

CLAUDIO BASSETTI /Switzerland

CLAUDIO BASSETTI /Switzerland

SLEEP AND DEMENTIA

The vignette of a patient with cognitive decline and a sleep disturbance is presented at the beginning, and its solution at the end of the lecture.

INTRODUCTION: the suggested mechanisms of sleep-related memory consolidation^{3,32,33} and the sleep-wake changes observed during aging are presented34. Experimental studies linking sleep loss with the accumulation of toxic proteins in the brain are shown^{35,36.}

EPIDEMIOLOGY AND CLINICAL ASPECTS: sleep-wake changes in the pre-symptomatic, early and advanced phases of dementia syndrome including M. Alzheimer and M. Parkinson are shown^{37,38.}

DIAGNOSIS/MANAGEMENT: the diagnostic work-up and treatment options for demented patients with sleepwake disturbances are discussed^{39,40}

REFERENCES

1.Xie L, Kang H, Xu Q, et al. Sleep drives metabolite clearance from the adult brain. Science 2013;342:373-377.

2.De Vivo L. Ultrastructural evidence for synaptic scaling across the wake/sleep cycle. Science 2017;355:507.510.

3.Rasch B, Born J. About sleep's role in memory. Physiol Rev 2013;93:688-766.

4.Yang G, Lai CS, Cichon M, et al. Sleep promotes branch-specific formation of dendritic spines after learning. Science 2014;344:1173-1178.

5.Boyce R, Glasgow SD, Williams S, Adamantidis AR. Causal evidence for the role of REM sleep theta rhythm in contextual memory consolidation. Science 2016;352:812-816.

6.Schmidt M. The energy allocation function of sleep: A unifying theory of sleep, torpor, and continuous wakefulness. Neuroscience and Biobehav Rev 2014;47:122-153.

7.Landolt H, Glatzel M, Blatter T, et al. Sleep-wake disturbances in sporadic Creutzfeldt-Jakob disesae. Neurology 2006;66:1418-1424.

8.Hermann DM, Siccoli M, Brugger P, et al. Evolution of neurological, neuropsychological and sleep-wake disturbances after paramedian thalamic stroke. Stroke 2008;39:62-68.

9.Bassetti CL. Primary and Secondary Neurogenic Hypersomnias. Sleep Med Clin 2012;7:249-261.

10.Bassetti CL, Peigneux P, Dogas Z. ESRS European Sleep Medicine Textbook. Regensburg: European Sleep Research Society (ESRS), 2014.

11.Hermann DM, Bassetti CL. Role of sleep-disordered breathing and sleep-wake disturbances for stroke and stroke recovery. Neurology 2016;87:1-10.

12.Bargiotas P, Schuepbach MWM, Bassetti CL. Sleep–wake disturbances in the premotor and early stage of Parkinson's disease. Curr Opin Neurol 2016;29:763-772.

13.Chokroverty S, Allen RP, Walters AS, Montagna P. Sleep and movement disorders: Oxford University Press, 2013.

14.Bassetti CL. Sleep-related Movement Disorders and Disorders of Motor Control. In: Wolters E. BC, ed. Parkinson Disease & Other Movement Disorders: Motor Behavioural Disorders & Behavioural Motor Disorders: University Press, 2014: 661-674.

15.Mahowald MK, Schenck CH. Dissociated states of wakefulness and sleep. Neurology 1992;42 (Suppl 6):44-52.

CLAUDIO BASSETTI /Switzerland

16.Terzaghi M, Sartori I, Tassa L, et al. Dissociated local arousal states underlying essential clinical features of non-rapid eye movement arousal parasomnia: an intracerebral stereo-electroencephalographic study. J Sleep Res 2012;21:502-506.

17.Bassetti C, Vella S, Donati F, Wielepp P, Weder B. SPECT during sleepwalking. Lancet 2000;356:484-485.

18.Bassetti CL. Sleepwalking (Somnambulism). In: Laures S, Tononi G, eds. The Neurology of Consciousness: Cognitive Neuroscience and Neuropatholog: Academic Press, 2009.

19.Zadra A, Desautels A, Petit D, Montplaisir J. Somnambulism: clinical aspects and pathophysiological hypotheses. Lancet Neurol 2013;12:285-294.

20.Iranzo A, Fernandez-Arcos A, Tolosa E, et al. Neurodegenerative Disorder Risk in Idiopathic REM Sleep Behavior Disorder: Study in 174 Patients. Plos One 2014;9:e89741.

21.Fernandez-Arcos A, Iranzo A, Serradell M, et al. The Clinical Phenotype of Idiopathic Rapid Eye Movement Sleep Behavior Disorder at Presentation: A Study in 203 Consecutive Patients. Sleep 2016;39:121-132.

22. Mayer C. Sleep related rhythmic movement disorder revisited. J Sleep Res 2007;16.

23.Halasz P.How Sleep Activates Epileptic Networks? Epilepsy Research and Treatment 2013;2013:dx.doi.org/10.1155/2013/425697. 24.Frauscher B, von Ellenrieder N, Ferrari-Marinho T, et al. Facilitation of epileptic activity during sleep is mediated by high amplitude slow waves. Brain 2015;138:1629-1641.

25.Glbbs SA, Proserpio P, Terzaghi M, et al. Sleep-related epileptic behaviors and non-REM-related parasomnias: Insights from stereo-EEG. Sleep Med Rev 2016;25:4-20.

26.Frauscher B, von Ellenrieder N, Dubeau F, et al. EEG desynchronization during phasic REM sleep suppresses interictal epileptic activit in humans. Epilepsia 2016;57:879-888.

27.Tassinari CA. Neuroethological approach to frontolimbic epileptic seizures and parasomnias: The same central pattern generators for the same behaviours. Rev Neurol 2009;165:762-768.

28.Höllinger P, Khatami R, Gugger M, Hess CW, Bassetti C. Epilepsy and obstructive sleep apnea. Eur Neurol 2006;55:74-79.

29. Tinuper P. Definition and diagnostic criteria of sleep-related hypermotor epilepsy. Neurology 2016;86:1-9.

30.Derry CP, Duncan JS, Berkovic SF. Paroxysmal Motor Disorders of Sleep: The Clinical Spectrum and Differentiation from Epilepsy. Epilepsia 2006;47:1775-1791. 31.Manni R, Terzaghi M, Repetto A. The FLEP scale in diagnosis nocturnal frontal lobe from epilepsy, NREM and REM parasomnias: Data from tertiary sleep and epilepsy unit. Epilepsia 2008;49:1581-1585.

32.Marshall L, Helgadottir H, Mölle M, Born J. Boosting slow oscillations during sleep potentiates memory. Nature 2016;444:610-613.

33.Mander BA, Rao V, Walker MP, et al. Prefrontal atrophy, disrupted NREM slow waves and impaired hippocampal-dependent memory in aging. Nat Neurosci 2013;16:357-364.

34.Mander BA, Marks SM, Walker MO, et al. Beta-Amyloid disrupts human NREM slow waves and related hippocampus-dependent memory consolidation. Nature Neuroscience 2015;18:1051-1057.

35.Kang JE, Limm MM, Holtzmann DM, et al. Amyloid-(beta) dynamics are regulated by orexin and the sleep-wake cycle. Science 2009;326:1005-1007.15

36.Yaffe K, Falvey CM, Hoan T. Connections between sleep and cognition in older adults. Lancet Neurol 2014;10:1017-1028.

37.Gagnon JF, Vendette M, Postuma RB, et al. Mild cognitive impairment in rapid eye movement sleep behavior disorder and Parkinson's disease. Ann Neurol 2009;66:39-47.

38.Ju YES, McLeland JS, Toedebusch CD, et al. Sleep quality and preclinical Alzheimer disease. JAMA 2013;70:587-593.

39.Riemersma-van der Lek R, Swaab DF, Twisk J, Hol EM, Hoogendijk WJG, Van Someren EJW. Effect of Bright Light and Melatonin on Cognitive and Noncognitive Function in Elderly Residents of Group Care Facilities. A Randomized Controlled Trial. JAMA 2008;299:2642-2655.

40.Guarnieri B, Musicco M, Caffarra P, et al. Recommendations of the Sleep Study Group of the Italian Dementia Research Association (SINDem) on clinical assessment and management of sleep disorders in individuals with mild cognitive impairment and dementia: a clinical review. Neurol SCi 2014;35:1329-1348.

41.Somers VK, Dyken ME, Mark AL, Abboud FM. Sympathetic-nerve activity during sleep in normal subjects. New England Journal of Medicine 1993;328:303-307.

42.Zunzunegui C, Gao B, Cam E, Hodor O, Bassetti CL. Sleep disturbance impairs stroke recovery in the rat. Sleep 2011;34:1261-1269.

43.Hodor O, Palchykova S, Baracchi F, Noain D, Bassetti CL. Baclofen facilitates sleep, neuroplasticity, and recovery after stroke in rats. Ann Clin Transl Neurol 2014;in press.

44.Pace M, Adamantidis AR, Facchin L, Bassetti C. Role of REM Sleep, Melanin Concentrating Hormone and Orexin/Hypocretin Systems in the Sleep Deprivation Pre-Ischemia. Plos One 2017;DOI:10.1371/journal.pone.0168430.

45.Pincherle A, Pace M, Sarasso S, Facchin L, Dreier JP, Bassetti CL. Sleep, Preconditioning and Stroke. Stroke 2017;48:3400-3407. 46.Mensen A, Pigorini A, Fcachin L, al. e. Sleep as a model to understand neuroplasticity and recovery after stroke: Observational, perturbational and interventional approaches. J Neuroscience Metods 2019;313:37-43.

47.Wu MP, Lin HJ, Weng SF, et al. Insomnia subtypes and the subsequent risks of stroke: report from a nationally representative cohort. Stroke 2014;45:1349-1354.

48.Leng Y, Cappuccio FP, Wainwright NW, et al. Sleep duration and risk of fatal and nonfatal stroke. Neurology 2015;84:1072-1079. 49.Alexiev F, Brill AK, Ott SR, Duss S, Schmidt M, Bassetti C. Sleep-disordered breathing and stroke: chicken or egg? J Thorac Dis 2018; doi:10.21037/ jtd.2018.12.66.

50.Birkbak J, Clark AJ, Rod NH. The Effect of Sleep Disordered Breathing on the Outcome of Stroke and Transient Ischemic Attack: A Systematic Review. J Clin Sleep Med 2014;10:103-108.

51.Parra O, Sanchez-Armengol A, Capote F, et al. Efficacy of continuous positive airway pressure treatment on 5-year survival in patients with ischaemic stroke and obstructive sleep apnea: a randomized controlled trial. J Sleep Res 2015;24:47-53.

52.Pace M, Camilo MR, Seiler A, et al. Rapid eye movements sleep as a predictor of functional outcome after stroke: a translational study. Sleep 2018;20:1-11.

53.McEvoy RD, Antic NA, Heeley E, et al. CPAP for Prevention of Cardiovascular Events in Obstructive Sleep Apnea. New Engl J Med 2016;375:919-931.

54.Brill AK, Horvath T, Seiler A, et al. CPAP as treatment of sleep apnea after stroke. A meta-analysis of randomized trials. Neurology 2018; in press.

55.Bravata DM, Sico JJ, FRagoso CA, al. e. Diagnosing and Treating Sleep Apnea in Patients With Acute

Cerebrovascular Disease. JACC 2018;7:e008841. DOI: 008810.001161/JAHA.008118.008841.

56.Khatami R, Luca G, Baumann CR, Bassetti CL, et al. The European Narcolepsy Network (EU-NN) database. JSR 2016;25:356.363. 57.Bassetti CLA, Adamantidis AR, Burdakov D, et al. Narcolepsy. Clinical features, etio-pathophysiology, diagnosis and management of a hypothalamic, immune-mediated disease. Nature Rev Neurol 2019; (in press).

58.Marti I, Valko PO, Khatami R, Bassetti CL, Baumann CR. Multiple sleep latency measures in narcolepsy and behaviourally insufficient sleep syndrome. Sleep Med 2009;10:1146-1150.

59.Kornum BR, Knudsen S, Ollila HM, et al. Narcolepsy. Nature Reviews/Disaese Primers 2017;3:1-19.

60.Latorre D, Kallweit U, Armentani E, et al. Autoreactive T cells in narcolepsy patients target antigens of hypocretin-producing neurons. Nature 2018;562:63-68.

61.Kallweit U, Bassetti CL. Pharmacological management of narcolepsy with and without cataplexy. Ex Op 2017;18:809-817.





"RoNeuro" Institute for Neurological Research and Diagnostic, Cluj-Napoca, Romania

Tel.: 0374 46.22.22

str. Mircea Eliade nr. 37, 400364 Cluj-Napoca, România Fax: 0374.461.674; Email: receptie@roneuro.ro

www.roneuro.ro