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### HIGHLIGHTS EDITORIAL TEAM

Petya Georgieva

Antonella Del Rosso

**ENLIGHT COORDINATOR** 

Manjit Dosanjh

Manjit Dosanjh

2018 ENLIGHT meeting, London, UK

DESIGN & LAYOUT Media Frontier

COVER:

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already be measured in terms of the legacy that we are (page 4) that took place in London with a special emphasis leaving behind us. Indeed, since the beginning, this network on our young participants and their research, the meeting has opened the way to initiatives that have contributed to that was held in Archamps (France) to discuss the PIMMS shape the current scenario of radiooncology. Among the 2 study, which is being supported by CERN (page 24). projects that have left a more visible mark is PARTNER, the the workshops in Australia, home of Bragg the magic EU-funded project we run ten years ago. PARTNER gave ingredient for hadron therapy (page 28), and in Lyon, the research and training opportunities to 29 young biologists, birthplace of very first use of radiation therapy for cancer engineers, physicians and physicists and allowed them to treatment (page 30). go on developing modern techniques for treating cancer. Since this issue goes out at Christmas time, I would like Today, these researchers are spread world-wide and have to extend my best wishes to all our members and their very interesting stories to tell (pages 8-17). Their stories families for a Merry Christmas and a Happy New Year! remind us of one of the pillars of our mission: educating Next year is already full of events for our community (page the next generation of experts in particle therapy and 27) and of course let me remind you of our next annual expose them to state-of-the-art technologies as early as meeting, which will be held in Caen (France) from 1 to 3 possible in their career. July, 2019.

Undoubtedly, another important achievement of our network are the strong links we have created among the different communities that compose ENLIGHT. They seem very "natural" and routinely applied now but, as Roberto Orecchia emphasises in his interview (page 22), they were definitely not a given when we started all this. What Roberto Orecchia describes in the article reminds us of what we need to accomplish our vision: traditionally separate communities can work together if they stay curious and creative but also pragmatic. A really exclusive blend of skills! Roberto's idea of a machine all radiooncologists are dreaming of will certainly trigger your curiosity.

As usual, I would like to conclude by thanking all the contributors and people in my team for their continued support, which makes it possible for us to share information among the members. I wish all of you an interesting reading!

Manjit Dosanjh

Masanih

## **2018 ENLIGHT LONDON** MEETING IN FOCUS





## ENLIGHT TRAINING DAY 2018





ENLIGHT dinner on the River Thames

This year, the European Network for Light Ion Hadron Therapy Annual Meeting and Training was hosted at University College London Hospital (UCLH) in the UK. UCLH, one of only two centres supported by the UK National Health System that will be offering proton beam therapy (PBT) in the near future, recently had its cyclotron lowered in to one of the deepest holes in central London. software, provided by Reynald Vanderstraeten from Varian Medical Systems. This session covered the impact and handling of range uncertainty in proton therapy planning, and the differences between robust and planning target volume optimisation. This exercise was a particularly valuable experience for myself as I am a medical physics PhD student with research interests in uncertainties in treatment planning.

At the beginning of the Training Day, we were guided to the The final workshop of the day was centred around the im-Cruciform Building, a stunning Grade II listed building located portant topic of clinical trials in proton therapy. We had an in the heart of London. Greeted with delicious pastries, bisintroduction to the CTRad strategy for PBT trials in the UK cuits and coffee, the morning session began with an eloquent from Prof Phil Evans and Maria Hawkins, and Gemma Eminoand welcoming introduction from Prof Manjit Dosanjh. This wicz presented us with the design and challenges of proton was promptly followed by talks from Richard Amos and Dr vs. photon clinical trials. Helen Bulbeck gave us a particularly Yen-Ching Chang on the physics and clinical potential of thought-provoking review of patient perspectives of proton protons, and their context in ion therapy treatment. therapy, and what it means to the general public following media coverage.

The Operational Lead of proton therapy physics at UCLH, Andrew Poynter, proceeded to give us an exciting update on the UCLH project: the developments in the construction of the new proton centre, and the required training of the staff to deliver the new modality of treatment.

Over the extended lunch break was the poster session. Within the Cruciform Building was a showcase of a variety of scientific posters submitted by attendees of the meeting presenting particle therapy related research projects. This proved to be an excellent opportunity to network, learn first-hand about other groups' exciting research and quiz the authors.

Following lunch, the practical session of the day involved a hands-on demonstration of Eclipse treatment planning



Young women scientists who took part in the ENLIGHT meeting













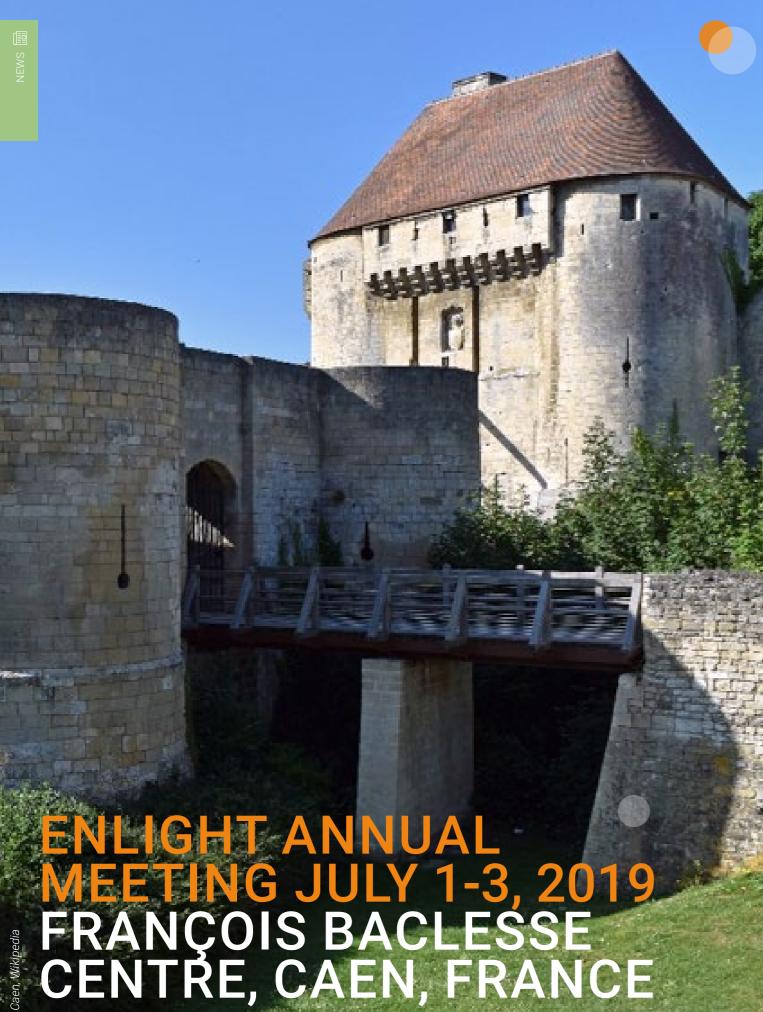




Johnny Lee of Radiotherapy Trials Quality Assurance (RTTQA) then gave us an entertaining and engaging understanding of the role of quality assurance in clinical trials, and Rush Patel provided an overview of the big data platforms and analysis in radiotherapy trials.

Megan Zoë Wilson, University College London | UCL

Discussions after the training day



Organisers : Jacques Balosso, Daniel Cussol, Manjit Dosanjh, Siamak Haghdoost, Yannick Saintigny, Juliette Thariat, Samuel Valable



Mont-Saint-Michel, Normandy, France

July 2019 will see the annual ENLIGHT meeting taking place experimental and clinical application of light ions over a large in Caen in Normandy, France, hosted by the University and scale of variety. the Cancer Centre François Baclesse of Caen. This location Caen and the Normandy Region are managing this project in has been chosen because Caen is hosting the third particle a rather favourable landscape including two university hostherapy centre of France, which opened to patient treatments pitals and two cancer centres in Caen and Rouen, the wellin protontherapy at the end of July this year. Two other cenknown research centres GANIL and CYCERON respectively tres, in Orsay and Nice, have been operating for 27 years. So, for nuclear physics and medical imaging of brain diseases. at the time of the meeting in Caen we will have a one-year experience to share with the participants.

This Centre, named CYCLHAD has a "one room protontherapy demonstrate the medical efficacy and utility of hadrontheraequipment" with a half revolution gantry of the Proteus®One py". This will be followed on Tuesday 2nd and Wednesday 3rd type. Actually, it is a far more ambitious project since the July by the two-day ENLIGHT meeting. building has been sized and built to host a second accelerator the C400, of a new type, to provide light ions in three different The training day will present topics on the currently most experimental rooms by the year 2023. It will by an isochrone critical item in particle therapy: the need for medical clinical super conducting cyclotron able to accelerate protons, He, data. Principles of clinical trials and advanced techniques for Li, B, C, N, O and Ne. Associated to the 4 beams/rooms of medical assessment will be presented and discussed. CYCLHAD, five hundred square metre laboratories will be The two meeting days will include fundamental topics on equipped to welcome local and visiting scientific teams. The physics, radiobiology including RBE determination, biology of CYLHAD centre will hopefully become a valuable centre for



The new CYCLHAD centre in Caen

The ENLIGHT meeting will start on Monday 1st July with a training day focusing on "How to produce clinical data to

radioresistance, quality assurance of beam path, advanced cyclotron technology, advanced positioning, ongoing clinical

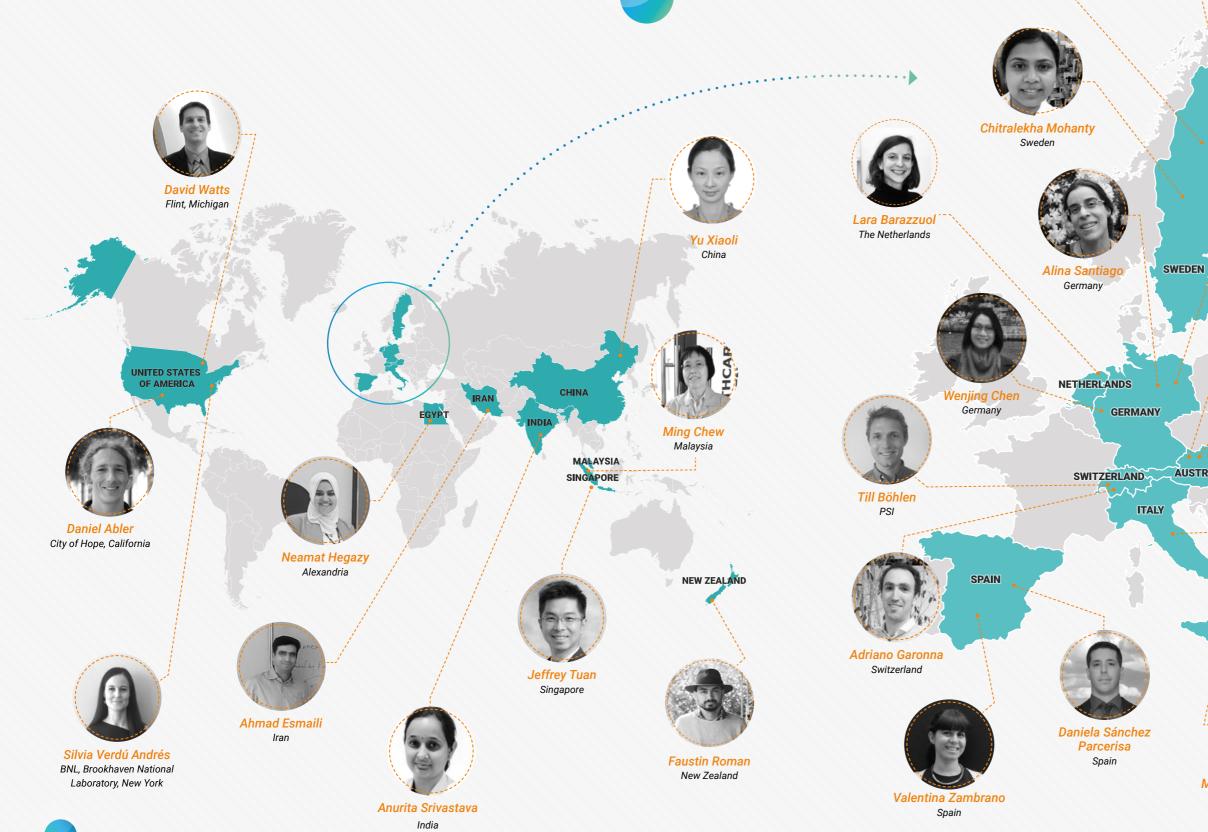
Jacques Balosso, Archade

## PARTICLE TRAINING NETWORK FOR EUROPEAN RADIOTHERAPY 10 YEARS LATER...



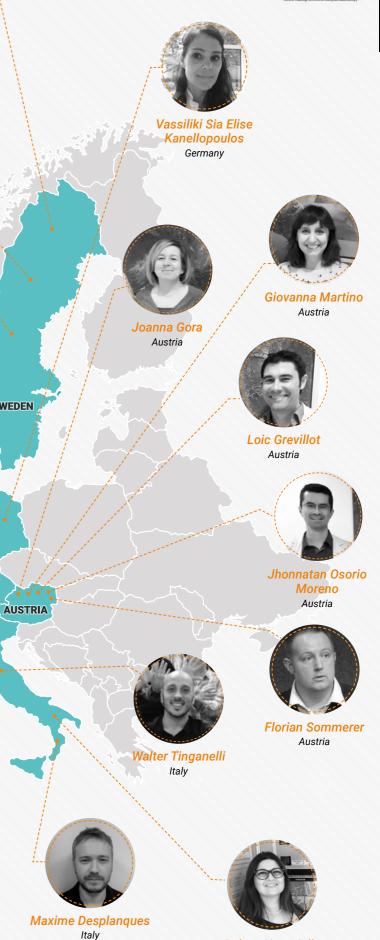


Marta Lazzeroni Sweden





ERSPECTIVE



Palma Simoniello Italy

### **PARTNERing in South-East Europe**

One of the early projects of ENLIGHT was PARTNER, a 4-year years ago, there is a lack of local expertise and the difficulty to Marie Curie Training project funded by the European Commission. For the first time, ten academic institutes and research centres and two leading companies were collaborating under the PARTNER umbrella to form a unique multidisciplinary and multinational European network. The project had the challenging goal of offering research and training opportunities to 25 young biologists, engineers, physicians and physicists and allow them to go on developing modern techniques for treating cancer.

Today, 10 years after the beginning of the project, the young researchers trained within PARTNER have become leading experts in various European hospitals, hadrontherapy centres or research institutes. Over the years, they have established a long-lasting network of specialised collaborators and they have continued to build expertise and train new researchers. In addition, the PARTNER's heritage is now becoming a model, which can be used to build expertise in other parts of the world and, in particular, in South-East Europe.

In many aspects, the current situation in the South-East Europe area presents analogies with the rest of Europe when, more than 10 years ago, ENLIGHT contributed to build the future of hadrontherapy. Today, in South-East European countries there is a strong political and scientific motivation to build a hadrontherapy centre which will allow patients in the region to society, including patients, physicians and scientists, with one be treated locally. However, similarly to central Europe several common initiative.

create links with experts spread across the rest of the world. To address the situation, the PARTNER project introduced a novel approach: starting building expertise by training young researchers.

Indeed, the common aim of the countries involved in the project of the South-East European hadrontherapy centre is to build competence, stop the brain drain, collaborate for a common good and, also, bring unity and peace. The ultimate goal is to bring state-of-the-art cancer treatment in a region where even conventional radiation therapy is not so common. The current vision for the new hadrontherapy centre includes its use as a particle accelerator for research for 50% of the beam time. This is an important aspect of the project as it will help the whole scientific community to grow and develop. In addition, such a potential would attract scientists from other regions where beam time is entirely devoted to treatment and research cannot be carried out on site. The centre will become a powerful network hub to connect the South Eastern European countries with the rest of Europe.

Looking back at the ideas that the PARTNER project brought in and the path shown by visionary scientific endeavours such as SESAME, one can be confident that the centre will be a fantastic opportunity for the entire region to benefit the whole



### LARA BARAZZUOL



THEN --> NOW University Medical Center Groningen, the Netherlands

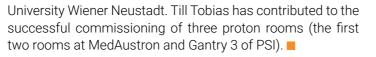
In 2016 Lara Barazzuol established her own group within the Department of Radiation Oncology at the University Medical Center Groningen (UMCG), in the Netherlands. Her research focuses on assessing the effect of radiation on the brain and aims to achieve an improved biological and molecular understanding of radiotherapy-induced neurocognitive dysfunction. Highlights from her career to date comprise several

University of Surrey

peer-reviewed articles and young investigator awards, including the 2012 Mercia Award in Medical Engineering, the 2015 British Institute of Radiology Nic McNally Award, the 2017 Adrian Begg Award and the 2017 Bas Mulder Award. Lara Barazzuol is currently project leader on grants from ZonMW and KWF (Dutch Cancer Society).



Till Tobias is a Swiss certified Medical Physicist (SGSMP), after having already the Austrian certification. He is working as Medical Physicist at PSI, specialized in TPS and workflow and lead in scripting. He is also lecturer at Applied





Neamat is currently lecturer of clinical oncology department to me! The PARTNER program has been a very important at Alexandria University, Egypt. She is also consultant of starting point in my career. This is a very unique experience brachytherapy in Future hands Hospital, Alexandria. In her and I am very grateful to ENLIGHT for that!" Today, Neamat words: "The knowledge acquired from my supervisors during shares her knowledge with her juniors' colleagues in her training in Vienna in the different areas of light ion beam home university and has started new research projects in therapy, and radiotherapy in general has been very important proton therapy.



per day in two rooms with three proton fixed beam lines (two horizontal and one vertical). We are involved in all aspects of medical physics: beam delivery, treatment planning, medical software, patient positioning and imaging systems. Currently, on top of ensure routine proton treatments, we are starting-up the commissioning phase of the first horizontal carbon ion beam line. This is a very exciting time and we are looking forward to the first carbon ion treatments in summer 2019! The PARTNER program has been an amazing starting point in our careers. The knowledge acquired by each of us in the different areas of light ion beam therapy and the We are working at the MedAustron light ion beam therapy facility, located in Wiener Neustadt, Austria and we are all PARTNER network developed have been very useful! It allows us today to implement light ion beam therapy treatments at part of the Medical Physics department. Patient treatments started at the end of 2016 with the first horizontal proton MedAustron. This is a very unique experience and we are very grateful to ENLIGHT for that! beam line. Today we are routinely treating about 25 patients





Upon returning from CNAO, Jeffrey went back to his usual proton beam therapy technical lead includes selecting the routine of managing patients in the National Cancer Centre equipment as well as developing clinical treatment protocols in Singapore, which takes care of 60% of the country's cancer for patient treatment. In his own words: "The training that I patients. The center will relocate to a new, bigger and better received with PARTNER really helped to prepare me for this cancer center in 2022, which will also include a state-of-thecurrent role. It is also through this same network that we can art Proton Beam Therapy Centre. Jeffrey's current role as forge future training and collaboration connections."



### LOIC GREVILLOT

- THEN IBA (Ion Beam Applications

### JHONNATAN OSORIO --- THEN CNAO MORENO





David is currently working as an independent consultant in treatment rooms, each with a 180-degree gantry. David has the United States. He collaborates with companies to develop proton therapy systems. One of the facilities David is current- has developed a particle physics card game whose rules ly working on is scheduled to treat a first patient before the close of 2018. It makes use of a compact proton synchrotron which can deliver protons up to 330 MeV to three patient

recently started his own consulting company in Canada and mimic the principles of Particle Physics and the Standard Model.



Palma is a senior Research Scientist at the Department of Tiganelli, another PARTNER researcher. When Palma is not Science and Technology (DiST) of University of Naples and in the lab or behind a microscope, she is in the classroom she is expecting to start a permanent position as associated teaching "Developmental Biology and Animal phylogeny" and professor in Cytology and Comparative anatomy. Her current "Cytology and Histology" to students of the Degree Course of Biology at University Parthenope in Naples. research focuses on radiobiology and particle therapy. She is currently involved in international projects with Walter



After his Phd, Daniel joined the computational bioengineer- tigate the effect of biomechanical forces on formation and ing group in the medical faculty of the University of Bern (CH) as post-doctoral research fellow. His work integrates tional models of tumor growth and tissue mechanics. Daniel approaches from mathematics, physics, and engineering to build patient-specific computational models. In June 2017, the Mathematical Oncology group of the Beckman Research Daniel joined the Glims project (http://glims.ch) as Marie Institute / City of Hope hospital in Duarte. Skłodowska-Curie Global fellow. The project aims to inves-

development of brain tumors using image-based computais currently based in Southern California (USA) and works in





Chitralekha works as production engineer at Atlas Antibodies, her own words: "I believe that the knowledge and experience the commercial part of Human Protein Atlas project. The gained during the project is going to help me in all my future company handles the production, marketing and sales of reendeavors. It has been a wonderful learning program to be search tools developed by the Swedish-based Human Protein in this multidisciplinary project where, I got the opportunity Atlas program. Chitralekha works with recombinant protein to interact with brilliant scientists of different scientific backexpression and purification with affinity chromatography. In ground."



After completing his Ph.D., Walter moved to Japan where applications also in radioprotection. About his participation he worked at the National Institute of Radiological Sciences (NIRS), in the Framework of the International Open Laboratory. For the last three years he has been working in Italy, to establish a network of friends, first of all, and colleagues, at the Trento Institute of Radiological Sciences (TIFPA). He coordinates with my TIFPA colleagues, various projects, network." among which a hibernation project that could have relevant

in PARTNER, Walter remembers his days as PARTNER researcher in Germany: "PARTNER Project was an opportunity which still today are an important radiobiology European





Daniel is currently a MSCA IF Fellow at the Universidad teaches several courses at the Faculty of Physics of the Complutense de Madrid working on ionoacoustic range University and collaborate with the particle therapy centers verification of proton beams and development of contrast under construction in Madrid. agents for PET and PG proton range verification. He also



### HITRALEKHA MOHANTY

Human Protein Atlas, Sweden

### DANIEL SÁNCHEZ PARCERISA

Universidad Complutense de Madrid, Spain







Centre for Biomedical Physics, School of Healthcare and Medical Sciences, Sunway University, Malaysia

Ming Chew is currently employed as a Lecturer at the Centre for Biomedical Physics, School of Healthcare and Medical Sciences, Sunway University, Malaysia. As a lecturer, Ming Chew teaches undergraduates, does research related work, supervises postgraduates' students, deals with funding applications and proposes new undergraduate degree courses. For research work she collaborates with public and private

hospitals, local and international nuclear agencies and local private institutes that deals with radiation equipment. Although, she does not have the opportunity to be directly be involved in particle radiation due to the country economy, she is still inspiring and motivating students to pursue particle radiation. 📕



Within the PARTNER project, Ahmad worked at the Clinical efforts to establish two new courses as Medical Radiation Bioengineering Unit of CNAO Foundation in Italy to assess Engineering course and Application of Radiation course by the performance of Patient Verification System and the taking required permissions from Ministry of Sciences, Re-Patient Positioning System. After that, he came back to his search and Technology. Ahmad also was as head of Medical hometown in Iran and became a member of Medical Radi-Radiation Division at Graduate University of Advanced Techation Division as assistant professor. At Kerman Graduate nology for almost 2 years. University of Advanced Technology (KGUT) he did serious



After the PARTNER programme, Adriano went on to design a new LEIR slow extraction system and beamlines for biomedical research. In 2014, he moved to the MedAustron therapy center in Austria, where he led the accelerator commissioning activities for proton medical therapy until 2016. Back to CERN. Adriano became the Technical Director for the TERA

Foundation. In the last two years he was involved in several studies including the mechanics and optics design of a normal conducting gantry for protontherapy and a feasibility study for the expansion plan of CNAO. Adriano thinks that the knowledge acquired during the PARTNER project was instrumental to his career.



After completing her PhD, Silvia joined the Collider-Accelerhave supported the design, fabrication and testing of several radio-frequency, superconducting crab cavity prototypes. For ator Department of Broohkaven National Laboratory (BNL, Silvia, "PARTNER was a nurturing platform that provided me New York, USA) to work on the crab cavities for the High Luminosity upgrade of LHC (HL-LHC). Since 2016 she is in with great tools to work on international projects involving tenure-track at BNL developing crab cavity systems for the multi-disciplinary, culturally diverse teams." future particle colliders HL-LHC and eRHIC. In this time, she



Alina joined the PARTNER project only for three months. treatment planning for lung tumors. In the beginning of 2019 And yet, those three months that she spent at CNAO in 2012 she will move to Essen, in order to join the team setting up the allowed her to get an excellent insight in all relevant aspects clinical programme for proton therapy treatment of thoracic of Medical Physics for particle therapy. When she came back tumors at the University Hospital, in cooperation with The to Marburg, Germany, Alina did her PhD in particle therapy West German Proton Therapy Centre Essen.





After her fellowship at CERN, Vassiliki stayed at home for medical physicist and hold a permanent position working three years enjoying her two wonderful little girls (the first mainly in clinical routine which includes treatment planning, was born during, the second at the end of her time at PARTpatient and machine quality assurance, commissioning of NER). In 2015, she started working at the university hospital new machines/software and imaging. For her: "PARTNER of the Technical University Munich (Klinikum rechts der Isar) was an amazing experience and I have met so many wonderin the department of radiooncology. Today, she is a senior ful people and friends during that time."



### Graduate University of Advanced Technology, Iran

### Broohkaven National Laboratory (BNL), New York, USA

### VASSILIKI SIA ELISE KANELLOPOULOS

### University Hospital of the Technical University Munich, Germany



Valentina is currently holding a Post-Doc position, started one year ago, at the Aragon Institute of Technology (ITAINNOVA), in Saragossa (Spain). ITAINNOVA is a non-profit public institution that directly depends on the Innovation Research and University Department of the Aragon Government, mainly focused on research and innovation for both public and

private areas. Valentina is currently focusing on data analysis and model order reduction strategies, combining software development together with tensor and matrix decomposition methods applied to different fields, such as fluid-dynamics, thermodynamics or mechanics.



search ever since the PARTNER project. Currently, her research Marta is a lecturer and researcher at Stockholm University and Karolinska Institute. After finishing the PhD within the focuses on the assessment of the tumour responsiveness to radiation and aims at identifying parameters to predict the PARTNER project, she worked four years as Post-Doc within another European network called cancerARTFORCE. Meantreatment outcome at an early stage of radiotherapy in order to contribute to the development of a practical solution for a while, she also studied to be a licensed Medical Physicist. Functional imaging has been the leading thread of her retruly optimized and adaptive treatment planning strategy.



After the PARTNER project, Maxime went on to do a PhD in bioengineering focused on particle therapy, based at the CNAO (Pavia, IT). After his PhD, Maxime left the research field to work for Varian Medical Systems in Milan (IT) where he am still working as a medical physicist for the physics European Helpdesk. More specifically, he is covering the physics cus-

tomer support for the dose calculation algorithms inquiries, related to the Varian's Treatment Planning System (Eclipse), for photons and also protons. In his words: "The experience and background I gathered during the PARTNER project has been very helpful to resolve practical and clinical issues."



ANURITA SRIVASTAVA THEN  $\cdot \cdot \rightarrow$ NOW

Anurita is currently working at Maulana Azad Medical College PARTNER project helped me realise and appreciate the in Delhi, a teaching public hospital where she trains young immense effort put in by each individual involved in estabradiation oncologists the 'art' of radiotherapy. Anurita also lishing different aspects of a 'new' technology. Earlier, I had helped establish framework for data entry and record keepnot worked with biologists or engineers or computer geeks, ing, initiated a systematic academic programme, supervised and I can only thank my PARTNER friends for enriching my the establishment of brachytherapy services apart from the knowledae." regular clinics and treatments. In her words: "Being part of



Faustin is currently in Auckland, New Zealand, enjoying life with his wife and his 3 kids. He is Director of Medical IT Advisors, a company he founded 2 years ago, and CIO of Patients First, a not-for-profit company that focuses on health information governance, risk and compliance, information security, digital transformation and emerging technologies. He also recently initiated and now chairs the first New Zealand Health Cybersecurity and Privacy industry group and won the first Maori hackatron with one of his other idea/start-up on blockchains. He is convinced that "none of this would have happened without the awesome PARTNER multi-disciplinary training and network!"



WENJING CHEN THEN  $\rightarrow$  NOW Siemens Healthcare

After the PARTNER project, Wenjing left Siemens and moved Heidelberg University Hospital to develop the Chinese marto Heidelberg where she continued to work as a PhD student ket, and provide non-medical services to Chinese patients. and with a PostDoc until the end of 2016. Meanwhile, she More recently, Wenjing has been working on an association enrolled in a part-time MBA programme to add some new (Sino-Europe Startups Association e.V.) to help new entrepreskills. Since 2017, she has been working on a startup project neurs in expanding network. for medical tourism. The company became a partner of



### Stockholm University/Karolinska Institute, Sweden

Maulana Azad Medical College, India



### DEXIN Med GmbH, Germany



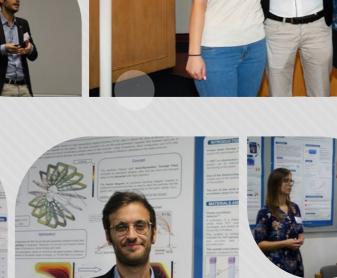


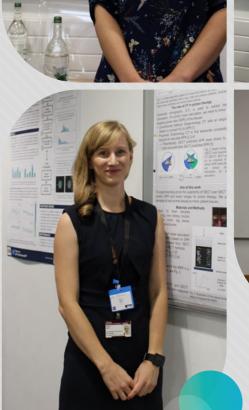


POSTER















"Like many young people in Krakow, who were involved in printing the compensator. The use of the 3D printed proton physics during their studies, I became fascinated with proton compensator, attached to the immobilization mask reduced therapy. It was inevitable - day after day, a modern cyclotron the spot size and in consequence the lateral penumbra (20center was being established in Krakow, the only such a 80%) by over 40%. center in Poland. During my studies I carried out my master's So far, two children have been treated with such compensathesis in the subject connected with proton therapy. After tors and the next compensator is in preparation. It is a great graduation, I started my PhD programme in Proton Radiofeeling to know that I have helped to provide treatment for therapy Group at the Institute of Nuclear Physics. That's how young patients with difficult cases of shallow tumors. my adventure with protons and radiotherapy began, from the beginning combined with 3D printing technology.

Cyclotron Centre Bronowice (CCB) – as a part of IFJ PAN - is the national center of hadrontherapy in Poland. This is the only place in our country where patients are treated with protons. In 2011 the first ocular patient was treated with 60 MeV protons from the self-developed AIC-144 60 MeV cyclotron. In 2015 the new facility with the IBA proton cyclotron (70-230MeV) and two gantries with Pencil Scanning Beam (PBS) technology started the operation.

The irradiation of superficial lesions, especially in children, requires application of a range shifter (RS) to deliver more shallow spots. RS is situated at a certain distance from the patient's body. However, the scattered beam is not conformal enough. Despite the potential advantages of using protons, such treatment plans could not be applied.

Then, being under time pressure, we started to think about how to better compensate the energy of protons. Maybe the range shifter could be placed directly on the patient's body or even exactly matched to it? Such a solution would significantly reduce the beam spread after passing through the absorber and would improve the dose distribution. As a result of this discussion, a new way of treatment planning for such cases was started – application of patient-specific 3D printed compensators for proton PBS of shallow situated tumors.

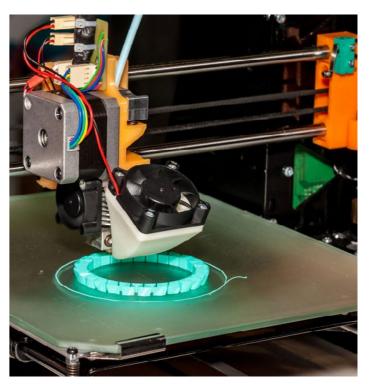
At the ENLIGHT 2017 meeting I presented very early results of the first study – an established procedure for designing and

### AGNIESZKA WOCHNIK

PhD student at the Institute of Nuclear Physics Polish Academy of Science (IFJ PAN), Krakow, Poland

PATIENT-SPECIFIC 3D PRINTED COMPENSATORS FOR PROTON PENCIL BEAM SCANNING OF SHALLOWLY SITUATED TUMOURS

We intend to conduct further research in this direction in order to better understand the properties of the materials used for printing. This will be the next stage of my adventure with protons and 3D printing technology."



Patient-specific 3D printed compensators for proton pencil beam scanning of shallow situated tumours



"I discovered superconductivity during my studies at the the cryogenics and the mechanics. Better conditions for University of Bologna. I literally fell in love with this amazing superconductors means higher magnetic field and, therefore, physics phenomenon and its applications. After my master's strong reduction of size and weight. GaToroid can claim a thesis at CERN, supervised by the magnet group leader very compact footprint, with a diameter of 3 metres for Luca Bottura, I had the chance to join Prof. Amaldi and his protons and 5 for carbon ions: less than half of the existing research group at TERA, working on hadron therapy: I fell in gantries! love for the second time. Could you imagine what happened Furthermore, GaToroid can accept incoming particles in a when Luca proposed me to design and develop his new conbroad range of energies, covering the complete treatment of cept of superconducting Toroidal Gantry? The possibility of proton and carbons ions. The beam can be delivered at very combining my passion and experience in superconductivity, high speed, without mechanical or electrical limitations of the with my interest and knowledge in medical applications? A gantry. dream come true!

Large acceptance, fixed configuration and use of supercon-Nowadays, the concept of proton and hadron therapy ganductors offer interesting reduction in size, weight and cost tries is based on magnetic transfer lines rotating around of gantries and related infrastructures, creating an attractive the patient. The established technology results in bulky and alternative to the state-of-the-art. The GaToroid concept massive structures, especially for heavy ions. Wonderful opens the door to different approaches to therapy, creating examples of engineering are the gantries of HIT, Germany, new opportunities for imaging, monitoring and treatment. and Chiba, Japan. The latter uses superconducting magnets to increase the achievable magnetic field and reduce the size After one year of PhD work, based on intense study and hard of the machine. However, the rotating nature of the gantry work, the dream of GaToroid is now more solid, becoming introduces significant complexity in terms of cryogenics and day by day closer to reality. What I am learning during these mechanical stability. years has inestimable value: I am very thankful to Luca for this unique opportunity of technical and human growth." GaToroid is a new concept of gantry, based on fixed toroidal

configuration, able to deliver the dose at discrete number of <sup>1</sup> L. Bottura, "A Gantry and apparatus for focussing beams of angles with neither rotation of the magnets nor the patient<sup>1</sup> charged particles," Patent, EP 18173426.0., May 2018 The basic principle is to use axial-symmetric magnetic field between each pair of coils constituting the torus to bend and focus accelerated particles down to the isocentre. The patient is placed inside the torus bore, similarly to an MRI configuration, but in null magnetic field. A single upstream bending magnet, rotating or combined vertical/horizontal, is used to steer the particles into the gantry with a proper angle, depending on the beam rigidity.

GaToroid works in complete steady state configuration: no rotation of mechanical parts and no variations of magnetic field. This feature allows one to push the potential of superconductivity to its limits, simplifying, at the same time,

## **ESTHER BÄR**

PhD student at University College London UCL, Department of Medical Physics and Bioengineering

**DUAL-ENERGY CT TO ESTIMATE PROTON STOPPING POWERS: A VALIDATION USING REAL TISSUES** 

"Proton therapy caught my interest during my training as a clinical scientist in Heidelberg, Germany. Since I first became aware of it, I was convinced that proton therapy is the future of radiotherapy. Although I strongly believed in it, I knew that there was room for improvement in order to make treatments better for the patients. While there are many ways to improve proton therapy, I chose to work on the imaging side.

Radiotherapy treatment planning, in general, requires a certain knowledge of the tissues that are in the beam's path. For radiotherapy with X-rays, the electronic densities of tissues are required for dose calculation. For proton therapy, however, the required tissue quantity is the stopping power (SP) of the material the beam is traversing.

In current clinical practice, the SPs of human tissues are estimated from a single-energy CT (SECT) scan of the patient. The CT numbers are converted into an estimate of the SP by the use of a lookup table. This procedure, although robust, cannot consider inter- and intra-subject tissue variations, because there is no one-to-one relation between the CT numbers and the SPs.

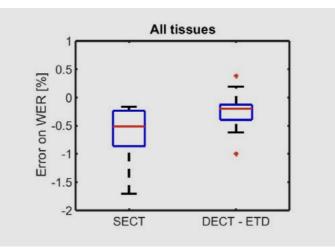
To improve SP estimation, dual-energy CT (DECT) was suggested. With DECT, two CT images of the patient are taken simultaneously at two different energies. In this manner, more information about the tissues is available and it can be used to estimate the SPs of tissues. Several groups have suggested mathematical models to estimate SPs from dual-energy CT and proven their mathematical consistency as well as their theoretical superiority over SECT.

Before DECT-estimated SPs can replace SECT-estimated SPs for proton treatment planning in a clinical environment, the accuracy of DECT predicted SPs needed to be demonstrated. Also this proof needed to be performed in a realistic clinical situation with tissues instead of phantoms, precise measurement techniques and accurate beam simulation.

Together with researchers from the University College London WFR tissues

(UCL), the University of Montreal and the Massachusetts General Hospital, we decided to take on this challenge. Under the supervision of Hugo Bouchard, Hsiao-Ming Lu, and Gary Royle, we developed an efficient measurement technique to measure the water equivalent range (WER) of a proton beam passing through animal tissue samples. We used twelve different soft tissue and bone samples and compared measured WER values to SECT- and DECT-predicted WER values. With this setup, we were able to show that DECT is superior to SECT in predicting SPs and beam ranges. In soft tissues, the errors on the predicted beam range were as small as 0.2% (as opposed to 0.5% with SECT), and in highly inhomogeneous bones we achieved an accuracy of 1.1% (1.4% with SECT). This strongly indicates that we should use DECT instead of SECT for proton therapy planning.

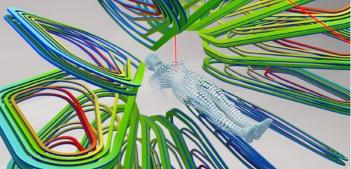
This project presents a necessary step towards implementing DECT for proton therapy planning and thus improving the accuracy of proton therapy. A continuous effort will be necessary in the future to improve imaging techniques for therapy planning and adaption. It's an exciting and inspiring field, with many challenges still to tackle and new imaging technologies emerging."



## **ENRICO FELCINI**

**CERN PhD student EPFL** Ecole Polytechnique Fédérale de Lausanne

### MAGNETIC DESIGN OF NOVEL **TOROIDAL GANTRY FOR** HADRONTHERAPY



Novel gantry

## **MULTIDISCIPLINARY APPROACH: NOW IT WORKS,** BUT...



New clinical strategies often require accepting different ideas and approaches. This is not always easy and it might not be immediately successful but Roberto Orecchia, Scientific Director of CNAO and IEO, has never been afraid of introducing novel solutions into the clinic. "Any new drugs. tools or techniques need to demonstrate their positive effect on the patient," he clarifies. "This can be in terms of purely medical effects or even global sustainability. We have to remain open to new proposals but we also need to stay constructively critical in order to preserve the patient's right to have the best cure available for the specific disease."

This is true for common drugs such as antibiotics or painkillers, but also for huge paradigm changers such as hadrontherapy and high-tech imaging. The field has been developing fast in recent years but when ENLIGHT was established, the success of this approach was not a given. "On one side, we had physicists who were literally "in love" with the pure technology and could not see the big challenges it represented for the medical community and the reality of hospitals distributed world-wide. And, on the other side, we had the clinicians, biologists and life science experts who were sceptical towards new techniques that had not come out of the medical sector," says Orecchia.

Medical doctors are used to having a very pragmatic approach to new technologies, they measure feasibility in terms of actual outcome on patients. Fundamental research works on the assumption that physics laws govern the Universe and will, sooner or later, bring progress and development to Roberto Orecchia, Scientific Director of the CNAO Foundation (Italian society. Both approaches are valid and work for the good of National Centre for Hadrontherapy) Pavia, Italy people but they deal with different timescales, procedures and constraints. "In my career I have faced many times questions from colleagues who were not convinced that we a positive outcome," explains Orecchia. "However, it took us should use a new technique or a novel technology," confirms several decades before the technique could really be accept-Orecchia. "However, medicine is science and scientists must ed by the medical doctors and eventually brought to a large stay curious in whatever field they are. Curiosity is not always number of patients. And this happened effectively only after a given in the medical field but I was lucky enough to be able the two communities - physicists on one side and clinicians to create the "critical mass" needed to nurture new ideas on the other – had been brought together and had talked to and let them grow. Conservative approaches are not wrong each other. ENLIGHT was instrumental to this strategy and as long as they ensure high standards to the patients. But we should be very proud of it." innovation is only possible if we try to go beyond what we feel Because we now have learned many things from the past and the two communities have established solid links, we can

comfortable with because we have been doing it for many vears." design the future in a better way. "Medical doctors should be Having an open-minded approach doesn't mean forgetting included "in the loop" at a much earlier stage by scientists about the fundamental steps that medical doctors have to who are developing new technologies for the medical field," go through before they can possibly accept to change estabconfirms Orecchia. "For example, the dream machine of every lished treatments. "We need to make sure that the scientific radio-oncologist of today would be a multi-ion device. With rationale is strong, that we perform an accurate technology such a device, we would ideally be able to select the most assessment and that we evaluate and validate the new techsuitable type of beam for the specific tumour we have to treat nique by performing comparative clinical studies," points out and we would be able to tune the energy, set the most precise Orecchia. "This is what we call evidence-based medicine and angle and choose the right dose to depose in the cancerous it's the irreplaceable guideline for all doctors." area."

When creating something totally innovative, the key to suc-We don't know whether such a machine will ever be built but, cess is an inclusive strategy. The example of hadrontherapy also thanks to multidisciplinary platforms such as ENLIGHT, is a very good one. "The use of protons in oncology was we are now sure that we are well equipped to take on the introduced by physicists who were doing their tests in their common challenge and continue working together. laboratories because they were sure that the fundamental gain introduced by the Bragg peak would have resulted in Antonella Del Rosso, CERN



## **NEW IDEAS FOR THERAPY** WITH IONS



A virtual particle therapy centre

A major new field in all of cancer therapy is immune therapy. Particle therapy with ion beams is attracting increasing interest and new facilities for treatment and research are being An important characteristic of tumour cells is their ability to hide from the immune system - but with recent advances in planned. However, the cost, size, and complexity of these molecular biology, targeted drugs can re-activate the immune facilities are indeed hampering the widespread adoption of system and cause it to attack the tumour cells throughout the this treatment modality. The few centres operating with ions body. Ideally, this will defeat even smallest metastases, offerare collecting a wealth of invaluable information to advance ing the potential of a true cure at least for a certain subset of the field, but they are based on technology developed in the cancer entities. A combination with radiotherapy can in principast century. Can we consider making the accelerator smaller and possibly less expensive by adopting new accelerator techple enhance this immune stimulation, as the cells damaged by irradiation release DNA fragments, which can in turn lead to a nologies? Can the wider ion therapy community - accelerator targeted immune response. As the action especially of heavier and medical experts - agree on a set of parameters and basic ions on the DNA is fundamentally different to photons, more requirement for a new generation of ion facilities? What is the and more complex DNA fragments are produced, providing experience from the present ion therapy centres and how can potentially a better immune stimulation. First hints on this this be translated into the requirements for new facilities? To increased synergy between immune stimulation and densely find answers to these questions sixty experts from all over the ionizing irradiation were already discovered, but this specific inworld met from 19-21 June 2018 at the European Scientific teraction and clinical implementation will be a major research Institute (ESI) in Archamps, France, for a Workshop on "Ideas field of the coming years. Research facilities offering flexible and technologies for a next generation facility for medical conditions for radiobiological research of immune-stimulating research and therapy with ions". agents in animal models are needed. Ideally, these facilities will The workshop, jointly organised by CERN, GSI and ESI, focused provide different ion species such as protons, Helium, Carbon on possibilities to advance towards the design for a next genand even heavier ions to study interactions with a wide variaeration medical and research facility for ion therapy in Europe, tion in energy density.

devoted to both research with different ions and patient treat-

There are clinical demands for faster, more efficient and flexment. ible therapy that could also open up the possibility for routine This workshop allowed experts of different related disciplines treatment of the 'big killers' such as lung cancer. This includes availability of technically still challenging gantry systems for to share current experiences, identify potential directions for ions. Technical options for increased irradiation speed such as improvement, discuss synergies between medical accelerdynamic intensity control or the extraction of multiple energies ators and other accelerator projects, and exchange ideas for future facilities. Overall, the general requests for increased from a single synchrotron cycle were already realized at HIT in Heidelberg and HIMAC, Chiba, Japan. The Japanese facility clinical efficacy and cost-effectiveness could be addressed in particular demonstrates a highly efficient workflow for the with further research on radiobiology, accelerator and medical therapy of moving organs, combining very fast irradiations physics, with the needs of the patient being at the focal point. with effective rescanning and marker-less gating for motion The finite range of particle therapy offers superior conformity control.

as opposed to conventional photon therapy. Tailored dose es-While for protons and Carbon ions by far the most clinical calation and also the specific advantages of heavier particles experience exists, other ion species could be of interest. In the are important to overcome radioresistant or hypoxic tumours, past, Neon ions were used at Berkeley, but were apparently too as has already been shown in a number of albeit rare headheavy, with projectile fragmentation leading to a worse ratio and-neck cancers.



Carbon Beam Gantry at HIT

of the dose in the entrance channel to the Bragg peak region. Oxygen on the other hand could offer a higher energy density in the Spread-Out Bragg Peak to combat hypoxia with acceptable entrance dose. The most interest currently focuses on Helium as the 'heavier proton'. Helium has a significantly minimised scattering compared to protons, far less fragmentation and less complex radiobiology than Carbon. It could thus possibly provide an ideal combination of small accelerators but still highly precise irradiation with especially a better lateral dose fall-off. Both species are available at the HIT facility for research purposes.

Large deficits in imaging exist especially in contrast to stateof-the-art photon therapy, where 3D imaging is seamlessly integrated in virtually every treatment. New options for image-guided therapy must be accommodated in the overall facility design but also by the accelerator. While in-room cone beam CT and CT-on-rails becomes more and more available in particle facilities, it is still not the standard as in photon therapy. This is true in spite of the fact that the precise irradiation with pencil beam scanning requires also more precision in patient setup and especially control of the beam range. Beam range estimation and monitoring of the beam itself in the patient is a major research topic. In-beam PET imaging of projectile and target fragments has been in use for several years, pioneered during the GSI pilot project. Faster imaging with at least the potential for adaptive therapy is offered by prompt emissions from nuclear reactions. First prototypes of a prompt gamma slit camera for range monitoring are in a clinical test phase, but also research on prompt particles is ongoing. How to incorporate such data into the treatment workflow is a further topic of medical physics research. The pinnacle of this development could be fast, adaptive particle therapy combined with online MR imaging and monitoring of prompt emissions to assess the beam range. This would finally enable a new realm of precisely eliminating tumours also in complex anatomical locations or moving organs.

As participants remarked, one could feel enthusiasm in the room, and could project on the enthusiasm at GSI where ion therapy has been pioneered since the early '90s. Leading figures of this initiative commented on their experience and recalled the day that the first patient was admitted for treatment - when, as an extra IT security measure, GSI switched

off Internet connection across the site in order to prevent any external disturbance of the patient treatment.

The available designs of very compact single-room installations for proton facilities was presented as a good example for future developments for ion beams. These facilities were designed to be installed into existing buildings, fitting for example into two conventional photon therapy bunkers.

Two approaches for a next generation accelerator were discussed. An option relying on proven delivery schemes could be based on a super-conducting synchrotron. First design studies have already been conducted in Europe and in Japan, where experience from the design of a super-conducting gantry clearly indicated the possible reduction in size. Another option could be to explore novel approaches such a high repetition frequency Linac-based design. Also, for this second option, theoretical studies and proposals exist, showing the potential of this solution. Especially intriguing could be the very fast change in beam energy in a Linac-design, which could even enable compensation of motion-induced range changes. Both avenues offer solutions, with potentially superior, but also more uncertain, results expected from revolutionary approaches.

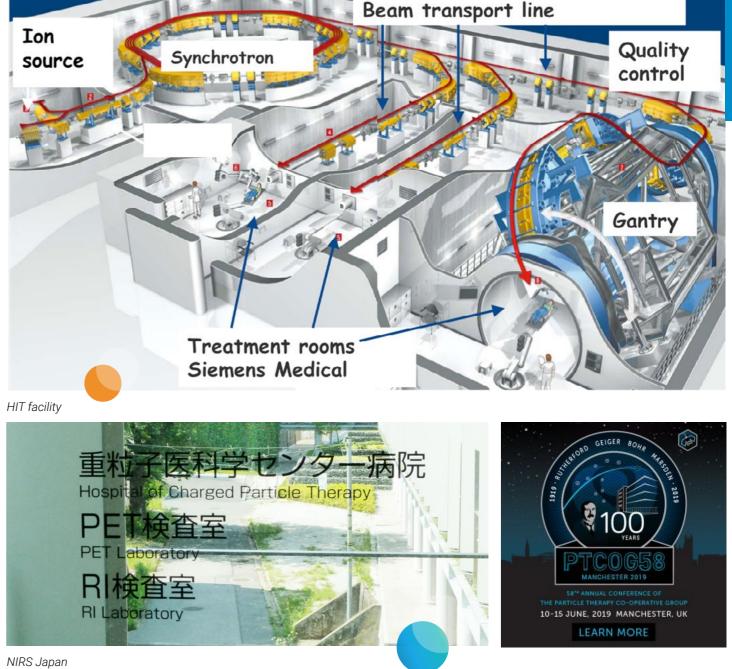
The closing discussion session clearly showed that the workshop addressed important issues on ion therapy in a timely manner. In particular, the community sees the need for and is eager to proceed towards establishing a dedicated collaboration resulting in a proposal to the European Commission. Discussing the outline of a possible R&D programme brought up the benefits of a study comparing the linac- or synchrotron-based designs.

To further pave the way towards this goal, a dedicated meeting is proposed in 2019 with the aim to define a common set of parameters and to receive feedback from different potential contributors. The success story of particle therapy in general and also ion beams in particular with more than 20,000 patients treated will continue with further innovations driving the field. A continuing expansion of ion beam facilities in the world will depend - not exclusively, but to a high degree - on our ability to make it also a commercial success, with smaller, less expensive and patient-oriented facilities designed to tackle the challenges of future cancer therapy.

> Maurizio Vretenar<sup>1</sup>, Yota Foka<sup>2</sup>, Christian Graeff<sup>3</sup> <sup>1</sup> CERN, <sup>2</sup> GSI, <sup>3</sup> GSI



CNAO treatment room



## **AGENDA**

NAME OF THE EVENT	DATE	PLACE OF THE EVENT	S S
26 - 30 April 2019	ESTRO 38 Annual Meeting	Milan, Italy	JRE EVENTS
10 - 15 June 2019	PTCOG 58 Annual Meeting	Manchester, UK	FUTURE
01 - 03 July 2019	ENLIGHT 2019	Caen, France	
15 - 18 September 2019	ASTRO's 61st Annual Meeting	Chicago, IL, USA	
27 October - 03 November 2019	2019 IEEE Nuclear Science Symposium and Medical Imaging Conference	Manchester, UK	

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# AUSTRALIAN NATIONAL PARTICLE THERAPO SYNDOSIUM

Westmead Hospital in Sydney was the venue of the Australia's second national particle therapy symposium. The hospital is currently undergoing major transformation works.

The second Australian National Particle therapy Symposium was held in Sydney in early July. Delegates represented consumers, government agencies, regulatory and training bodies, health services, research organisations (including Trans Tasman Radiation Oncology Group Cancer Research) and universities across Australia, New Zealand and overseas. The Symposium aimed to make the clinical case for carbon ions as well as protons, and to explore the non-clinical as well as clinical research opportunities that a particle therapy facility including one with carbon ions, could bring Australia. National and international collaboration amongst the clinical and research communities was a strong theme throughout the Symposium.

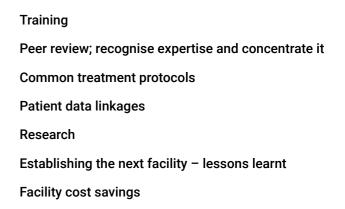
The first proton therapy facility in Australia will be the Bragg Centre for Proton Therapy in Adelaide, expected to treat its first patients in 2022. Consortia of universities, Australia's Nuclear Science and Technology Organisation and academic teaching hospitals in Sydney, Brisbane and Melbourne have well developed business cases for particle therapy facilities, with the Westmead precinct in western Sydney aiming to include carbon ions as well as protons. There has been no national tender for a particle therapy facility in Australia and initiatives are being driven primarily by clinicians.

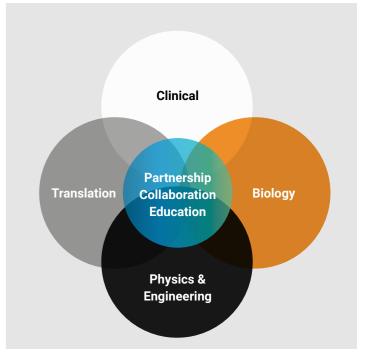
Evidence of benefit for particle therapy is accumulating and as slow adopters, Australian patients are missing out on this treatment. Professor Peter Choong (Chair of the Bone and Soft Tissue Sarcoma Service at the Peter MacCallum Cancer Centre in Melbourne) presented the reasons their service now refers some patients to Japan for carbon ion therapy. Professor Shannon MacDonald from Massachusetts General Cancer Center and Harvard Medical School and Dr Piero Fossati from MedAustron presented updates on the indications for treatment by proton therapy and selecting patients for treatment by carbon ion therapy, and well as the clinical research potential for these therapies. Dr Peter Urschutz provided an overview of how the MedAustron facility developed – from concept to clinical operation.

CERN was at the foundation of the Symposium. Professor Emmanuel Tsmeslis, CERN's Head of Relations with Associate Members and Non-Member States, explained CERN's role as a gateway to science, technology and innovation, and its mission to push back the frontiers of knowledge, develop new technologies for accelerators and detectors, train scientists and engineers of tomorrow and unite people from different cultures and countries. Professor Tsmeslis demonstrated Australia's existing links to CERN through the involvement of scientists from the Universities of Melbourne, Sydney and Adelaide in the ATLAS experiment. Professor Manjit Dosanjh, CERN's Medical Applications Senior Advisor and ENLIGHT Coordinator, illustrated CERN's involvement in medical applications through accelerating particle beams, detecting particles and large-scale computing. She also explained how ENLIGHT has leveraged physics collaboration into a multidisciplinary medical environment to identify challenges, share knowledge and best practice, harmonise data, provide training and education, innovate to improve and lobby for funding. Professor Dosanjh advised there is still need for radiobiology studies due to the paucity of long-term clinical data and gave practical advice on the requirements for a radiobiology facility as well as a wish-list for accelerator requirements. She informed the Symposium participants about the six work packages of the recently formed European Particle Therapy Network (EPTN). The mission statement of EPTN "to promote clinical and research collaboration between the rapidly increasing number of European particle therapy centres and toensure that particle therapy becomes integrated in the overall radiation oncology community" represents the vision for particle therapy in Australia.

Verity Ahern, Western Sydney Local Health District, Australia

### **COLLABORATION**





The vision





Group photo of the participants

## PROTON IMAGING WORKSHOP LYON 2018

On June 14-15 this year, a workshop focused on ion imaging took place in Lyon, France. It was attended by about 50 researchers from around Europe, the USA, and Southeast Asia. Attendance was free of charge thanks to funding through the EU's Marie-Curie fellowship program as well as the LabEx Primes network financed by the ANR. The participants ranged from medical physics experts from major proton therapy facilities, engineers and particle physicists with experience in detector design, to mathematicians specialized in inverse problems and image reconstruction. The event attracted experienced researchers as well as PhD students sharing their recent work. It was organized by Nils Krah and Simon Rit, both from the Creatis lab in Lyon, France.

The workshop setting was overall rather informal, with room for discussion, thus creating a very constructive atmosphere and facilitating a rich exchange of ideas. The first day was rounded up by a dinner in the beautiful center of Lyon with the interesting twist that the restaurant was actually part of a reputable gastronomy school. With the waiters bustling around everyone, it felt as lively as in some experimental hall at CERN.

A first session, dedicated to proton imaging setups, demonstrated that hardware is already quite advanced, with proton CT scanner prototypes being assembled in a clinical environment in the USA and a new generation prototype under development in the United Kingdom. An interesting alternative approach from Norway made use of hardware which was formerly used for a detector in a high energy physics experiment at CERN.

Several speakers presented work focused on data processing and tomographic image reconstruction. While the basic methodology is

already established, several innovative approaches were proposed. To reconstruct more detailed information about the properties of the patient's tissue than is currently the case or to optimize the irradiation procedure in order to lower the dose to the patient while keeping the image quality at a sufficient level, being only two examples.

One session was dedicated to imaging with heavier ions, such as helium, carbon or oxygen, that are available in a few facilities around the world. When compared to protons, these particles have a more complex interaction with the human tissue and this brings along the need for dedicated post-processing methods. Recent developments in this direction were shared. It is noteworthy that some of the presented experimental work made use of the Medipix/Timepix detector which was originally developed at CERN.

A final session focused on efforts to realize proton imaging at comparatively low cost. Such systems make use of detector hardware that is often already available in proton therapy facilities and researchers showed that two-dimensional proton radiographic images can be obtained which could complement the currently available arsenal of X-ray imagery.

Motivated by the success of this year's workshop, a second edition is in preparation for 2019, in Manchester, UK. The event will take place during the educational session of the PTCOG conference and is therefore expected to attract the attention of even more experts in the field of hadrontherapy. More details will be announced through the newly created website ionimaging.org and interested people are invited to sign up for the ion-imaging mailing list (instructions can be found on the website).

Nils Krah, Simon Rit, Creatis lab - CNRS, Lyon, France



#ESTRO38 WWW.ESTRO.ORG

### 26-30 April 2019 Milan, Italy

**ESTRO** 

38

Targeting optimal care, together



# **ENLIGHT** Advisory Committee

**Bleddyn Jones** Radiation Oncologist, Oxford

Manjit Dosanjh Biologist, CERN ENLIGHT Coordinator, Geneva



Jacques Balosso Radiation Oncologist, Centre François Baclesse, Caen and CHU, Grenoble



**Marco Durante** Physicist and Radiobiologist, Trento





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**Pawel Olko** Physicist, Krakow



**Richard Poetter** Radiation Oncologist, Vienna



**Piero Fossati** Radiation Oncologist, MedAustron, Wiener Neustadt

### THE EUROPEAN NETWORK FOR LIGHT ION HADRON THERAPY

### A multidisciplinary platform aimed at a coordinated effort towards ion beam research in Europe.

The European Network for Light Ion Hadron Therapy (ENLIGHT), which had its inaugural meeting at the European Organization for Nuclear Research (CERN) in February 2002, today has more than 600 participants from nearly 25 European countries. Harnessing the full potential of particle therapy requires the expertise and ability of physicists, physicians, radiobiologists, engineers, and information technology experts, as well as collaboration between academic, research, and industrial partners.

The ENLIGHT network has been instrumental in bringing together different European centres to promote hadron therapy and to help establish international discussions comparing the respective advantages of intensity modulated radiation proton and carbon therapies. A major success of ENLIGHT has been the creation of a multidisciplinary platform bringing together communities that were traditionally separated, so that clinicians, physicists, biologists, and engineers work side-by-side. Special attention is also given to the training of young researchers and professionals of oncologic radiotherapy.

For more information and contact details please visit the ENLIGHT website at cern.ch/enlight (or scan the QR code)

Join the ENLIGHT network Register to become a member here.

https://indico.cern.ch/e/RegisterENLIGHT



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