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ORAL PRESENTATIONS - ABSTRACTS

1)

Chasing quantum gravity with a pencil

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I shall review why in Dirac materials, like graphene, special relativistic-like matter and space may emerge. I shall then show how aspects of gravity theories may be reproduced as well. These include effects typical of quantum fields in the presence of black-holes, leading to a Unruh-Hawking kind of phenomenon, as proposed by our research group at the MatFyz of Charles University, more than a decade ago.

I shall then mention as many fresh results of our group, as time permits. From the time-loop to spot torsion, to the generalized uncertainty principles of quantum gravity stemming from and underlying (lattice) length; from a model of grain-boundaries and their relation to (A)dS and Poincaré spacetime algebras, to Unconventional Supersymmetry and the role of the two Dirac points.

Acknowledgements: I am deeply indebted to my coauthors Pablo Pais, Paolo Castorina, Luca Smaldone, Giovanni Acquaviva, Ruggero Gabbriellini, Boris Ivetic, Salvatore Mignemi, Marcelo Ciappina, Adamantia Zampeli, Gaetano Lambiase and to Raffaele Agostino, Francisco Correa, Arundhati Dasgupta, Gaston Giribet, Artem Grebenko, Paco Guinea, Vit Jakubsky, Siddhartha Sen, Guillermo Silva, Maria Vozmediano and Jorge Zanelli, for the many stimulating and informative discussions on the experimental and theoretical physics of this “graphene analog enterprise”.

Reference: [Most recent review] G Acquaviva, A Iorio, P Pais, L Smaldone, Hunting Quantum Gravity with Analogs: The Case of Graphene, Universe 8 (2022) 455

2)

ELI Beamlines: the high-peak, high-average power laser facility of the Extreme Light Infrastructure

Daniele Margarone

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The ELI Beamlines Facility is a pillar of the ELI (Extreme Light Infrastructure) ERIC pan-European Research Infrastructure hosting the world's most intense laser sources. ELI Beamlines developed and operates four cutting edge high-peak, high-average power femtosecond laser systems and offers a unique combination of primary (lasers up to 10 PW peak power) and secondary (high-energy particles and X-rays) sources to the international user community. Currently, several beamlines are operational and being upgraded to reach their full performances, while other beamlines are in their commissioning phase. Laser-driven particle accelerators have gained interest in the recent years thanks to their versatility and innovative features. This interest has pushed forward the development of beamlines where users can exploit the unique parameters (e.g. ultrashort bunch duration and ultrahigh dose rate) of laser-driven particle accelerators (ion and electron beams) and radiation (XUV to gamma-ray sources) for a wide range of applications. The current performance of particle and radiation sources available at the ELI Beamlines user facility will be presented and discussed along with their potential use for multidisciplinary applications. The high repetition rate capability of the available primary and secondary sources will be highlighted in combination with a range of advanced target delivery solutions and diagnostics in operation in extreme laser-plasma conditions ($>10^{21}$ W/cm² at >1 Hz and $>5 \times 10^{18}$ W/cm² at 1 kHz)

3)

Commissioning experiments at the ELIMAIA user beamline

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The ELI Multidisciplinary Applications of laser-Ion Acceleration (ELIMAIA) beamline is the laser-driven ion target area available at ELI Beamlines facility (Czech Rep.), having as final goal to provide a fully characterized and tuneable ion source generated by the HAPLS PW-class laser (>10 J in <30 fs), working at relativistic intensities ($>10^{21}$ W/cm²) and in repetition-rate (up to 10 Hz), for multidisciplinary applications.

We will report here the commissioning of the beamline, necessary to evaluate the performances of the system. Using targets of different composition and thickness, we were able to optimize the performance of the laser-plasma Ion Accelerator, obtaining in the optimal case proton energy cut-offs well beyond 30 MeV fluxes above 10^{11} /sr/shot. Moreover, we have demonstrated an excellent reliability and shot-to-shot stability (1-2% in energy) of the Ion Accelerator up to a repetition rate of 0.5 Hz for several hundreds of consecutive shots, along with on-shot target positioning and data acquisition and analysis systems.

These results demonstrate the robustness of the developed technology available for users at the ELI-Maia beamline, thus paving the way towards its future use for fundamental and applied research, including biomedical ones.

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4)

A multi-MeV alpha particle source via proton-boron fusion driven by a 10-GW tabletop laser

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Nuclear fusion between protons and boron-11 nuclei has undergone a revival of interest thanks to the rapid progress in pulsed laser technology. Potential applications of such reaction range from controlled nuclear fusion to radiobiology and cancer therapy. A laser-driven fusion approach consists in the interaction of high-power, high-intensity pulses with H- and B-rich targets. We report on an experiment exploiting proton-boron fusion in CN-BN targets to obtain high-energy alpha particle beams (up to 5 MeV) using a very compact approach and a tabletop laser system with a peak power of ~ 10 GW, which can operate at high-repetition rate (up to 1 kHz). The secondary resonance in the cross section of proton-boron fusion (~ 150 keV in the center-of-mass frame) is exploited using a laser-based approach. The generated alpha particles are characterized in terms of energy, flux, and angular distribution using solid-state nuclear-track detectors, demonstrating a flux of ~ 105 particles per second at 10 Hz, and ~ 106 per second at 1 kHz. Hydrodynamic and particle-in-cell numerical simulations support our experimental findings. Potential impact of our approach on future spread of ultra-compact, multi-MeV alpha particle sources driven by moderate intensity (10^{16} - 10^{17} W/cm²) laser pulses is anticipated.

Acknowledgements: This research was funded by the Ministry of Education, Youth, and Sports of the Czech Republic through the project “Advanced Research Using High-Intensity Laser-Produced Photons and Particles” (CZ.02.1.010.00.016_0190000789), the European Union's Horizon 2020 research and

innovation program (Grant agreement no. 871124, LASERLAB-EUROPE), the project “Target Engineering for Proton-Boron Nuclear Fusion Studies” sponsored by the UK Royal Society, and the EUROfusion Consortium, which was funded by the European Union via the Euratom Research and Training Program (Grant agreement no. 101052200—EUROfusion).

5)

The FLUKA.CERN collaboration

Roberto Versaci, A. Cimmino, D. Horvath, B. Lefebvre

ELI Beamlines Facility, The Extreme Light Infrastructure ERIC, Dolny Brezany TESTO:

In physics, Monte Carlo simulations are an effective way to calculate macroscopic quantities starting from the description of elemental phenomena. They have been at the hearth of scientific research in nuclear and particle physics for the past 80 years.

ELI Beamlines is a founding member of the FLUKA.CERN collaboration, which develops the FLUKA Monte Carlo code, a worldwide-recognized standard. The FLUKA Monte Carlo code is able to simulate the transport of several tens of particles: from the light neutrinos to the heavy ions, including the nasty low-energy neutrons. It is able to simulate a large amount of fundamental processes: electromagnetic interactions of photons, electrons, and muon; hadron-hadron, hadron-nucleus, and nucleus-nucleus interactions, low-energy neutrons interactions, and particles decays. The code is able to simulate electric and magnetic fields.

FLUKA has a wide range of applications including but not limited to particle physics, accelerator design, detector development, radiation protection, dosimetry, radiation damage, cosmic rays, and medical applications.

This contribution will provide an introduction to the FLUKA Monte Carlo code and its many applications. Then it will describe the activities of the ELI Beamlines group particularly focusing on those involving collaborations with external institutes.

6)

Radiation protection at ELI Beamlines

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In 2021, the European Commission established the Extreme Light Infrastructure (ELI) project as an European Research Infrastructure Consortium (ERIC), thus creating a one-of-a-kind legal entity, ELI ERIC. ELI ERIC is the largest international civilian laser-based user facility network. The establishment of ELI ERIC enable researchers and industries to access the world’s largest collection of high-power and ultra-fast lasers. Located in the Czech Republic, the “ELI Beamlines” pillar of the consortium, aims at investigating high-field high-density physics, developing high-brightness sources of X-rays, as well as secondary proton, electron, and ion beams, for interdisciplinary applications in physics, medicine, biology, and material sciences. Lasers as powerful as those used in the facility will generate a significant amount of ionizing radiation that can dangerous to both humans (personnel, visitors, and general public) and technology. The task of the ELI Beamlines radiation protection group is to guarantee that the whole laboratory and its surroundings are safe from the produced ionizing radiation. In this

contribution, the radiation protection challenges will be discussed and the mitigation strategies implemented will be presented.

7)

The long march toward zero friction

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Friction causes massive energy dissipation and mechanical abrasion between machine component parts in the world every year (costing approximately $3.3 \cdot 10^{13}$ kWh[1]). Understanding the mechanism of the frictional processes and searching for an optimum material combination, ideally providing a near frictionless state, are thus essential. Recent works show that near-zero interface friction can be realized in twisted 2D materials or 2D heterostructures due to their ultra-flat interface potential energy surface. However, the origin of friction is complex. Suppressing one primary source will make friction from other mechanisms to surface. Thus, reaching a pure zero friction state is going to be a long march.

In this talk, I will report the friction phenomenon in superlubric MoS₂/graphite and MoS₂/h-BN van der Waals heterostructure interfaces. In such systems, instead of reaching a frictionless state with a suppressed interface friction, mechanisms like the edge pinning effect[2] and the shape of the structural potential energy[3] start to dominate the friction processes and provide friction during the sliding and twisting. Those phenomena explain why it is hard to realize low friction on the macro scale. I will also further discuss the main challenges and provide possible ways toward a pure zero friction state[4].

References:

[1] K. Holmberg, A. Erdemir, "Influence of tribology on global energy consumption, costs and emissions". *Friction*, 5, 263 (2017).

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[3] M. Liao, A. Silva, L. Du, P. Nicolini et al., "Twisting Dynamics of Large Lattice-Mismatch van der Waals Heterostructures". *ACS Appl. Mater. Interfaces*, 15, 19616 (2023).

[4] M. Liao, P. Nicolini, T. Polcar, "Separating anisotropic and isotropic friction between atomic force microscope tips and atomically flat surfaces". Under review in *Phys. Rev. B* (2023).

8)

Turbulent flows of superfluid helium-4: an introduction

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The investigation of turbulent flows of superfluid helium-4 is an active and challenging research field,

which is not only interesting in its own right but that has also broader implications, related, for example, to neutron stars or to turbulence in classical viscous fluids, such as water and air. Indeed, superfluid helium-4 is characterized by unique properties, e.g. in certain conditions, it apparently flows without friction, as if it were a perfect (inviscid) fluid. In order to explain some of its remarkable features, it is often described as a quantum liquid, i.e. its turbulent flows are defined by the presence of quantized vortices, which are line singularities within the superfluid and can be viewed as the carriers of the flow vorticity. Special emphasis is given to the visualization of turbulent flows of superfluid helium-4, which in recent years has significantly contributed to our understanding of the underlying physics (the first laboratory in Europe to be exclusively devoted to such measurements has been established at Charles University: its design and implementation was the focus of my work, when I was hired in 2009, and I am now responsible of its scientific activities). Selected experimental findings are reviewed, especially to highlight close similarities, as well as striking differences, with classical turbulent flows of viscous fluids. For example, it has been shown that quantum features are apparent at flow scales smaller than the mean distance between quantized vortices, while, at larger scales, a classical-like picture is observed in some cases, especially when thermal effects can be neglected.

9)

Uncovering the Hidden Potentials of Materials: Exploring the Electrical Properties of the objects That Surround Us

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In the last 15 years, I have been on a journey exploring the properties of various materials commonly used in daily life, but often overlooked by the public. This journey has taken me from the lab bench to the real world, where my colleagues and I have applied our knowledge to try developing innovative solutions.

I started by working on conductive graphite embedded in polymers for applications such as electromagnetic shielding, antistatic coatings, and barriers against gas or fire. However, low conductive, insulating materials also play a critical role in modern technology. For example, high permittivity barium titanate embedded in plastics is essential for flexible electronics, where the ability of preventing the passage of electric current and transmit the electromagnetic field is crucial in devices like transistors.

More recently, I have worked on cement/carbon nanotube composites for self-sensing applications, where the materials themselves can detect chemical and structural changes and communicate them to supervisors. In this case, the electrical performance helps maintain buildings and urban elements, increasing their lifetime while reducing costs and pollution.

I have also explored the potential of collagen/chitin nanocrystals as scaffolds for tissue regeneration, where their piezoelectric properties determine the quality and speed of the healing process. Mechanical stress applied to the material generates a small electric current that stimulates cell growth, reproduction, and tissue repair.

In this presentation, I will share my journey and showcase some of the applications developed using these often-overlooked materials in solving real-world problems.

Acknowledgements:

I want to thank the department leader Dr. Jiří Pflieger and all the colleagues of the department of Polymers for Electronics and Photonics for their scientific contributions. For the financial support, my thanks go to MYAS, GAČR and TAČR who funded our research projects all these years.

10)

Hydrogen sulfide and sperm biology: from poison to remedy

Eliana Pintus and José Luis Ros-Santaella

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Hydrogen sulfide (H₂S) is a colorless and water-soluble gas that typically smells like rotten eggs and is naturally found in volcanic springs and decomposed organic matters. After being regarded as a mere poison for decades, H₂S is now recognized as an endogenous gaseous molecule having both a physiological and pathological role in cells and tissues. Because of its antioxidant properties, H₂S has shown the ability to palliate the damages induced by oxidative stress, which is characterized by an imbalance between reactive oxygen species (ROS) and antioxidant levels. Sperm cells are particularly susceptible to the injuries provoked by oxidative stress because of their unique plasma membrane composition and their limited antioxidant defense. Although small levels of ROS are physiologically required for sperm function and fertilization process, their overproduction, which can arise because of pathological conditions or semen manipulation for assisted reproductive techniques, is detrimental for the male fertility.

In this presentation, I will overview the most recent advances about the role of H₂S on the male reproductive system, the effects of different H₂S donors on sperm quality and their potential application for treating sperm fertility disorders, with a special focus on those associated with oxidative stress. Overall, we found that H₂S protects sperm cells against the deleterious consequences of oxidative stress, although the effects are both donor and dose dependent. Moreover, some H₂S donors have shown a clear positive effect on sperm mitochondrial activity, which can be particularly useful to explore the role of H₂S in sperm mitochondria-dependent events such as apoptosis or fertilization.

Acknowledgements: We would like to thank our former and current Ph.D. students and our colleagues Prof. Matson (Virginia Polytechnic Institute and State University, USA), Prof. García-Vázquez (University of Murcia, Spain), and Dr. Novy (Czech University of Life Sciences Prague) for their collaboration.

11)

Implication of sperm size and shape in animal andrology

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Spermatozoa are the most diverse cells in the animal kingdom. They are ejected from the body into an unknown environment where they live as free-living organisms. Accordingly, spermatozoa of each species are adapted to their specific fertilization environment. The environments where sperm cells carry out fertilization are very diverse, so they have developed very specific adaptations to them, with a high variability in their structural components that result in a high morphological diversity among animal species. Sperm size also drastically varies across species ranging from 6 to ~60,000 µm, being the longest sperm found in the tiny fruit fly *Drosophila bifurca*. Intraspecific studies have investigated whether sperm size determines sperm lifespan, velocity, and fertilization capability, but, in general, contradictory results or no correlations were found. Similarly, studies concerning the implication of

sperm size in their ability to withstand the cryopreservation process are scarce and focused on the sperm head only.

The present communication aims to describe the relationships between intra-male variation in sperm size and sperm quality, the implication of sperm cell dimension in sperm freezability, and the effects of nutrition on sperm shape. Finally, we aimed to assess the main sperm traits in unique species with interest in conservation breeding programs. Overall, we found that the sperm size homogeneity is related to increased testis size, spermatogenic activity, and sperm quality. Our results also show a strong relationship between the volume of the sperm flagellum with sperm freezability, that is, sperm with a higher flagellum volume freeze worse. Moreover, we reported that a high-energy diet administered to pubertal fallow deer promotes an increase in the sperm midpiece where the mitochondria are located. The assessment of sperm morphometry can reveal new insights into the biological processes and the multiple adaptations to the fertilization environment of these cells.

Acknowledgements: We would like to thank our former Ph.D. students and our colleagues from Czech University of Life Sciences Prague (Czech Republic), University of Murcia (Spain), and University of Castilla-La Mancha (Spain) for their collaboration.

12)

Thermo-hydro-mechanics of landslides under climate change

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Landslide initiation and dynamics are investigated with methods that are specific to the scale of investigation. Hydro-mechanical models are the go-to for individual landslides as they rely on the well-studied coupling between hydrological input – via atmosphere-soil interaction – and mechanical response. At larger scales, these models gradually give way to simpler physics-based approaches and finally to geostatistical models. The role of temperature in controlling evapotranspiration and thus the hydrological balance is well recognized and featured in models; conversely, direct thermo-mechanical couplings are systematically neglected unless changes in water phase are involved. This contrasts with experimental evidence of a fully-coupled thermo-hydro-mechanical behaviour of most geomaterials. Temperature-dependent processes are potentially relevant to slope stability, especially in clayey slopes in temperate and warm regions.

Here, the thought-provoking hypothesis is that temperature fluctuations and trends induced by climate change may cause, in short to long terms, a disturbance to mechanical equilibria in slopes (by altering permeability, water retention capacity, compressibility, shear strength). Together with other known effects, such as altered precipitation patterns and changes in land use, this disturbance could affect landslide activity and the distribution and frequency of slope movements. To verify this hypothesis across the scales, systematic field monitoring of temperature-related variables is necessary, together with geostatistical analyses entailing thermal remote sensing products. At the same time, fully-coupled approaches need to be upscaled to permit physically-based catchment- or regional-scale studies accounting for appropriate temperature-related variables and the inherent heterogeneity in materials and boundary conditions.

13)

Metal soaps in oil paintings over the centuries

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Metal carboxylates formation is a well-known issue in oil paintings, and it is one of the most recurrent topic in conservation science research of the last 20 years (1). During the lifetime of an oil painting, free carboxylic acids ($R-COOH$) derived from drying oils can react with the metal cations from some pigments (e.g., lead-based and zinc-based ones) leading to the formation of metal carboxylates, which are characterized by the general formula $M+(RCOO-)$ (1). These products of pigment-binder interaction can reveal themselves in various ways, namely, aggregates, efflorescent crusts, crystalline exudates, blisters, and cracks (2). These phenomena have been detected in numerous paintings and are now acknowledged as a part of the dynamics of the aging of oil paints (2). Despite numerous researches focused on this degradation process, there are still many unresolved questions concerning the formation and behaviour of metal carboxylates. In this presentation, the symptoms manifested by the presence of crystalline metal carboxylates (metal soaps) together with the detailed characterization of saponified paint layers will be showed. Furthermore, a comparison of different analytical methods involving X-ray fluorescence (XRF), X-ray powder diffraction (XRPD) and micro-spectroscopic methods ($\mu FTIR$ and $\mu Raman$) will be provided, highlighting both their advantages and limitations. The study of the saponification processes will be also described within the experimental drying of model paints to reproduce and monitor both the reactions and the factors influencing or inhibiting the process (e.g., relative humidity, temperature, light, and so on).

References:

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2. Izzo, F. C. et al. ChemistryOpen 10, 904–921 (2021)

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14)

Multi-text medieval manuscripts between textual criticism, codicology and paleography: the case of XV century manuscript Florence, Biblioteca Marucelliana, C 150, containing philosophical and rhetorical texts in vernacular Italian.

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When dealing with complex objects such as multi-text manuscripts, that is to say the most common form of book in the late Middle Ages and Renaissance, only a multidisciplinary approach that includes textual criticism, codicology, paleography and other field of studies (such as history of the language, history, source studies), can provide a historically aware understanding of the object itself, which quite often is the result of multilayered juxtapositions in a synchronic and diachronic way. The little known XV century manuscript preserved in Florence, Biblioteca Marucelliana, C 150, containing philosophical and rhetorical texts in vernacular Italian, will be used as case study in order to show how to untangle complex structures.

[keywords: manuscript studies, textual criticism, codicology, paleography, history of the language, history, literature]

15)

Giuliano de' Ricci reader of a lost work.

Ludovica Radif

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As a extremely nuanced and fascinating but also potentially close to our sensibilities. The beginning of the 16th century marks a turning point that I would call “authorial”, with a hermeneutic maturity compared to the past, and it is precisely when the text we are discussing today is located. A borderline case that a philologist can come across. The soul and skeleton of his analyses are the manuscripts, be they part of a direct tradition or an indirect one, such as quotations, a manuscript (or printed text) that at least partially reproduces the text. Nevertheless, in this case we are going to examine, we are not in such a situation, we do not possess any manuscript, but we do know one of his readers, who tells us about it. It deals with a very famous and important author on the European scene and a work that I am sure was very original and unique in the scenario of the time: author was Niccolò Machiavelli founder of political theory; the text is the lost comedy *The Masks*; the play was inspired by the ancient comedy of Aristophanes, something special in the context of his production. The *Histories of Italian literature* usually dedicate very few lines to that “literary failure”. We certainly face a gap, but this gap in my opinion can be at least partially filled. It is not only the words expressed (in Biblioteca Nazionale Centrale di Firenze, MS Palatino, Grande Priorista a famiglie E.B. 14.1/I-IV: I, f. 160v) that are eloquent, but perhaps even more so what is kept silent. We must therefore consider Machiavelli as a not unremarkable representative of Aristophanes’ revival, *Fortleben*, during the Renaissance.

16)

The evolution of political families in Central and Eastern Europe: Dynamics of electoral volatility and political orientation

Mattia Collini

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Party politics in Central and Eastern Europe has been characterized by instability, fragmentation and volatility, at least when compared with the traditionally more stable Western Europe. However, are these phenomena affecting all in the same way or can we see some differences among ‘blocs’ or ‘political families’? This question is particularly relevant in a time of political changes both in Eastern and Western Europe. This paper aims to be a comprehensive assessment of these phenomena, based on empirical evidence from the region. Blocs are a key aspect of the research, which explores how much the concept of ‘bloc’ can be applicable, and what are the main characteristics of old and new political families in CEE. Thus, the first goal of the paper is to present a comparative diachronic analysis of the electoral dynamics of political blocs/families in Central and Eastern Europe. The paper covers national elections in six Central and Eastern European countries (Czech Republic, Slovakia, Poland, Romania, Bulgaria, Hungary) from their democratic transition to 2020. This long timespan allows tracing the entire evolution of party systems and political families from their origin to the present days.

Electoral volatility, (intra)bloc fragmentation, and political positions, are considered among the main variables to assess whether there are differences and/or recurring trends within the systems, as well as the emergence of newcomers. In particular, regression analyses will be used to search for correlations.

17)

The smoldering inflammation and the route to cancer.

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Chronic inflammation is recognized as a cancer-promoting factor. The full range of inflammatory processes is still under definition. Local changes of immune activation can induce fast collagen remodeling in the tissue. Reciprocally, collagen accumulation can affect the local immunity. We have shown that immunological activation of a tissue (by bacteria or chemicals) quickly modifies the colon mucosa structure. Using a rat model of chronic colitis (dextran sodium sulphate – DSS – induced colitis) and of carcinogenesis (azoxymethane – AOM - carcinogen for the colon) we have shown that, in both models, inflammation produce remodeling of the collagen scaffold, even when the mucosa appears recovered at the distance of 1 month from the acute induction. This occur despite apparently normal or reduced levels of local cytokines as a smoldering inflammation. Immunological data and confocal microscopy suggest that normally the microbiota can elicit homeostatic regulation in the healthy rat colon mucosa. The modulation can be due to an “inflammatory threshold” establishing the range of tolerance toward the pro-inflammatory activation. The smoldering inflammation has dis-balance between pro-inflammatory and regulatory signals and, despite low cytokine levels overcomes the inflammatory threshold. Underhand, smoldering inflammation is capable to continue to damaging the tissue structure and leading to either chronic inflammatory diseases or cancer development.

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18)

The different layers of pathological mechanosensing.

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The onset and progression of aging pathologies is highlighted by continuous extracellular matrix (ECM) transformation. ECM pathological remodeling results in dramatic changes in the chemistry, local stiffness, and nanostructure of the organ scaffold. Deposited by the tissue fibroblasts, the newly produced matrix displays impaired compliance and aberrant signalling, thus hinders cell functionality. The response of tissue cells to the ensuing biomechanical stress leads to the alteration of their transcriptional and post-transcriptional landscape, with the aim of preserving organ function (1).

Our group has tried to dissect the processes driving ECM remodelling and unveil the molecular pathways highlighting cell response to the new conditions. Cardiac disease represents a paradigmatic example of how ECM remodelling hinders cell functionality, here affecting muscle contractility and organ pumping function. Through human specimens and induced pluripotent stem cells, we identified reproducible modifications in the 3D nanostructure and mechanics of heart ECM that contribute to organ failure (2).

We found the hyperactivation of Hippo effector YAP in patient-derived cardiac fibroblasts promotes ECM pathological remodelling and favors the fibrotic process and fuels heart failure. Additionally, we showed ECM maladaptive remodelling and YAP reactivation drive dramatic changes in cardiomyocyte transcriptional and post-transcriptional landscape to increase their contractility and preserve organ pumping function. We also highlighted how this process rewires the alternative splicing of numerous genes involved in cardiomyocyte contractility, calcium handling and mechanosensing (3).

ECM remodelling is also key to tumorigenesis, so that altered matrix stiffness is considered predictive of malignancies. We analyzed primary prostate cancer tissues obtained from patients undergoing radical prostatectomy to underline reproducible structural changes in the tumor ECM and promoting the loss of the glandular architecture. We established patient-derived prostate cancer tumoroids (PCTs) that mimic key features of the native tumor microenvironment (TME) and proceeded to investigate the cooperative role of TGF- β signalling and ECM *desmoplasia* in fostering tumor progression and dissemination (4).

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19)

An induced pluripotent stem cell-based model to study the role of EXTRACELLULAR MATRIX in myocardial fibrosis.

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Cardiac fibrosis is the consequence of chronic insults to the myocardium, characterized by the abnormal accumulation of extracellular matrix (ECM). Differentiation of cardiac fibroblasts (cFbs) into myofibroblasts drives the pathological ECM remodelling, a process highlighted by biochemical and structural changes which compromise cardiomyocyte (CMs) activity and lead to heart failure. Here,

we adopted induced pluripotent stem cells (iPSCs) to investigate how fibrotic ECM affects CMs pathophysiology. We derived cFbs from induced pluripotent stem cells (iPSCs-cFbs) and optimized a protocol to differentiate them into myofibroblasts based on the fine-tuning of bFGF and TGF- β 1 signalling pathways. Next, we obtained ECM by implementing a decellularization procedure of either quiescent or activated iPSCs-cFbs and analyzed the pathological changes occurring during the deposition of diseased cardiac ECM. Then, we developed iPSCs-CMs and cultured them on healthy or fibrotic decellularized ECM (dECM). Flow cytometry was adopted to study how the biomechanical properties of dECM affect CMs phenotype. We further validated our model using iPSCs-CMs derived from Duchenne muscular dystrophy (DMD) cardiopathic patients. By capitalizing on this approach, we might be able to recapitulate the accumulation of fibrotic tissue during heart diseases and investigate the contribution of pathological ECM to the progression of many cardiac disorders. Indeed, dECM derived from cFbs (dECM-cFbs) represents a bio-inspired scaffold for iPSCs-CMs culture, proposed as an animal-free, isogenic model of cardiac fibrosis for personalized medicine applications. *Adnowledgements: This project has received funding from the European Union's Horizon 2020 research and innovation programme under the Marie Skłodowska-Curie grant agreement No 860715. The material presented and views expressed here are the responsibility of the author(s) only. The EU Commission takes no responsibility for any use made of the information set out.*

20)

Primordial proteins: simple but not simpler

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All extant cells known to humankind build proteins from the same 20 coded amino acids. However, the study of origins of life implies that earlier cells functioned with a smaller alphabet, before the fixation of the Central Dogma. This profound evolutionary transition in our cells' history therefore raises urgent questions: how could early proteins support a proto-biosphere and how easy or hard is it to build a functional protein from a prebiotically plausible alphabet?

I will uncover the properties of proteins from an early subset of the canonical amino acids and additionally, the effects of selected unnatural (prebiotically plausible) amino acid incorporation will be elaborated. Besides the impact of different amino acid alphabets on the peptide/protein biophysical properties, we have inspected the capacities of prebiotically plausible sequences to bind RNA as representatives of their early functional potential.

Our work indicates that structured conformations were readily available already to early protein alphabets, capitalizing on interactions that are less frequent or rare in today's biology. In addition, we have identified factors which have likely contributed to the selection of the canonical alphabet from the prebiotically plausible alternatives.

21)

Applications and development at IPHYS Bioimaging facility

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The IPHYS facility provides an extensive light microscopy and data analysis service. The facility belongs to the Czech-Bioimaging infrastructure and actively participates in Prague Node – EuroBioimaging. It participated in research areas such as confocal microscopy, label-free microscopy, correlative microscopy or stereology. The activities of facility will be presented on two selected examples covering Cajal body formation (Davide Basello) and the development and integration of STED modality into the existing customized Bruker Ultima microscope (Daniel Hadraba).

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22)

Development of a new optogenetic brain-on-a-chip approach to study GABA/NMDAR altered ratio in Schizophrenia.

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Schizophrenia affects about 1% of the worldwide population and includes impaired decision-making and cognitive inflexibility. These dysfunctions are due to an unbalanced excitation/inhibition in the neural transmission of the prefrontal cortex and the hippocampus. GABA is an inhibitory neurotransmitter, while NMDAR is a glutamatergic receptor involved in cognitive processing. Optogenetic stimulation of GABA showed to be effective in re-establishing excitatory/inhibitory balance. Although animal models study in vivo brain physiology, the real-time observation of neural circuits is hampered by the low accessibility to deeper areas. The fabrication of nanofluidic chips advanced innovative in vitro-silico models for neural circuits (brain-on-a-chip, BoC), allowing a high-level time/space resolution. Within the 3Rs concept, neural systems built on microchips allow a significant leap forward in developing human-relevant brain models of pathologies. The following logical step is to integrate the optogenetics technique with a BoC, exploiting the high target reliability of optogenetics.

23)

Deletion of beta2-containing nicotinic acetylcholine receptors in striatal interneurons controls striatal-dependent behaviors.

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The importance of cholinergic signaling in the striatum for behavioral control and cognition has been extensively studied, but little is known about the functional role of nicotinic acetylcholine receptors (nAChRs) expressed by local neurons. The principal striatal neurons, the medium spiny neurons

(MSNs), express only very low levels of nAChRs, hence the vast majority of nAChRs are expressed by striatal interneurons (INs), either cholinergic (CINs) or GABAergic (GABAINs), that overall represent less than 5% of the striatal population. We hypothesize that acetylcholine released by CINs activates nAChRs expressed by CINs themselves and GABAINs and that this activation is important for modulating striatal-based behavior.

At first, we used double fluorescent in situ hybridization (FISH) targeting the most prevalent nicotinic subunit, beta2, in combination with markers identifying CINs or GABAINs, to define which neuronal types express nAChRs, in the mouse striatum. To determine the functional role of striatal nAChRs, we deleted beta2 nicotinic subunit by injecting Cre-expressing AAV viral vector into the dorsal striatum of beta2-flox/flox mice, which we tested in a battery of behavioral tasks. Mice with deletion of beta2* nAChRs showed increased anxiety-like behavior, together with a decrease in sociability ratio and a deficit in discrimination learning. As last task before sacrificing the animals, we tested the sensitivity to a stimulant drug, amphetamine, and analyzed the expression of a neural activity marker, c-Fos, in the dorsal striatum. The deletion of beta2* nAChRs increased amphetamine-induced hyperlocomotion along with c-Fos expression in MSNs and in striatal INs.

We conclude that beta2-containing nAChRs are primarily expressed by CINs in the striatum and that, even if in a small number, they modulate striatal signaling and striatal-based behavior.

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24)

Assessing the non-linearity in brain data across modalities.

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Resting-state fMRI functional connectivity (RSFC) is a technique used in neuroimaging to assess the functional connectivity between different brain regions while the subject is at rest. We can use RSFC to study the brain's intrinsic functional architecture and to investigate how brain regions communicate. For example, RSFC is useful in the study of several neuropsychiatric disorders, including epilepsy, depression, and schizophrenia. The most common way to estimate functional connectivity is Pearson's correlation of the BOLD signal between region pairs. However, correlation can account correctly only for linear dependences. In the last ten years, scholars proposed alternative methods for the estimation to address the limitations of correlation. One of the most adopted is the information-theoretic measure of Mutual Information. For infinite time series, MI can account for any possible pairwise interaction between the time series. Yet, in applications, the time series' length is limited by technical and conceptual reasons. We investigate the actual power of MI approaches on different brain data varieties. We go beyond functional connectivity exploring the amount of non-linearity observable in the brain, from fMRI data to EEG and iEEG. By comparing different MI estimators and correlation, we can learn some lessons about the actual usefulness of the newly proposed approaches.

25)

Vaccine Hesitancy and Medical Specialists

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Healthcare professionals represent one of the most influential groups of actors in childhood vaccination programmes. While recent social scientific scholarship focused its primary attention on the role of paediatricians (e.g., Deml 2020; Leask et al. 2012; Lehner et al. 2022), including vaccine-hesitant healthcare professionals (MacDonald, & Dubé 2015; Verger et al. 2015, 2016), the role of medical specialists remained unexplored. Against this backdrop, this contribution aims to systematically explore the varieties of involvement of medical specialists in childhood vaccination. More specifically, this contribution addresses the following questions: When do the parents and paediatricians address medical specialists concerning vaccination? What is the role of medical specialists concerning vaccination? How do paediatricians approach medical specialists? This study draws on ethnographic observations and semi-structured interviews with healthcare professionals in the Czech Republic, where childhood vaccination is mandatory. Four ideal-typical forms of interactions between paediatricians and medical specialists were identified, in particular, a) uncritical reliance on medical specialists, b) problematization of the role of medical specialists, c) contestation of medical specialists and d) the recognition of complexity, understanding and reflexive approach. Furthermore, the study suggests that the interactions between paediatricians and medical specialists contribute to negotiating and reproducing existing professional identities.