

Scientific Report

EOC-USI 2022



In partnership with:



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List of frequently used abbreviations

AFRi	Area Formazione medica e Ricerca / Medical Education and Research Division	PI	Principal Investigator
CTU -EOC	Clinical Trial Unit EOC	RCT	Randomized controlled trial
EOC	Ente Ospedaliero Cantonale	SAKK	Swiss Group for Clinical Cancer research
ICCT	Istituto Cardiocentro Ticino / Cardiocentro Ticino Institute	SCTO	Swiss Clinical Trial Organization
ICP	Istituto Cantonale di Patologia / Cantonal Institute of Pathology	SNSF	Swiss National Science Foundation
ICT	Area Informatica e tecnologia della Comunicazione / Information and Communications Technology Division	SUPSI	Scuola Universitaria Professionale della Svizzera italiana
IIMSI	Istituto Imaging della Svizzera Italiana / Imaging Institute of Southern Switzerland	USI	Università della Svizzera italiana
INSI	Istituto di Neuroscienze Cliniche della Svizzera Italiana / Institute of Clinical Neurosciences of Southern Switzerland		
IOR	Institute of Oncology Research		
IOSI	Istituto Oncologico della Svizzera Italiana / Oncology Institute of Southern Switzerland		
IPSI	Istituto Pediatrico della Svizzera Italiana / Institute of Pediatrics of Southern Switzerland		
IRB	Istituto di Ricerca in Biomedicina / Institute for Research in Biomedicine		
LRT	Laboratori di Ricerca Traslazionale / Laboratories for Translational Research		
LRU	Local Research Unit		

Preface

This annual scientific report summarizes the research activities at the Institutes and Departments of the Ente Ospedaliero Cantonale (EOC) and at the Università della Svizzera italiana (USI) in the field of medical research for the year 2022. This format reflects the increasing integration between research conducted at the two institutions following the creation of the USI Faculty of Biomedical Sciences and the signature of a research cooperation agreement in 2020. Overall, the EOC scientific production in 2022 confirmed the upward trend recorded the previous year. In 2022 the EOC Institutes and Departments, including both the clinical research groups and the laboratories for translational research (LRT-EOC), produced high-quality research attaining broad recognition at national and international level. Furthermore, several research projects are underway involving multidisciplinary teams and collaborations with research groups worldwide. Collaborative efforts will further fuel the EOC research activities in the upcoming years.

Together with patient care and education, research is an essential part of EOC activities. Bringing research close to practice facilitates the development of clinical studies and the monitoring of the effectiveness of clinical interventions and care processes, guarantees the quality and safety of services provided to patients and ensures that patients benefit from innovative diagnostic methods and therapies.

The growth in quantity and quality of research activities at EOC is paramount for the USI Faculty of Biomedical Sciences and it is in line with the 2022-2025 EOC strategic plan for a cantonal hospital with mandate for academic education and research that aspires to become a university hospital.



EOC Institutes and multisites Departments



Oncology Institute of Southern Switzerland (IOSI)

Prof. Silke Gillessen Sommer, MD

Medical and Scientific Director

Prof. Anastasios Stathis, MD

Chair Clinical Research

Luigi Tortola, PhD

Scientific Project Manager

The Oncology Institute of Southern Switzerland (IOSI) is one of the largest clinical cancer research centers in Switzerland with a longstanding experience in the conduct of clinical trials. The IOSI clinics of Medical Oncology (Head: Prof. Dr. S. Gillessen), Haematology (Head: Prof. Dr. G. Stüssi), Radiation Oncology (Head: Prof. Dr. T. Zilli) and Palliative Care (Head: PD Dr. C. Gamondi) have their own clinical research activities and each team, based on its scientific interests, can propose trials to be activated or promote investigator-initiated trials. The IOSI scientific research board (composed of senior physicians, the head of research nurses, the CRU manager, the scientific project manager and a finance representative) decides about the trials to be opened and is responsible for the strategy of the clinical research of the institute. To support investigators at IOSI and promote clinical research, a well-structured clinical research unit (CRU) is active within the IOSI and coordinates the research activities. The IOSI-CRU staff includes research nurses, study coordinators and a pharmacist and provides the infrastructures and support for researchers of the institute. Finally, the New Drugs Development Unit (Head: Prof. Dr. A. Stathis) active in the Research Day Hospital at the San Giovanni Hospital takes care of the activation and conduct of early phase clinical trials including first-in-human phase I clinical trials.

The three main areas of clinical research of our institute comprise new drugs development, prostate cancer and lymphoproliferative diseases (lymphoma and chronic lymphocytic leukaemia), but we have also started to strengthen our clinical research in additional specialties such as lung, breast and gastrointestinal cancer.

During 2022, 26 new clinical studies (among them 5 phase I and 11 phase II-III interventional clinical trials) were activated, which added to the already active, resulted in 65 open studies at the end of 2022, 48 of them being interventional phase I-III clinical trials.

Together with studies evaluating new drugs mainly sponsored by pharmaceutical industry, we also run several investigator-initiated trials assessing biomarkers in haematological, prostate, breast and gynecological malignancies in collaboration with the Institute of Oncology Research (IOR).

The IOSI has an excellent history of collaboration with both the pharmaceutical industry and non-for-profit academic organizations. Of particular relevance are the collaborations with the IOR, the Institute for Research in Biomedicine (IRB), the International Extranodal Lymphoma Group (IELSG), Swiss Group for Clinical Cancer Research (SAKK), European Organisation for



Research and Treatment of Cancer (EORTC) and the Clinical Trial Unit of the Ente Ospedaliero Cantonale (CTU-EOC). Many of our medical staff have a central role in the USI Faculty of Biomedical Sciences and are also associated with ETH Zurich, University of Berne, University of Geneva, University of Zurich, University of Padua and University of Manchester.

The high volume of clinical research activity highlights the devotion of IOSI to clinical research, with an extensive publication record of IOSI researchers in a multitude of areas related to both solid tumours and hematological malignancies that was maintained also in 2022, with numerous high impact factor publications. Researchers from the IOSI present regularly results of their research or are invited as speakers in major international conferences.

In 2022, some important awards were obtained from physicians affiliated with the IOSI. The Swiss Society of Hematology and the Miescher Foundation for Research in Hematology assigned their 2022 award to Dr. A. Condoluci, while Dr. R. Pereira Mestre was awarded with the SAKK/AbbVie Award. In addition, 5 EOC awards were assigned in 2022, two young investigator awards (Dr. A. Condolucci and Dr. M.C. Piroso) and three senior awards (PD Dr. S. De Dosso, Prof.

Dr. D. Rossi and Prof. Dr. A. Stathis). The training and career development of younger colleagues that will be the future principal investigators is an essential aspect as well as the training of international fellows through our international fellowship program with 2-3 fellows training each year in clinical research at the IOSI.

Finally, several grants were obtained by IOSI investigators, in particular, PD Dr. C. Gamondi from the Horizon program for the project PAINLESS, PD Dr. U. Vogl from the foundation Fond'Action Contre Le Cancer for the clinical trial ACTIDIET-PRO, and Dr. R. Pereira Mestre from Fondazione Daccò for the project SUNSHINE BIO. For the years to come, the development of investigator-initiated trials represents a major objective of clinical research for the IOSI. For this reason, the IOSI is aiming to increase the number of investigator-initiated clinical trials over the next years in collaboration also with the CTU-EOC. With the aim of further promoting the transfer of preclinical knowledge to the clinic, regular scientific meetings with the heads and PIs of IOR and the IRB (now Bios+) have continued in 2022. This resulted in the publication of several collaborative original research articles involving IOSI as well as IRB / IOR.

Institute of Clinical Neurosciences of Southern Switzerland – Neurocenter (INSI)

Prof. Alain Kaelin, MD PhD
Medical and Scientific Director

The mission of the INSI is offering the highest quality of cure to the patient with the application and promotion of an inter- and multidisciplinary approach. Translational and clinical research as well as educational activities are a priority of the INSI with a close interaction between research activities and patient care. Academic collaboration with the USI, SUPSI as well as other national and international universities are contributing to the success of the INSI. Also in 2022 there was an intense and successful clinical and translational research activity, with a large number of scientific articles published in high-ranked journals and numerous competitive grants obtained.

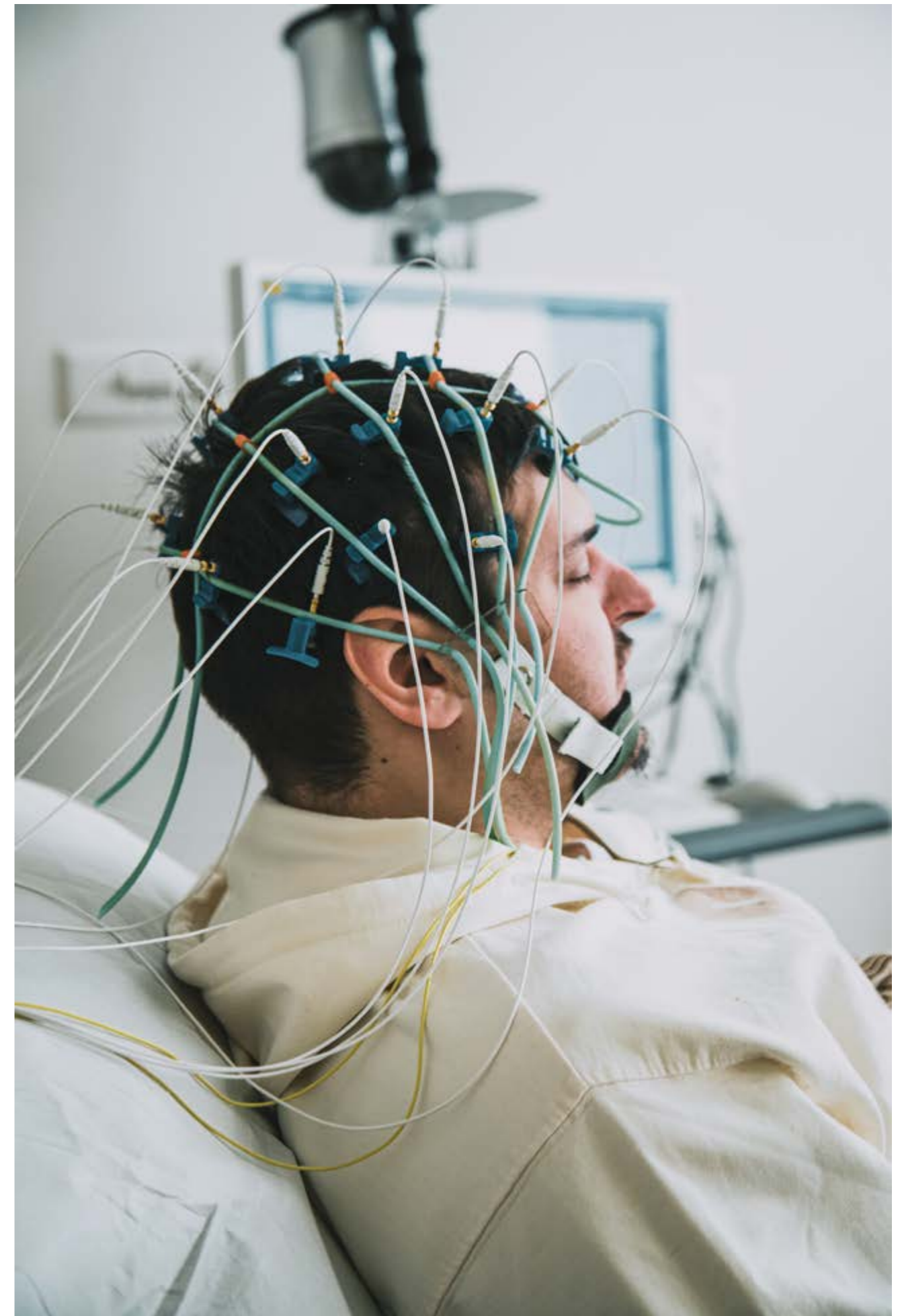
Neurodegeneration research group (Prof. Dr. P. Paganetti)

The team searches for proteins involved in aberrant cellular mechanisms associated to aging-related human disorders, neurodegeneration and cancer, as well as other rare proteinopathies. Currently we are mainly investigating Tau as model protein to address two unanswered questions: the putative role of Tau in the management of a lesion to the DNA or in the epigenetic regulation of DNA transcription and the identification of the cellular organelle where toxic forms of Tau propagate during disease progression. Our recent data clarify, among other, the molecular mechanisms utilized by Tau to regulate the stability and

activity of p53, as well as the epigenetic repression of gene expression and cell senescence induction. In cells exposed to pathogenic forms of Tau, e.g. when included in extracellular vesicles, we observed for the first time the molecular encounter of internalized toxic Tau with endogenous normal Tau in lysosomes. We are also interested to study the role of lysosomal dysfunction as it occurs in rare pediatric diseases or in the aging brain. As a Tau-independent project, we are searching for natural compounds able to increase the expression of breast cancer-associated proteins with the goal to de-risk cancer in families carrying BRCA1/2 mutations. [See also chapter 3 on the Laboratories for Translational Research.](#)

Neurodegenerative diseases research group (Prof. Dr. G. Melli)

Parkinson's disease (PD) is a chronic progressive neurodegenerative disease associated with intra-neuronal accumulation of aggregated alpha-synuclein (α -Syn) protein, but unfortunately current knowledge still does not address the underlying pathogenic mechanisms. Consequently, PD lacks both an effective causal treatment and definitive early diagnostic criteria. In the recent years a large amount of evidence has shown that peripheral tissues like skin autonomic nerves, gastro-intestinal mucosa,





salivary glands show immunoreactivity to pathogenic forms of α -Syn. Thus, peripheral tissues, that can be easily biopsied, contain nervous tissue prone to the pathology, and bear potentially important information on disease mechanisms. Our laboratory has a strong translational attitude and aims to set-up novel pathway biomarkers of disease by analyzing skin biopsy and peripheral fluids via advanced new biotechnology. Our group has published several papers on the detection/characterization of α -Syn and tau and its pathogenic species in the skin of PD and other neurodegenerative diseases like atypical parkinsonisms plus we demonstrated that PD patients present a somatosensory small fiber neuropathy that correlates with disease duration and progression. More recently we published several papers on the specific profile of surface proteins of blood- and CSF-derived extracellular vesicles (EVs), which

stratifies patients with PD and atypical parkinsonisms. These EV markers are related to circulating inflammatory and immune cells. We are now profiling prodromal and early phases of disease to identify specific signature and we are investigating how peripheral inflammation and immune dysregulation can contribute to brain neuronal cell death in PD. These projects take advantage of a biobank and clinical database for PD and atypical parkinsonisms that has been established at INSI in Lugano since 2015. [See also chapter 3 on the Laboratories for Translational Research.](#)

Parkinson's disease research group

(Prof. Dr. G. Melli, Prof. Dr. A. Kaelin)

Our research group aims to set-up novel biomarkers for Parkinson's disease by analyzing skin biopsy and peripheral fluids like plasma and cerebrospinal fluid using advanced new biotechnology. Our lab hosts master students USI and from universities abroad for master thesis internship and PhD students in Neuroscience USI. In 2022 we worked on the development of a novel diagnostic model based on plasma-derived extracellular vesicles and machine learning algorithms for Parkinson's disease, supported by national and international grants and we have published an important pioneering article on the characterization of tau protein in skin biopsy that, has raised international interest. The Parkinson research group aims to set-up novel biomarkers for Parkinson's disease by analyzing skin biopsy and peripheral fluids like plasma and cerebrospinal fluid. In 2022, Elena Vacchi successfully defended the very first PhD thesis in Neuroscience at the USI and a new student from India, Ankush Yadav started his PhD. We mainly continued our projects on the development of a diagnostic models based on plasma-derived extracellular vesicles and on skin biopsies, identifying the importance of tau

protein in the skin of patients with atypical parkinsonism.

[See also chapter 3 on the Laboratories for Translational Research.](#)

Epilepsy research group

(Dr. P. Agazzi, Dr. C. Prosperetti)

We are involved in three clinical studies. First, we participate to a swiss multicenter study enrolling adult patients with a first epileptic seizure who undergo electroencephalography (EEG) and brain magnetic resonance imaging (MRI), with specific modalities, to try to identify biomarkers of epilepsy. On the other hand, we collaborate with the cardiological intensive care unit of EOC in a study on the prognostic evaluation of cardiac arrest throughout new blood markers of neuronal death. Finally, we are studying the differences between males and females in subtypes and prognosis of brain glial tumor patients discussed in our neuroncological group.



Headache research group

(Prof. Dr. C. Zecca)

The research focuses on genetic and clinical predictors of efficacy of new migraine treatments, also exploring their safety profile in the post-marketing setting. Epidemiology and pathophysiology of migraine are also investigated. Our Center also participates in various industry-sponsored studies in patients with migraine.

Movement Disorders research group

(PD Dr. S. Galati)

The priority of our clinical research is given to the influence of neuronal plasticity changes during sleep on movement disorders such as dystonia and dyskinesias in Parkinson's Disease. Two translational research projects are ongoing in patients with Parkinson Disease affected by levodopa-induced dyskinesia using high-density recordings and acoustic stimulation.

Multiple Sclerosis research group

(Prof. Dr. C. Gobbi, Prof. Dr. C. Zecca)

The research focuses on diagnosis and treatment, exploring new MRI techniques, understanding better risks associated with new multiple sclerosis (MS) therapies in

aged MS patients and in the context of the COVID-19 pandemic, and searching for innovative neuromodulation treatments of overactive bladder. We also participate in a nationwide investigator-initiated project on MS epidemiology. The Center is also a recognized study site for several industry sponsored international clinical trials.

Ophthalmology research group

(PD Dr. M. Menghini, Dr. G. Grimaldi)

The focus of the ophthalmology research group lies on multi-modal retinal imaging, and retinal degenerative diseases, such as age-related macular degeneration (AMD). We aim to identify surrogate clinical markers that are predictive of the natural disease course and response to anti-vascular endothelial growth factor (VEGF) therapy. In patients with early-stage AMD for example, we analyze imaging parameters that are predictive of progression to the "dry" late-stage of the disease, so-called geographic atrophy. In addition, we conduct various studies analyzing the relationship between function as assessed by scotopic fundus microperimetry, and morphology as assessed by multi-modal imaging in patients with retinal degeneratives, and post vitreoretinal surgical procedures. Funding of the ophthalmology research group is through an unrestricted research grant by BAYER AG Switzerland.

Neuropsychology and Behavioural Neurology research group

(Dr. L. Sacco, PD Dr. G.C. Riccitelli)

The priority of our clinical research is given to the detection and treatment of cognitive and behavioural dysfunctions related to acquired brain damages and diseases in order to better understand the mechanisms responsible for the onset of cognitive alterations, and to identify new rehabilitative-therapeutic strategies. In this context, we are investigating the effect of non-invasive brain stimulation approaches on social cognition abilities in patients with mild cognitive impairment, with the goal of slowing down the progression of the disease. Moreover, we are developing new tests, in collaboration with the Universitäre Altersmedizin (University of Basel), for the detection of social cognition deficits. Furthermore, our research group is conducting scientific collaboration to other EOC departments, and to external institutions such as the collaboration with Laboratories for Translational Research, Eye and Brain Mapping (University of Fribourg). In addition, we are actively involved as co-investigator site in the following multicentric clinical trials:

- The EMBARK study: to evaluate the efficacy and safety of Aducanumab (BIIB037) in subjects with early symptomatic Alzheimer's disease.
- The EORTC 1635 Trial: to investigate cognitive profiles in patients with lower grade glioma following resection: Wait or Treat?
- The MoCA-DCI study: to assess the impact of delayed cerebral ischemia on neuropsychological outcome after aneurysmal subarachnoid haemorrhage.

Educational initiatives such as university masters, symposia and courses, as well as periodic meeting with the general population are a further basic pillar of our mission.

Neuroradiology research group

(Prof. Dr. A. Cianfoni, PD Dr. E. Pravata)

The Neuroradiology Service of the INSI is committed to providing high-level morphological and functional advanced

diagnostic neuroimaging, image-guided spine interventions, as well as endovascular diagnostic and therapeutic procedures. In 2022, several diagnostic and interventional studies have been published in the areas of multiple sclerosis diagnosis, image-guided spine interventions, stroke mimics imaging and brachial plexus neuropathy. These included a study about the risk of false positive results in multiple sclerosis lesion detection, and another study about neuroimaging of neuralgic inflammatory brachial plexopathy.

The following internal projects were started and/or continued:

- A second work package originating from the "High resolution MRI of the optic nerves in Multiple sclerosis" project, focusing on the optic nerve high-resolution normal anatomy by MRI, has been completed. This project was previously founded by the ABREOC and Swiss Multiple Sclerosis Society and received a third grant from the VELUX foundation in 2020. A scientific partnership with the University of Genoa (IT) was also started with this project.
- The "Swiss Cohort" multiple sclerosis registry, sponsored by the University of Basel, a multicentric registry collecting MRI, laboratory and clinical information about patients with multiple sclerosis.
- The diagnostic Neuroradiology team, together with the INSI Neurology service, continued its partnership cooperation in the newly started "SWISS FIRST" multicenter study, regarding advanced MRI biomarkers extraction in first-episode epilepsy patients.

Other active collaborations with the Inselspital of Bern, and the I.T.A.B. (Institute of Advanced Biomedical Technologies) of University of Chieti (IT) in the field of MRI brain tumor research, were also carried on. On the interventional side, there has been continuous effort on ongoing studies mainly on the minimally invasive treatment of osteoporotic vertebral fractures. The feasibility phase of a randomized controlled trial comparing a minimally invasive interventional technique, called SAIF, to the standard surgical stabilization for unstable vertebral fractures is ongoing and close



to completion, in cooperation with the neurosurgery and pain center EOC services. There has been a continuous output of scientific publication on minimally invasive approaches to treat complex osteoporotic vertebral fractures and to perform image-guided interventional procedures in the region of the skull base.

Neurosurgery research group

(Dr. A. Cardia, Dr. F. Marchi)

Research topics concerned cranial and spinal diseases and treatments. The clinical research about cranial neurosurgery has been dedicated to the fields of oncological and vascular neurosurgery. The Neurosurgical service is involved in national and international study protocols (International ReSurge Glioma Network, SWITCH international study for brain hemorrhage, Swiss SOS for intracranial aneurysms and the Swiss trial on space-expanding flap in decompressive hemicraniectomy for stroke). Local research included the prospective study on liquid biopsies for brain tumor (young research EOC-AFRi grant won by Dr. F. Marchi) in collaboration with the EOC Cantonal Institute of Pathology, the intraoperative quantification of fluorescence in brain tumor study in collaboration with SUPSI and the ongoing project on neuromonitoring during aneurysmal clipping surgery. Clinical research in spinal neurosurgery is focused on new intraoperative imaging and navigation technologies in complex spinal surgery in collaboration with the Neuroradiology service and the prospective study on localized amyloid in lumbar spinal stenosis (in collaboration with IOSI and the EOC Cardiocentro Ticino Institute) which reached in 2022 the end of the first step of the project. Moreover, is still active the TRIBECA study (in collaboration with the Maastricht University and the Zuyderland Medisch Centrum in Herleen (NL) that evaluates the impact of a psychological preoperative intervention in patients at risk for chronic pain after lumbar fusion.

Pain Management research group

(PD Dr. P. Maino, PD Dr. E. Koetsier)

Our Pain Management research group has initiated multiple international, multicenter randomized controlled trials, with the aim of improving the efficacy of some chronic pain treatments. The main focus of the research activity is on minimally invasive treatments of pain, such as for example neuromodulation and on the psychological factors that have an impact on the outcome of pain treatments. Internationally, low back pain and neck pain are the leading causes of disability. Our research mission is therefore to explore innovative minimally invasive therapeutic options to treat chronic low back and neck pain resistant to conservative therapy. We are conducting two international randomized controlled trials regarding highly innovative mini-invasive treatments of pain caused by degenerated disc disease. Additionally, we conduct several studies to assess and improve the efficacy of dorsal root ganglion stimulation and spinal cord stimulation to treat back pain and painful polyneuropathy. Furthermore, we are also exploring the impact of psychological factors on chronic pain and how best to address patient expectations. Our research group is also conducting several multidisciplinary research projects (involving colleagues from neurosurgery, neuroradiology, palliative care, orthopedics and neurology). Our research group has also a close academic cooperation with the University of Maastricht (Prof. Dr. S. van Kuijk) and with SUPSI (Prof. M. Barbero).

Peripheral Nerves research group and Center Myosuisse

(Prof. Dr. G. Melli, PD Dr. P. Ripellino)

In 2022 we concluded the NERVES study, focusing on neuropathies related to hepatitis E. In addition, our group published a retrospective multicenter study on neuroimaging findings (ultrasound, MRI) of neuralgic amyotrophy in the acute phase. We continued the translational projects to characterize T cells in Guillain-Barré syndrome (with the ETH Zurich) and to identify genetic determinants of hepatitis E course (with the CHUV in Lausanne). We started a randomized controlled trial

(ongoing) on oral Edaravon for Amyotrophic Lateral Sclerosis (ALS). Furthermore, we actively participated in the Swiss Register of neuromuscular diseases. Our center also includes the “skin biopsy laboratory” for the diagnosis of small fiber neuropathy and the “clinical autonomic nervous system laboratory” for the evaluation of dysautonomic disorders. In this field, we are working on a research project for detection and characterization of amyloid fibrils in skin biopsy of patients with transthyretin and light chain amyloidosis. We collaborated with IOSI (PD Dr. B. Gerber) and with the Charité-University Hospital Berlin, for an observational study on early identification of patients with wild type transthyretin amyloidosis.

Sleep research group

(Prof. Dr. M. Manconi, Prof. Dr. S. Miano, Dr. A. Castelnovo)

The research mission of the Sleep Medicine is to explore brain function during sleep and sleep-related disorders mainly by using innovative electrophysiological approaches. The main field of interest of our research are sleep related movement disorders such as restless legs syndrome and periodic limb movements. In 2022 our group produced a significant amount of high-level scientific articles. These results have been obtained thanks to consolidated local and external collaborations and thanks to the financial support of two competitive grants from the Swiss National Foundation. New therapeutic therapies are under investigation for restless legs syndrome: a phase 2b trial with anti-glutamatergic compound and a study on the effect of electrical spinal stimulation. In recent years, we developed a strong expertise in using High-Density EEG. Other sleep related fields are also represented, among others, by projects on: sleep and pregnancy, sleep and stroke, sleep and attention deficit hyperactive disorder, and infra-slow oscillation. Two important local collaboration have been consolidated, one with the IOSI to evaluate the impact of hormonal treatment on sleep in patients with prostate cancer and in those with breast cancer, and the second on REM behavior disorders in Parkinson Disease.

In collaboration with Bern University, the post-doc Master in Sleep and Consciousness successfully progressed from diploma for advanced studies into a master of advanced studies. The international sleep summer school is in preparation and it will be held in Lugano in July 2023.

Stroke research group

(Prof. Dr. C. Cereda, Dr. G. Bianco)

The research activities of the Stroke Center focus on patient-oriented clinical research: we are principally interested in understanding the role of multimodal advanced neuroimaging (perfusion imaging) for the diagnosis and selection of best treatment for acute ischemic stroke patients together with the Neuroradiology research group. We are currently leading studies addressing to the role of multimodal imaging in patients with ischemic stroke in the posterior circulation (basilar artery stroke), in gender differences in acute stroke treatment and stroke related to small vessel disease. We are also especially interested in recognizing new serum biomarkers to optimize the diagnostic process of transient ischemic attack (TIA). The Center also offers a platform for industry promoted international trials and promotes several other clinical studies and participates to numerous national and international projects mainly dedicated to acute stroke management, stroke prevention and stroke treatment.

Cardiocentro Ticino Institute (ICCT)

Prof. Stefanos Demertzis, MD

Scientific Director

Prof. Marco Valgimigli, MD PhD

Chair Clinical Research

Cardiology

(Prof. Dr. G. Pedrazzini)

Clinical Electrophysiology

Prof. Dr. A. Auricchio)

The Clinical Electrophysiology research group is focused on cardiac disorder diagnosis and management including atrial fibrillation, cardiac implantable devices, management of lead and device issues, remote patient monitoring, computer modelling, and advanced signal processing. The group collaborates with Fondazione Ticino Cuore, the InterDisciplinary Institute of Data Science at USI (Prof. A. Mira) and MeteoSwiss; a research cluster on out-of-hospital cardiac arrest (OHCA) has been established. The project makes use of advanced spatial statistics methods to identify patterns of temporal-spatial distribution of cardiac arrests in Canton Ticino. This project is supported by SNSF. Furthermore, additional research activities on OHCA in Switzerland have been successfully carried out in collaboration with the Swiss "Interverband für Rettungswesen" Association. Finally, the The Clinical Electrophysiology research group has a longstanding research partnership with EULER Institute of USI to conduct computer modeling studies in atrial fibrillation and heart failure.

Interventional Cardiology

(Prof. Dr. M. Valgimigli, Dr. M. Moccetti (until 31.8.2022), PD Dr. D. Sürder, from 1.1.2023)

The interventional cardiology research group participates in several national and international trials, which are coordinated by the research cardiovascular service (SRC) and runs multiple international interventional studies under the leadership of Prof. Dr. Valgimigli as global Principal Investigator (PI). The group is still running the NATURE trial, which is investigating the role of a new thrombectomy device for the management of patients with ST segment elevation myocardial infarction with large thrombus burden and involves eight sites in Switzerland and Italy. Furthermore, the group is actively participating as recruitment sites in multiple randomised trials and registries, among which SWISS TAVI, MITRASWISS, SOS AMI, PROTECT 2 and 4, PICSO 5, Amundsen, Victorion Plaque, MOCA 1 and Ex-VIVO. This research group has published this year as first or senior author(s) multiple sub-analyses of the MASTER DAPT Trial, which involved 130 sites in Europe, Asia, South America and Australia and investigated the duration of dual and single antiplatelet therapy in patients at high risk of bleeding who underwent PCI. Two individual patient data meta-analyses were published in the two most prestigious cardiological journals, including the European Heart Journal and Circulation addressing the optimal antiplatelet treatment in patients at high

bleeding risk and the value of radial versus femoral access site in patients undergoing cardiac catheterization, respectively. In 2022, this group has published a First-in-Human study on the use of a stent retriever as new thrombectomy technique for patients with large thrombus burden. Since late 2022, we host 2 interventional fellows (Dr. A. Landi and Dr. S. Cagliani) with each having 50% active role in the clinical and 50% in the research field.

Cardiac surgery

(Prof. Dr. S. Demertzis)

Interprofessionalism & Communication

Interprofessional electronic patient communication: The COVID-19 pandemic restrained in an unprecedented manner the face-to-face communication of families seeking information on their hospitalized loved ones and the interprofessional care teams (physicians, surgeons, nurses, physiotherapists). Those restrictions were generally perceived negatively. We developed a software-based communication solution to proactively and regularly convey all relevant information to the authorized family members. A prospective RCT, significantly supported by the USI (Prof. Schultz - Faculty of Communication) and SUPSI (Prof. Bianchi - Nursing Science) was conceived and completed. The positive results were reported in 2022 (A. Pozzoli, C. Zurfluh, S. Demertzis) at the Annual Conference of the Swiss Society of Cardiac Surgery (SGHCCH).

Extracorporeal Circulation

1) Intraoperative monitoring of the gaseous emboli (micro-air bubbles) that travel through the heart-lung machine and are filtered by the several machine components was introduced. Despite filtering, in some cases clouds of tiny bubbles (generally <200µm) still enter the patient and we are assessing their correlation with systemic inflammatory reactions or subtle neurological effects (S. Demertzis, S. Vandenberghe, M. Puthettu). Funded by the Swiss Heart Foundation.
2) Protecting the brain while treating the heart is a block of two projects focusing on quantification of air-bubbles captured in the heart chambers during open-heart surgery. One of the subprojects (AIRCATCH)

is announced to be funded by the Fondo EOC-USI (Fondazione Daccò) and the other is funded by the Foundation Fidinam (S. Demertzis, S. Vandenberghe, R. Krause – USI).
3) Analysis of venous drainage in extracorporeal circulation using different types of venous cannulas (E. Ferrari in collaboration with the University of Lausanne).

Coronary Artery Bypass Grafting

The European Registry of Clinical Outcomes after external support of saphenous vein grafts (VEST) after coronary artery bypass grafting coordinated by S. Demertzis (PI) concluded patient enrollment and is registering the foreseen follow-up interviews up to 5 years after surgery.

Cardiac Valves

1) Two ongoing research projects focus on surgical techniques to reduce early complications of rapid-deployment sutureless aortic valves, paravalvular leakage and heart-blocks requiring implantation of a pacemaker (E. Ferrari).
2) Predictability of failure of surgical mitral valve repair analyzing preoperative imaging using machine learning. Initialization of a common project in collaboration with the USI (Prof. Krause - Euler Institute) (T. Torre, S. Demertzis).
3) Analysis of clinical data and outcome after implantation of transcatheter aortic valves (TAVI) in general for severe aortic stenosis (focusing on alternative access than the femoral artery) and in presence of concomitant significant tricuspid valve regurgitation (E. Ferrari).
4) Analysis of the predictability of clinical outcome after percutaneous implantation of Mitraclip for severe mitral regurgitation, depending on intraprocedural measurements of the pressure in the left atrium (E. Ferrari in collaboration with University of Tokyo and University Hospital San Raffaele of Milan).
5) Analysis of long-term outcomes under different anticoagulation / antiaggregation regimes after TAVI registered in the SWISS-TAVI registry is underway (E. Ferrari in collaboration with the University of Bern).
6) Analysis of the radiologically documented position of TAVI-in-TAVI implantations and the corresponding clinical outcomes (E. Ferrari in collaboration with the Washington University).

Thoracic Aorta

1) Analysis of clinical outcomes after thoracic aorta endoprostheses (TEVAR) for different aortic pathologies (T. Theologou, E. Ferrari, S. Demertzis).

2) Retrospective analysis of aortic size indices in patients with acute aortic dissection type A (S. Grego, S. Demertzis).

Cardio-anesthesiology and intensive care

(Prof. Dr. T. Cassina)

A prospective clinical study focusing on the kinetics of Neurofilament Light Chain after cardiac arrest as prognostic parameter of neurological damage is ongoing (in collaboration with INSI). Our group has provided support for a MD thesis (Dr. T. Preda) on traumatic injuries after manual and mechanical cardiopulmonary resuscitation. We were involved in Swiss Latin Airway Guidelines: a holistic approach for improving airway management. We participate on a RCT (STEP CARE trial) on sedation, temperature and pressure after cardiac arrest and resuscitation. Dr. G. Via is member of the panelist of the Society of Critical Care Medicine (SCCM) Task Force on the Ultrasound Guideline Focused Update and member of the panelist of the American Society of Echocardiography (ASE) writing group on Nomenclature Standards for Cardiac Point-of-Care Ultrasound Imaging. We have also conducted a study on cost-effectiveness and diagnostic accuracy of focused cardiac ultrasound in the pre-participation screening of athletes.

Lugano cell factory (LCF)

(Dr. L. Turchetto)

LCF is a Good Manufacturing Practice (GMP) facility authorized by Swissmedic for the production of Advanced Therapy Medicinal Products (ATMP)/transplant products. After optimizing and validating the manufacturing process for production of hyaline cartilage beads (Cartibeads) from human chondrocytes for the startup company Vanarix SA, LCF became the first clinical manufacturing site of microtissues for knee cartilage regeneration. The first

patient has already been successfully implanted in 2022 upon Swissmedic's approval of the first Cartibeads-based Swiss trial. LCF has also established a collaboration with the University of Manchester (UK) to act as a GMP manufacturing center for a gene and cell-based therapy product related to the treatment of Duchenne Muscular Dystrophy. LCF performed process validations and also contributed to preparing the regulatory dossier recently submitted to the regulatory authorities for approval of phase 1 clinical trial. The LCF developed a novel protocol for the production of clinical-grade human exosomes derived from cardiac progenitor cells [patent application N° 102017000035315_ (IT/CH/PCT) entitled "Device and method for exosomes production and purification"]. In this context, LCF is presently participating to a European consortium (MARVEL project/ H2020 grant) devoted to improve the purity of the GMP grade exosomes for diagnostic and/or clinical applications. The LCF is also qualified to operate as cardiovascular tissue bank (aortic and pulmonary valves, as well as pericardium for human use), which already delivered several cardiovascular tissues for patient treatment.

Laboratory research

[See chapter III on the Laboratories for Translational Research](#)



Imaging Institute of Southern Switzerland (IIMSI)

Prof. Filippo Del Grande, MD
Medical and Scientific Director

The Imaging Institute of Southern Switzerland (IIMSI) includes the Clinic of Radiology (Head: Prof. Dr. F. Del Grande), the Clinic of Nuclear Medicine and Molecular Imaging (Head: Prof. Dr. L. Giovanella) and the Medical Physics Service (Head: Dr. S. Presilla).

Clinic of Nuclear Medicine and Molecular imaging

Integrated diagnostics
(Prof. Dr. L. Giovanella)

- Artificial intelligence and machine learning to develop integrated diagnostic expert systems/neural networks. Collaborations: University of Padua, ICT EOC, Roche Diagnostics.
- Molecular imaging, molecular pathology and circulating biomarkers of thyroid cancer. Collaborations: Universities of Marburg, Essen, Paris Descartes, Zurich, Udine, Thyroid Committee-European Association of Nuclear Medicine - Vienna, EO Ospedali Galliera Genoa.

Thyroid diseases

(Prof. Dr. L. Giovanella)

- Radioiodine-refractory thyroid cancer and re-differentiation strategies. Collaborations: Center of Excellence on Radioiodine Theranostics-Society of Nuclear Medicine and Molecular Imaging.
- Mitochondrial imaging in thyroid diseases. Collaborations: EO Ospedali Galliera Genoa, Thyroid Committee-European Association of Nuclear Medicine - Vienna,

University of Turin, University of Messina, Universities of Augsburg, University of Giessen.

PET radiomics and predictive imaging of lymphomas

(Prof. Dr. L. Ceriani)

- Development and validation of new PET-metrics, radiomic analysis and Artificial Intelligence tools. Collaborations: IELSG, SAKK, IOR, IOSI, Amsterdam University Medical Centre, Federazione Italiana Linfomi, United Kingdom Cancer Research Institute.
- Integration of PET/CT data, biological and genomic features (i.e., liquid biopsy) using Radiomics, Genomics and Artificial Intelligence tools to model new diagnostic/prognostic approaches. Collaborations: IELSG, SAKK, IOR, IOSI, Amsterdam University Medical Centre, Federazione Italiana Linfomi, United Kingdom Cancer Research Institute.

Theragnostics

(PD Dr. G. Paone)

- Clinical/research activities in theragnostic field for the therapy working group of Swiss Society of Nuclear Medicine (SGNM) particularly focused on PRRT and PSMA therapy (collaborations with Universitatsspital Bern, Novartis/AAA).
- Personalized dosimetry studies focused on selective internal radiotherapy with Y-90 (collaborations with Istituto Nazionale Tumori Milan, Boston Scientific and Mirada group).

- Production and updating therapeutic flow-charts in NETs and combined therapy protocols in mCRPC.

Evidence-based nuclear medicine

(Prof. Dr. G. Treglia)

- Production of evidence-based documents (systematic reviews and meta-analyses of diagnostic test accuracy) and guidelines on PET/CT. Collaborations: Cochrane and European Association of Nuclear Medicine, University of Lausanne, University of Groningen, University of Brescia, University of Mashhad, Fondazione Policlinico A. Gemelli Rome and EO Ospedali Galliera Genoa.
- Research-related activities for the PET working group of the Swiss Society of Nuclear Medicine focused on the achievement of reimbursement for new PET indications.

Clinic of Radiology

The Clinic of Radiology of IIMSI offers diagnostic and interventional imaging examinations in the main four sites of EOC (Bellinzona, Locarno, Lugano, Mendrisio). The Clinic of Radiology offers a complete coverage of the body imaging, with radiologists divided into subspecialties, to enhance specific skills and competence, in order to offer the patients and their referring physicians a highly specialized radiological service. Among the different subspecialties, the below mentioned are the most active in research, as reflected by their scientific production.

Musculoskeletal imaging

(Prof. Dr. F. Del Grande)

The group of radiologists dedicated to musculoskeletal imaging, are conducting research projects, monocentric as well as multicentric, with the objective of improving image quality without prolonging the time of acquisition, with the help of artificial intelligence mediated approaches. Furthermore, the advanced technology of the Institute allows to test new technologies, developments and sequences aiming to a better understanding of different features of muscles, tendons, ligaments and nerves,

such as (but not limited to) diffusion tensor imaging and dixon sequences.

Oncologic imaging

(Prof. Dr. S. Rizzo)

The group of radiologists dedicated to oncologic imaging are conducting several research projects with the aim of understanding how quantitative imaging of body composition is associated with surgical and oncological outcomes in cancer patients. With this end, different approaches are under evaluation and are the core of many ongoing research projects, one of which funded by the Swiss Cancer League. Furthermore, the group is involved in national and international collaborations evaluating artificial intelligence mediated approaches (including radiomics, machine learning and convolutional neural networks) for prognostication in different cancer settings.

Uro-genital imaging

(Prof. Dr. R. Wyttenbach)

The radiologists dedicated to the urogenital diseases are involved in a multicentric project aiming to find an artificial intelligence based way to detect prostate cancer with magnetic resonance imaging.

Breast imaging

(Dr. S. Schiaffino)

The radiologists of the breast imaging team are involved in the research projects of the CSSI (Centro di Senologia della Svizzera Italiana) and in international multicenter studies. Part of the team is developing systematic reviews and meta-analyses in the high-risk breast lesions' field.

Institute of Pediatrics of Southern Switzerland (IPSI)

Prof. Giacomo Simonetti, MD
Medical and Scientific Director

Surgical and medical pediatrics focuses on the health of neonates, infants, children, and adolescents, their growth and development, and their opportunity to achieve full potential as adults. Pediatric care encompasses a wide spectrum of health services ranging from preventive health care to the diagnosis and management of acute and chronic (often congenital) diseases.

IPSI has a dedicated research unit, conducted by Prof. Dr. Simonetti, PD Dr. F. Vanoni, PD Dr. B. Goeggel Simonetti, PD Dr. M. Rizzi, Dr. A. Stefani-Glücksberg, Dr. C. Balice, and scientific assistant A. Severi Conti. The pediatric research unit is included in the SwissPedNet, a Swiss based network that promotes pediatric clinical research in Switzerland.

The research unit is mainly oriented on clinical research in different fields of pediatrics: cardiology, infectious disease, neuropsychiatry, nephrology, nursing sciences, oncology, rheumatology and pediatric surgery.

The principal areas of research for each subspecialty are:

Cardiology-Nephrology

(Dr. C. Leoni-Foglia, Dr. A. Stefani-Glücksberg, Prof. Dr. M.G. Bianchetti and Prof. Dr. G.D. Simonetti)
Childhood hypertension, cardiovascular risk factors in the young, Kawasaki disease. We participate in the Swiss Pediatric Heart Cohort and the International Pediatric Brugada Registry. Prof. Simonetti

collaborates in the HyperChildNet board, a COST Action of the EU with the aim at improving awareness of childhood hypertension.

General Pediatrics

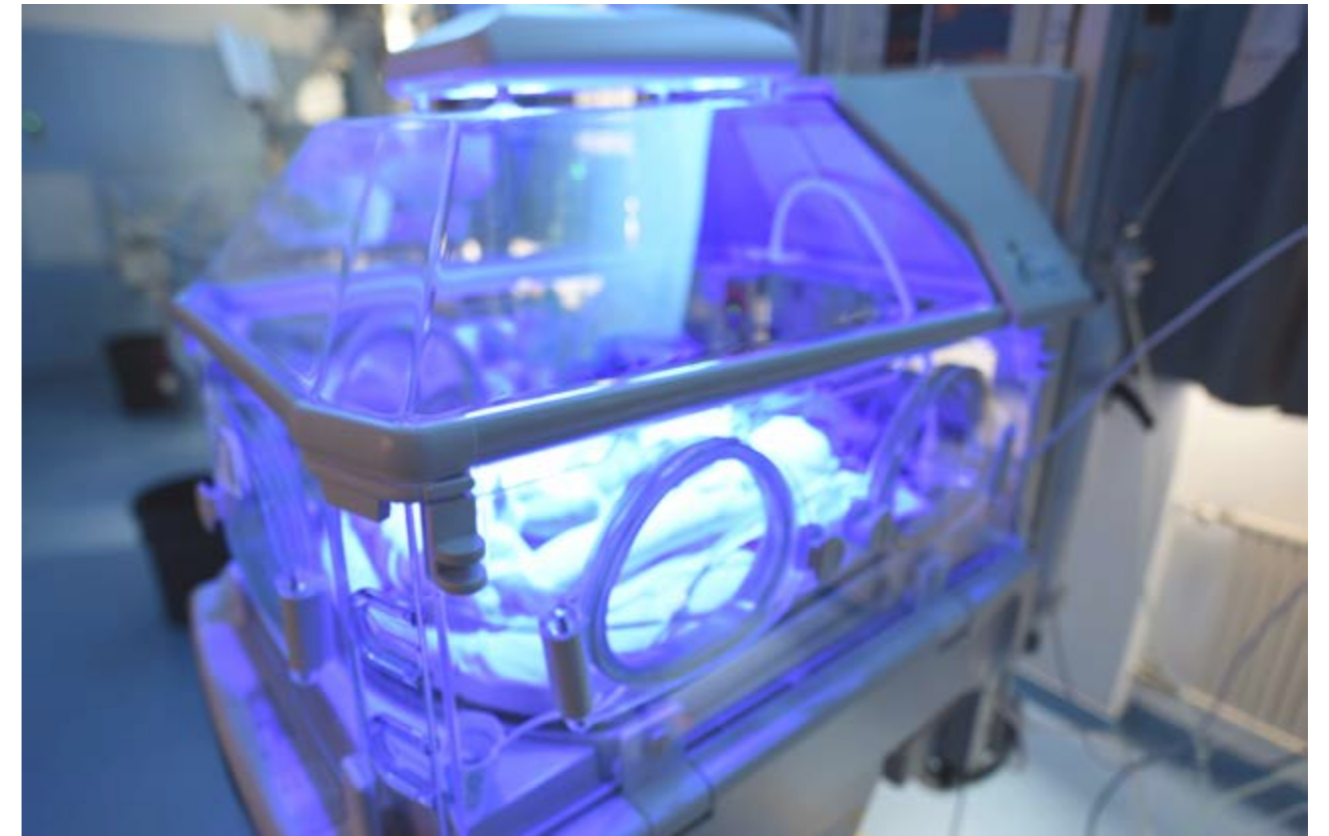
(Prof. Dr. M.G. Bianchetti, Prof. Dr. G.D. Simonetti, PD Dr. F. Vanoni)

Systematic reviews of rare diseases, or rare complications of frequent pediatric diseases. Moreover, fever phobia among parents in collaboration with the Institute of Public Health at USI in Lugano will be analyzed. We conducted also a prospective, observational, single center, cohort study on the bioimpedance measurements to assess the change of fluid status in pediatric patients undergoing general anesthesia. We are involved in the MenuCH-Kids study set up which will take place between 2023 and 2024 and will involve a sample of children from Ticino. This study will provide important information on the diet and nutritional status of children in Ticino.

Infectious disease

(Dr. L. Kottanattu)

We participate in a multicentric RCT on pediatric inflammatory multisystem syndrome temporally associated with SARS-CoV-2 (RECOVERY study). We are involved in several registries (MoCHIV, CITRUS) that analyze the prevalence of rare infectious diseases in childhood. In 2020 we started a seroprevalence study of COVID-19 in pediatric population in collaboration with Humabs, covering the different waves of pandemic.



Neuropsychiatry

(PD Dr. B. Goeggel Simonetti, Prof. Dr. G.P. Ramelli)

In collaboration with SUPSI we promote a project on the early diagnosis of autism (Auto-Play-D). As members of national and international research consortiums we participate in studies on neurovascular diseases in childhood and young adults (IPSS, SNPSR, RCT PASTA Trial, intracranial dissection, pediatric FMD), epilepsy, autism spectrum disorders and cerebral palsy, inflammatory brain disease (Swiss-Ped-IBrainD), long-term follow-up of high-risk neonates (Swiss Neo Net), head trauma.

Onco-hematology

(PD Dr. M. Rizzi and Dr. P. Brazzola)

We are involved in 31 international studies (oncology protocols)/registry. We contribute to a non-interventional study for treatment of patient with hemophilia A with and without inhibitors (EMILL, Roche).

Rheumatology

(PD Dr. F. Vanoni)

We participate in an international cohort rheumatology diseases registry. We strictly collaborate with the CHUV for the development of observational prospective and retrospective studies on treatment and outcome in inflammatory diseases.

Pediatric Surgery

(Dr. M. Mendoza, Dr. V. De Rosa and Dr. E. Montaruli)

Minimal invasive surgery, orthopedic and surgical trauma. The main goal of the research unit is to promote high-quality pediatric clinical research in the Italian speaking part of Switzerland and to help young fellows and medical students of Swiss Universities to participate in clinical research in pediatrics. In the context of the Master medical school, we are also involved in 10 master thesis supervision and promotion of clinical research.

Institute of Pharmacological Sciences of Southern Switzerland (ISFSI)

Prof. Alessandro Ceschi, MD, MSc
Medical and Scientific Director

The Institute of Pharmacological Sciences of Southern Switzerland (ISFSI), includes the Division of Clinical Pharmacology and Toxicology - with its clinical services including the Pharmacogenomics and Pharmacogenetics Unit, the Regional Pharmacovigilance Centre, and a small Research Unit - and the Central Pharmacy Service - with the Hospital Pharmacy, Clinical Pharmacy, and Public Pharmacy. We advise other departments and healthcare professionals on safe drug use and promote the rational use of medicines. The final goal of our activities is to personalise drug therapy by optimising beneficial effects while minimising adverse effects and costs.

During 2022, the ISFSI Research Unit has been continuing the signal detection activity for the surveillance of COVID-19 vaccine safety, which started in July 2021 on behalf of and in collaboration with Swissmedic. Since then, the Research Unit has conducted a systematic analysis of VigiBase®, the World Health Organization's global pharmacovigilance database, performing data mining through disproportionality techniques on safety reports originating in Switzerland and reporting adverse events following immunization associated with Spikevax® and Comirnaty® vaccines. Noteworthy, this activity revealed an early safety signal of disproportionate reporting for paraesthesia

with Spikevax® in Switzerland. This occurred few months before the European Medicines Agency assessed and validated the same signal, which finally led to the addition of paraesthesia to the European summary of product characteristics of Spikevax®. Moreover, shortly after the booster vaccination campaign began in Switzerland in December 2021, an increasing number of cases of delayed urticaria were reported in association with Spikevax® to Swissmedic. Therefore, the ISFSI Research Unit, in collaboration with Swissmedic, performed a descriptive analysis of that case series suggesting that, albeit the inherent limitations of a spontaneous reporting system, the booster vaccination with Spikevax® might be a trigger for delayed urticaria. Subsequently, the Swiss product information of Spikevax® was updated.

Furthermore, ISFSI was actively contributing to a range of studies on different aspects of COVID-19 with a focus on immunological and vaccine-related aspects.

ISFSI is also an official sentinel centre of the European Drug Emergency Network Plus (Euro-DEN Plus) Project, which collects data on Emergency Department (ED) acute recreational drug presentations across Europe. Whilst not representative of the general population of drug users as

a whole, by using a sentinel centre model covering, to date, twenty-three European and neighbouring countries, the Euro-DEN Plus database provides a snapshot of the epidemiology of ED presentations related to acute recreational drug toxicity in Europe, and gathers systematic and structured toxicology surveillance data relative to such events. During 2022, we performed a matched case-control- study in the Euro-DEN Plus database to assess determinants of intensive care unit (ICU) admission following presentation to the ED because of acute recreational drug toxicity (manuscript to be submitted). Within the Euro-DEN Plus, we also contributed to two other studies; one assessing the clinical impact of ethanol co-use in patients presenting to the ED with acute toxicity involving the use of central nervous system depressant drugs, and the other describing ED presentations with acute recreational drug toxicity involving new psychoactive substances.

A further branch of the research activity conducted at ISFSI was in collaboration with INSI EOC within the Swiss Stroke Registry, a national web-based registry that includes all consecutive patients admitted to Stroke Units and Stroke Centres across Switzerland. During 2022, we concluded an observational cohort population-based study aimed at assessing sex differences in the safety and efficacy outcomes of intravenous thrombolysis in acute ischemic stroke patients in relation to preadmission use of antiplatelet agents. From this initial work in the field of sex differences in relation to outcomes of acute ischemic stroke treatment(s), we have just designed an observational study to be conducted within the European EndoVascular treatment and

Thrombolysis for Ischemic Stroke Patients (EVATRISP) registry to assess sex-specific impact of pre-stroke antiplatelet therapy on functional and safety outcomes of endovascular therapy.

In 2022, the ISFSI Research Unit conducted two studies on drug safety in pregnancy by using VigiBase®. A pharmacovigilance study assessed the safety of immune checkpoint inhibitors in pregnancy observing that up to date there are no signals of increased reporting of spontaneous abortion, foetal growth restriction and prematurity with immune checkpoint inhibitors compared to both other drugs in general and other antineoplastic agents. A second study, which aimed to provide an updated safety assessment of the use of anti-migraine monoclonal antibodies targeting the calcitonin-gene related peptide system in pregnancy, showed no safety signals of foeto-maternal toxicity according to VigiBase® safety reports (manuscript under revision).

Lastly, by the end of 2022, ISFSI joined the REporting of A Disproportionality analysis for drUg Safety signal detection project (READUS PV, <https://readus-statement.org>) becoming member of the multinational working committee involved in the development and validation of the first reporting recommendations for studies using disproportionality analyses in databases of spontaneously reported adverse events.

Institute of Laboratory Medicine (EOLAB)

Dr. Franco Keller, PhD
Director

The Institute of Laboratory Medicine (EOLAB) was set up to meet the requirements of modern clinical laboratory medicine. EOLAB recognizes the importance of research into its activity to be continuously up-to-date and to offer the best service and collaboration to all medical departments. The EOLAB specialists are actively involved in several scientific society boards as soon as nationally and international scientific society working groups.

The collaboration between EOLAB, the EOC Clinical Trial Unit, and ICT area allowed the creation of the EOC Biobank during 2020. The headquarters of the EOC Biobank is located inside the Service of Microbiology (SMIC). During 2022 EOLAB continuously collaborated with EOC Biobank providing infrastructure, logistic support and technical staff.

In the 2022 EOLAB activate 7 new collaboration studies with a total of 27 collaboration in progress during 2022.

The Service of Microbiology (SMIC) contributes with its known-how and data to EOC internal projects as well as to external national and international projects. During 2021-2022 microbiological examinations were pursued by HIV and HCV patients in the frame of national HIV and HCV cohort studies and national projects. Case reports on clinically significant cases with uncommon bacterial and fungal strains were published in collaboration with clinicians from EOC and colleagues of Swiss

university hospitals. SMIC collaborated with INSI on a research project to establish a proven correlation between Hepatitis E viral infections and neurological manifestation and to evaluate Hepatitis E virus (HEV) involvement in Guillan-Barré Syndrome. The project has been extended toward additional viral agent, which could be implicated as immuno triggers in nevralgic amyothrophy. Several projects as been conducted on SARS-CoV-2 infections on immunocompetent as well as non-immunocompetent patients. Collaborations were established with EOC researchers as well as with colleagues of university hospitals. In particular, SMIC actively participated to the Swiss SARS-CoV-2 sequencing genomic project. Whole genome sequencing has been validated and applied to several isolates of SARS-CoV-2 during the first phase of the national sequencing project. Resulting sequences and genetic typing have been transferred to national and international databases.

This project offered also a very useful instrument for monitoring SARS-CoV-2 to the cantonal health authorities. SMIC also hosts the National Reference Laboratory for Legionella and contributes to national collaborations and international projects on the epidemiology and spread of Legionnaires' disease in Switzerland.



The Clinical Biochemistry and Pharmacology Laboratory (BFC) is involved in a European network for the harmonization of steroid hormone quantitation in mass spectrometry and close collaboration with the pharmacology unit of the University of Turin and Amedeo di Savoia Hospital in Turin on pharmacology and pharmacogenomics fields.

Cantonal Institute of Pathology (ICP)

Prof. Luca Mazzucchelli, MD
Medical and Scientific Director

The EOC Cantonal Institute of Pathology (ICP) is actively involved in various research projects that aim to identify new methods to be introduced into routine diagnostics or to identify new markers that can be used both in prognosis and in predicting the efficacy of chemotherapeutic treatments: ICP research is essentially aimed at the molecular characterisation of diseases, mainly those in the field of oncology. IOSI/IOR are privileged research partner of the ICP, with particular emphasis on cancers of the gastrointestinal tract and thoracic cavity, hematolymphoid neoplasms and urogenital cancers. Currently, ICP participates in almost 50 clinical trials and also collaborates with various Swiss and international pathology institutes.

Digital pathology and artificial intelligence

(Prof. L. Mazzucchelli, Dr. V. Martin)

Digitalization of pathology offers new opportunities to improve clinical service but also to develop research projects. The ICP is currently going through a gradual transition to digital pathology. In parallel, working groups have been formed in collaboration with Delle Molle Institute for artificial intelligence (IDSIA) to develop new diagnostic algorithms but also artificial intelligence applications to exploit the data that will be produced. The potential is enormous and the ICP remains open and available for any project in collaboration with other EOC institutes/departments. Specifically, the Institute is working on a project to efficiently reconstruct the

histological cross-sections of large tumours that do not fit a single microscope slide and thus they have to be split in multiple smaller fragments. This reconstruction process is currently performed by pathologists in an approximate manner. An AI-assisted tool would allow more accurate assessment of important parameters useful in patient management, such as the relationship of a tumor to resection margins or tumor regression in patients receiving neoadjuvant treatment. Such a tool has enormous potential in terms of the increased quality of anatomo-pathological diagnoses, saving time in issuing diagnoses, improving patient treatment, and standardizing important parameters evaluated in clinical trials. A grant application has recently been submitted to Innosuisse.

Analysis of circulating tumour DNA in patients with non-small cell lung cancer (NSCLC) undergoing immunotherapy with checkpoint inhibitors

(Dr. M. Frattini)

For patients with metastatic lung cancer and a high expression of the immunological marker PD-L1, therapy with the immune checkpoint inhibitor pembrolizumab or atezolizumab is the standard therapy. Recent phase II clinical trials and interim analysis from phase III trials have demonstrated a great benefit from the addition of tiragolumab (another checkpoint inhibitor) to atezolizumab in patients characterized by PD-L1 expression >50%. Today, the response of these patients



to therapy is checked every 9-12 weeks using computer tomography. This means that in a substantial proportion of patients, tumour progression is only detected when the general condition is worst and further tumour therapy is no longer possible. Recent data seem to indicate that a quantitative decrease in the analysis of DNA in plasma (ctDNA) correlates with the response to immunotherapy. The aim of the study is to use the quantitative progression of ctDNA to optimise the therapy. Patients with an early drop in ctDNA (>50%) compared to the initial level will continue with immunotherapy with the combination of atezolizumab and tiragolumab checkpoint inhibitors. Patients without a corresponding drop in ctDNA will be randomly assigned to one of two treatment groups: either they will also be further treated with atezolizumab+tiragolumab alone or they will receive platinum-based chemotherapy in addition to the combination of checkpoint

inhibitors. With this study, we want to investigate whether early intensification of therapy in patients with insufficient drop in ctDNA after 2 cycles of immunotherapy may lead to a better prognosis and we want to investigate whether ctDNA measurement is a suitable biomarker for risk stratification and therapy adjustment. This is a study co-directed with IOSI (Dr. P. Froesch), more than 10 Swiss centres (through SAKK involvement) and more than 10 European centres (from France, Spain, Germany, Italy). The trial will receive the financial support of Roche.

Sensitive screening in treatment of skin cancer

(Dr. M. Frattini)

Melanoma is a severe disease, relatively insensitive to standard chemotherapeutics, but recently vemurafenib, dabrafenib and trametinib have shown promising results. However, the efficiency of these therapies is strongly correlated with the presence of somatic mutations (SMs) in the BRAF oncogene in the cancer cells. Furthermore, recent data indicate that trametinib may benefit patients affected by SMs in the NRAS gene and moreover, SMs occurring in the KIT gene are correlated with susceptibility to other targeted drugs like imatinib. The aim of the project is to develop, in collaboration with a Danish company, three assays 20-200 fold more sensitive than available methods (Next Generation Sequencing and FDA-approved assays) also facilitating non-invasive monitoring.

Sensitive diagnosis, prognosis and treatment planning on open platform of glioma brain tumours

(Dr. M. Frattini)

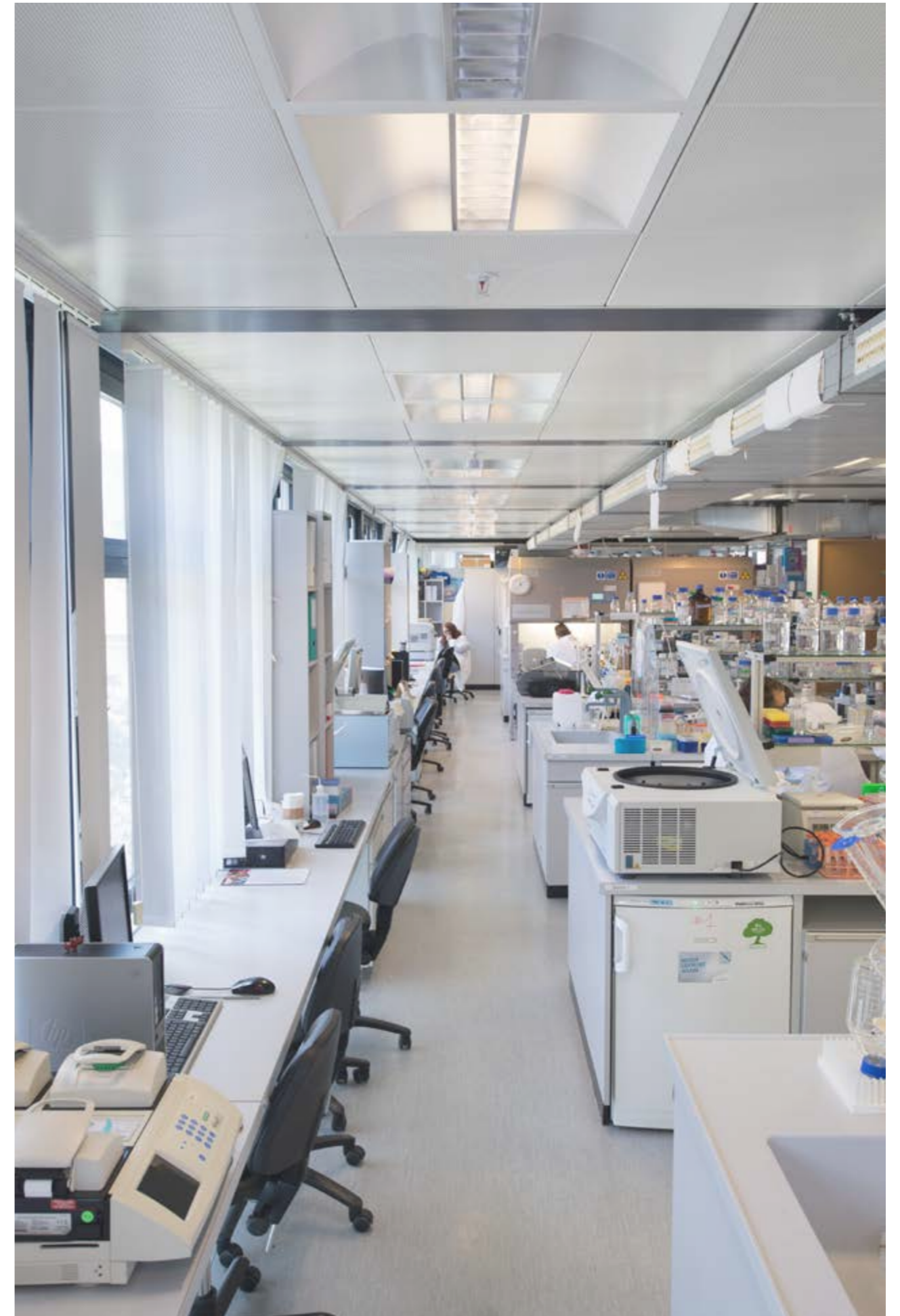
Brain tumors are one of the most aggressive and common cancers. About 28% of brain tumors are gliomas; glioblastoma (GBM) subtype is found in 17% out of this group. The classification and the treatment of gliomas is strictly correlated to the evaluation of several molecular markers (i.e., mutations in IDH1, IDH2, hTERT, H3F3A, HIST1H3B genes; MGMT (O6-methylguanine DNA methyltransferase) promoter methylation). The genetic status of IDH1, IDH2, H3F3A and HIST1H3B is fundamental for a proper classification of the tumor; MGMT promoter methylation status has clinical relevance because patients with hypermethylation benefit more from the combination of chemo (based on temozolomide) and radiotherapy; finally, TERT promoter genetic status is linked to adverse prognosis and to diagnosis of oligodendroglioma or glioblastoma. Nowadays, the available molecular assays are essentially made on tissue samples and suffer from method variation, low sensitivity and technical barriers.

On these bases, the main purpose of our project is to improve these analyses providing a new, more sensitive and faster real-time PCR based methodology in collaboration with PentaBase ApS, a Danish company with which we have excellent collaborative relationships from years. In collaboration with INSI (Dr. F. Marchi) and IOSI, liquid biopsies will be collected from patients affected by gliomas and evaluated with the newly developed assays.

Personalized cancer monitoring

(Dr. M. Frattini)

Monitoring of cancer progression is mainly done by bi-monthly imaging of the tumor. Cancer patients can be better treated if their cancers are monitored closely. PCR-based genetic analysis of cancer specific mutations is a fast, inexpensive alternative. However, this method is only available for patients carrying typical (hotspot) mutations in a few cancers. Using simple blood samples, we will enable sensitive and inexpensive monitoring of all cancers for true personalized treatments. Based on next generation sequencing (NGS) data from the clinics from the solid, diagnostic biopsy, in collaboration with a Danish company we will design, manufacture and validate tumor- and patient specific, real-time PCR-based monitoring and ctDNA purification assays for blood samples. We will sell at least twenty assays per patient to the clinic, allowing for bi-weekly monitoring and early response to resistance acquisition of tumors.



Department of Medicine EOC

Prof. Pietro Cippà, MD
Head of Department

The Department of Medicine EOC supports and promotes translational and clinical research in several areas related to internal medicine. The research activity of the different units is coordinated by the Local Research Unit (LRU) of the Department of Medicine. We report here the most relevant clinical research projects of the year. The work on translational research performed by the Hepatology (Prof. A. De Gottardi) and the Nephrology (Prof. P. Cippà) groups is included in the section dedicated to the Laboratory of Translational Research EOC.

General internal medicine

(Dr. B. Balestra, Prof. Dr. L. Gabutti, PD Dr. O. Giannini, Prof. Dr. A. Ognà, Prof. Dr. M. Pons)

Quality and appropriateness of care, inspired by the Choosing Wisely campaign, involving in particular medication and laboratory test prescriptions, are among the most important research lines of Prof. L. Gabutti. His group is also working on salt consumption, natriuria in stress hypertension, arterial stiffness, and cardiovascular consequences of magnesemia. Arteriosclerosis (CATCH study, Prof. Dr. A. Gallino) and cardiovascular risk factors, including studies on discontinuing statins in multimorbid older adults without cardiovascular disease (STREAM and STREAM-SubATS) is another topic of interest. PD Dr. O. Giannini has started a new project aiming at a detailed characterization of the adaptive immune response to seasonal influenza vaccination in healthy volunteers and in patients on hemodialysis in collaboration with Humabs BioMed SA (Kidney-Flu

Infectious diseases

(Prof. Dr. E. Bernasconi)

Our group is involved in the OCTOPLUS study, a randomized multi-center controlled trial comparing conventional X ray, low-dose computer tomography and lung sonography in the diagnostic accuracy in patients with pneumonia. In collaboration with the University Hospital of Geneva we are studying the immune response to the new Herpes Zoster vaccine (Shingrix) in HIV-infected persons and controls (SHINGR'HIV). We participate to international studies developing a new antiretroviral dual therapy (islatravir/doravirine). We started a collaborative study with the IRB aiming to evaluate and characterize autoantibodies targeting cell trafficking in HIV-1 infection. We continue the development of an antibiotic stewardship program following the COMPASS study. We continue to actively contribute to the Swiss HIV Cohort Study (SHCS) (www.shcs.ch), one of the world's largest epidemiological and clinical studies in the field of HIV.

Nephrology

(Prof. Dr. P. Cippà)

The InterACTIVE-HD2.0 study, an international collaboration supported by the Interreg program, aims at improving quality and accessibility to hemodialysis by implementing different strategies, including algorithms to predict dialysis complications (Dr. S. Pianca, Dr. D. Giunzioni). We started a new line of research focused on peritoneal dialysis to support an evidence-based



approach for a personalized selection of the dialysis modality (Dr. A. Bellasi). We investigate the role of FGF23 in cardiac arrhythmias in dialysis patients (Dr. A. Bellasi). Moreover, we work on experimental and translational models of kidney disease, and we are developing artificial intelligence-based algorithms to take advantage of the knowledge obtained by single cell technology in the interpretation of kidney biopsies (Dr. A. Rinaldi).

Gastroenterology and Hepatology

(Prof. Dr. A. De Gottardi)

We developed novel applications of artificial intelligence to improve the accuracy of colonoscopy for the detection of benign polyps and colorectal cancer lesions (Dr. G. Lollo). The data obtained in two international clinical trials will contribute to set novel standards of care in colorectal cancer

screening. Predictive models in patients with advanced chronic liver disease are being developed to identify trajectories of movement, temperature, voice and cognitive functions, as assessed by using wearable devices, to detect behavioral changes that predict and prevent the occurrence of complications (Dr. A. Galante, Dr. D. Pizzagalli). Moreover, we are including patients in two international randomized controlled trials testing the efficacy of new molecules to treat portal hypertension.

Pneumology

(Dr. M. Bernasconi)

Night-time sleep disordered breathing linked to chronic kidney disease, and non-invasive high-frequency ventilation applied to radiotherapy and radiological diagnostics are among the main research interests of Prof. Dr. A. Ognà. The research of Prof. Dr. M.

Pons is focused on obstructive and central sleep apnea in patients with ischemic stroke. Research on dyspnea management and palliative care for patients with advanced chronic obstructive pulmonary disease is ongoing in collaboration with the Palliative and Supportive Care Clinic IOSI-EOC, CHUV and HUG (PD Dr. T. Fusi-Schmidhauser). Moreover, EOC contributes to the Swiss Severe Asthma Registry (Dr. P. Gianella).

Endocrinology

(Prof. Dr. P. Trimboli)

The main research area is related to thyroid disorders, including studies on the performance of different levothyroxine formulations to replace hypothyroidism, the diagnosis of cancer by biopsy and ultrasound, the treatment of benign nodules by radiofrequency, thyroid inconveniences induced by COVID-19 and thyroid implications of gluten-free diet. We recently started a new project about the application of wearable devices in endocrine and metabolic disorders, including severe hypoparathyroidism, gestational diabetes and thyroid diseases.

Dermatology

(PD Dr. H. Beltraminelli)

PD Dr. C. Mangas is mostly active in the field of melanoma and skin cancer. In collaboration with research unit from IOSI, she is a first investigator in three international and multicentric clinical studies in adjuvant, neoadjuvant and metastatic setting. Moreover, we have started a study to analyze the impact of dermatological toxicity in cancer patients. PD Dr. H. Beltraminelli participated to several national and international studies (melanocytic lesions, Martorell-ulceration, artificial intelligence, education) and is involved in some humanitarian projects,

especially, he coordinates a project aiming at the development of dermatopathology in Sub-Saharan Africa. Dr. I. Terrani is following a master's thesis on isotretinoin therapy.

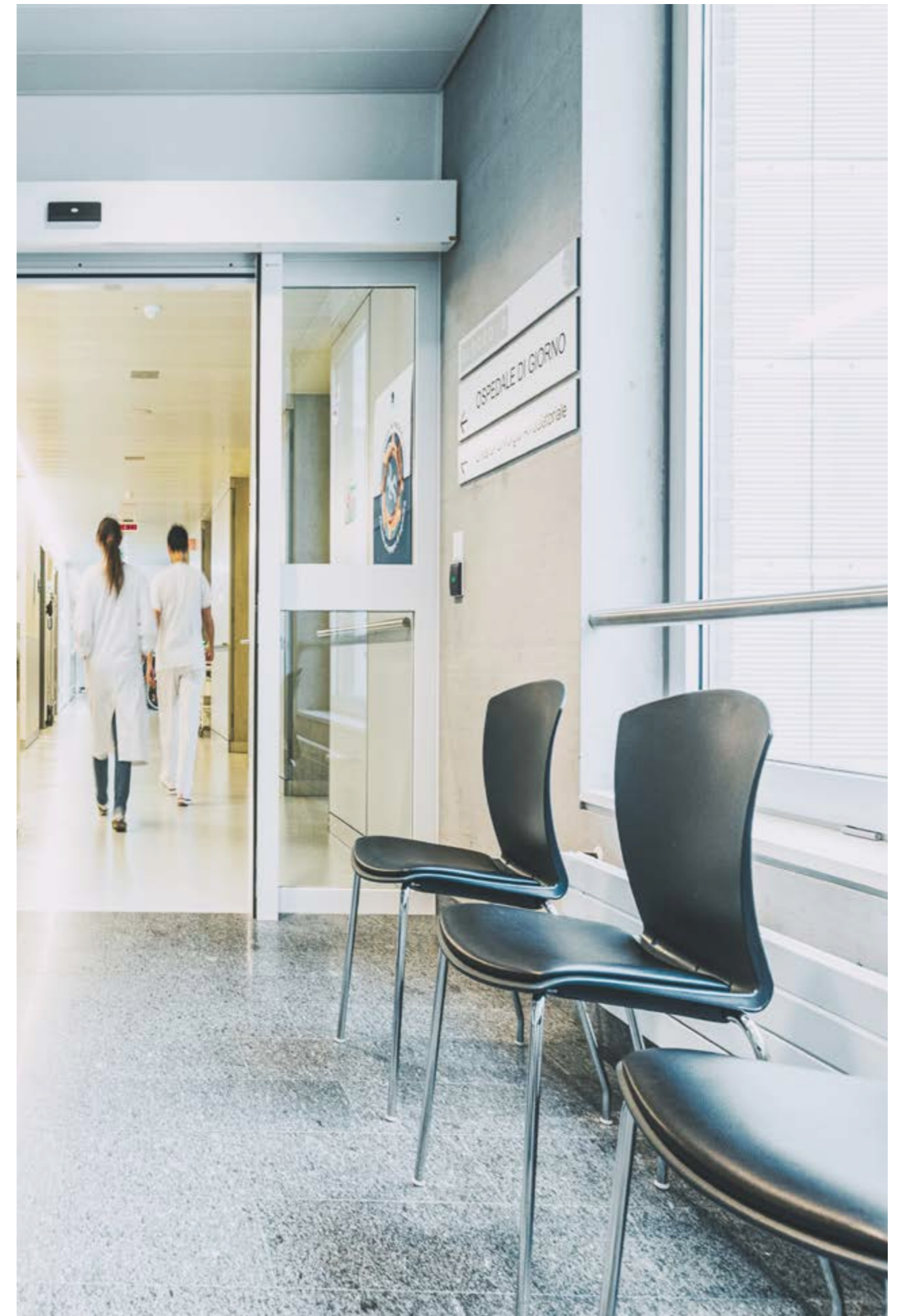
Clinical Nutrition

(Dr. N. Ossola)

Dr. N. Ossola and Dr. M. Quarenghi are involved in the e-swissHPN project (eHealth Care for Patients on Home Parenteral Nutrition), in collaboration with the University of Bern. The goal of the project is to evaluate the quality and effectiveness of videoconference consultations with patients on home parenteral nutrition. Other projects in collaboration with other research groups of the EOC are focused on the prehabilitation for patient undergoing major abdominal surgery, the intestinal absorption of drugs after bariatric surgery and the impact of a Mediterranean/low lipid diet combined with a physical activity program on prostate cancer.

Laboratory Research

[See chapter III on the Laboratories for Translational Research.](#)



Department of Surgery EOC

Prof. Pietro Majno-Hurst, MD
Head of Department

Visceral Surgery

In 2022 we pursued our clinical research activity with our own clinical trials, by participating in international trials and international registries, and with collaborative projects with our lab (see also scientific report of LRT-EOC). We gave an important number of presentations in international meetings and received prizes for high quality video-publications on surgical techniques.

Colorectal surgery

(Prof. Dr. D. Christoforidis, Dr. S. Popeskou, PD Dr. F. Mongelli et al.)

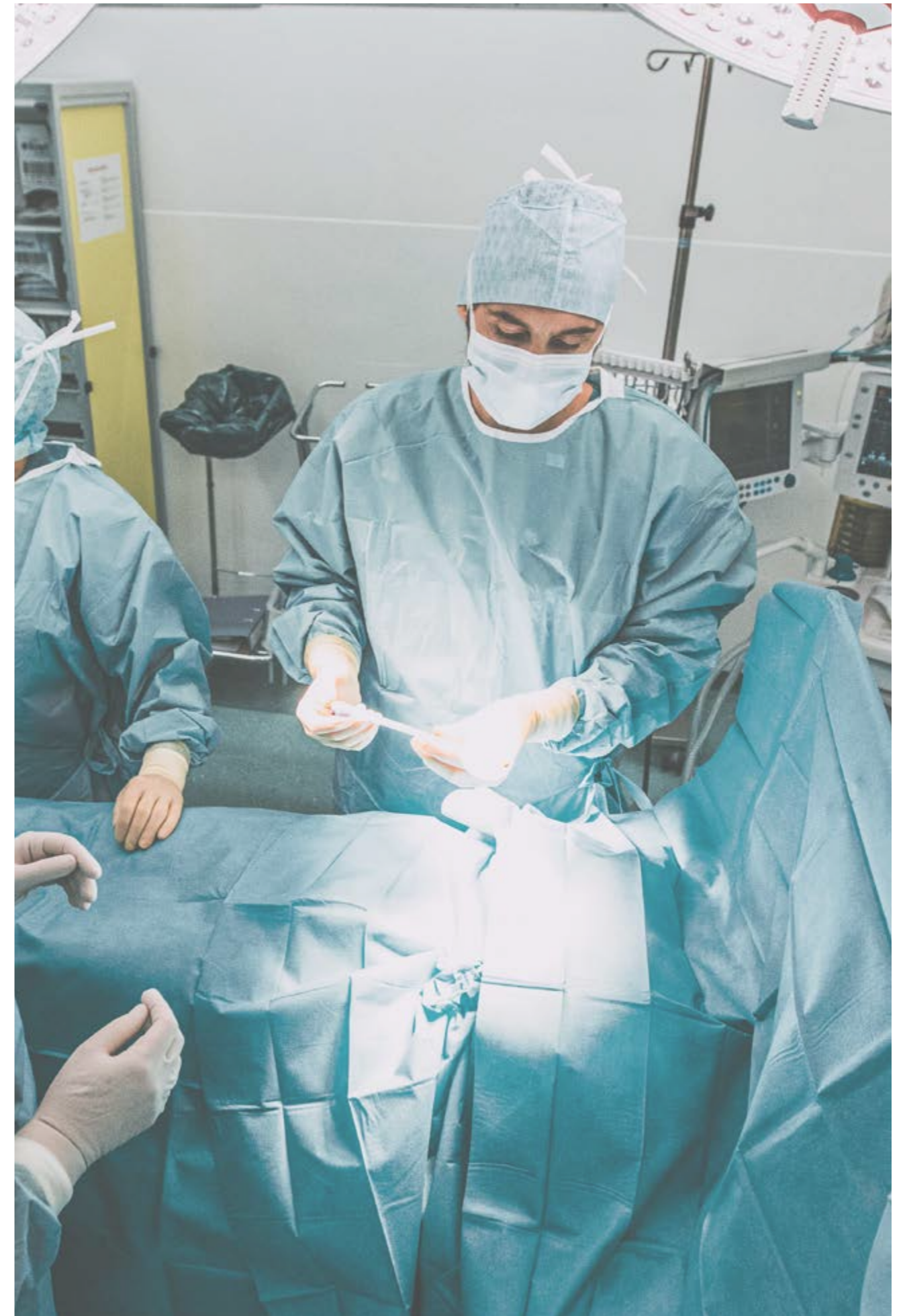
Along the lines of the past years, our research projects focused mainly on rectal cancer, and outcomes in minimal invasive surgery. In translational research, the MIBRECA study, looking at correlations between rectal cancer associated microbiome and response to pre-operative chemoradiotherapy has been recently concluded with very interesting results. the MicrobAL study, looking at correlations between the microbiome and anastomotic leakage after left colorectal resection both retrospectively in operated patients and in a mouse model is in full action. In clinical research, several projects are running: the RCT comparing percutaneous vs. laparoscopic transversus abdominis plane (TAP) block to improve post-operative pain control after colectomy has been completed; the study on outcomes after laparoscopic colorectal surgery in octogenarians has been almost concluded, just like a study looking at a 10-year follow-up of simple

diverticulitis and predictive factors of adverse outcomes. Both these latter projects are subjects of a Master thesis in Medicine. Finally, we took the national lead to participate and coordinate the APOLLO trial (Acute PresentatiOn of CoLorectal Cancer: an internatiOnal snapshot) on behalf of the Euro SURG Collaborative, that will run during 2023.

Bariatric surgery

(Dr. M. Marengo, Dr. F. Garofalo, Dr. F. Volonté, PD Dr. F. Mongelli)

Research topics in 2022 focused on revisional and conversional bariatric surgery, post-operative pathway optimization, surgical and anesthesia techniques, and multidisciplinary assessment. Currently, several studies are running, and we collaborate with other bariatric centers in Switzerland. A double-blind RCT assessing the efficacy of transversus abdominis plane block was successfully concluded and showed noteworthy results. We are enrolling patients for the SLIM Trial, a Swiss RCT investigating the effectiveness of long and short alimentary limb in Roux-en-Y gastric bypass. We concluded a study investigating the role of indocyanine-green fluorescence in patients undergoing revisional and conversional surgery and another study assessing the effect of primary bariatric surgery in patients with hypothyroidism was recently completed. Finally, a study regarding the long-term effect of gastric pouch resizing and distalization is ongoing.



The “research effort” produced the publication of several articles in some of the most quoted journals in bariatric surgery worldwide.

Robotic visceral surgery

(PD Dr. D. La Regina, Dr. R. Pini, PD Dr. F. Mongelli)

Several studies on surgical techniques and multimodal anesthesia methods are running in the innovative field of robotic visceral surgery. We concluded a randomized trial investigating locoregional anesthetic blocks (erector spine vs. transversus abdominis block) in patients undergoing robotic-assisted transabdominal preperitoneal inguinal hernia repair. Further studies either prospective or retrospective regarding surgical techniques, postoperative surgical pathways, economic assessments, and long-term incisional hernia incidence in robotic visceral surgery are running.

Hepato-pancreatico-biliary surgery

(Prof Dr. P. Majno, Dr. A. Cristaudi, Dr. R. Roesel, Dr. L. Bernardi et al)

We focused our research on surgical techniques implementing minimally-invasive approaches and improving surgical outcome with some on-going and some new projects. Our project on the impact of patient’s biography on caregivers and patient satisfaction is enlarging the horizon also to a qualitative analysis which is on-going. The Bi-Le trial, a multicentric Swiss trial that examines the impact of cholangiography using a white lipid solution to control biliostasis after liver resection, on the rate of post-operative biliary fistula is proceeding successfully. We started a multicentric trial (NutriWhi Trial) on post-operative immediate enteral nutrition in pancreatic head resections, which often put a challenge for managing post-operative gastroparesis. We obtained the best surgical procedure awards for minimally invasive pancreatic resection at the meeting for validation of European guideline for this kind of surgery. An interesting and accurate review on the impact of body nutritional status indicators as sarcopenia on hepatic oncologic surgery has been published on a high Impact Factor journal. Finally, we studied the learning curves of two minimally invasive approaches

(laparoscopic and robotic) in collaboration with the University of Pisa. We published a technical note on continuous suture who obtained a high impact award (Altmetric) in the British Journal of Surgery.

Vascular Surgery and Angiology

(Dr. L. Giovannacci, PD Dr J. Van den Berg, Dr. A. Ruffino, Dr. G. Prouse, Dr. A. Robaldo, PD Dr. H. Stricker, Dr. R. Canevascini et al.)

In the field of basic vascular research our main topic has been the introduction of a novel computational hemodynamic model that takes vessel compliance into account, testing its difference compared to traditional rigid vessel wall models. Furthermore, through the collaboration with the Swiss Stem Cell Foundation, we investigated and described the preparation of peripheral blood mononuclear cells to achieve a good manufacturing practice grade clinical product, ready to be used in patients with critical limb ischemia. We also evaluated the production in hypoxic conditions to increase functional activity and angiogenic potential. In the field of vascular clinical research there has been a special focus on recently developed minimally invasive techniques. We participated as one of the leading centers in testing the safety and effectiveness of aspiration thrombectomy systems in patients with acute lower limb ischemia. We also participated in a pilot study introducing the use of temporary stents in the treatment of infrapopliteal arterial disease (DEEPER pilot study). Regarding open surgery, we started a retrospective study to evaluate the mid and long term outcome after treatment of juxtarenal aortic aneurysms requiring supra-renal clamping. We were also involved in the development of the 2023 practice guidelines of the European Society for Vascular Surgery for the management of carotid and vertebral arterial disease.

Plastic, Reconstructive & Aesthetic surgery

(Prof. Dr. Y. Harder, PD Dr. R. Schweizer et al.)

Projects in clinical research: our department continues its efforts in clinical research, assessing efficacy and outcome of different reconstructive procedures after

breast cancer surgery, in different ongoing collaborative projects: the study assessing advantages and disadvantages of pre- and subpectoral implant-based breast reconstruction in a randomized multicentric study funded by SNSF/SAKK (collaboration with Prof. Dr. W. Weber, University Hospital Basel); the multicentric observational cohort study on immediate breast reconstruction following mastectomy (IRMA: collaboration with Dr. B. Lipp-von Wattenwyl; CSSI, EOC); and the rate of ischemia-associated delayed wound healing or skin necrosis after repetitive local heat preconditioning of the skin in reduction mammoplasty (collaboration with Prof. Dr. H.G. Machens, Munich). Other than that, we finished data collection for the study assessing MR-compatibility and user friendliness of a newly launched breast tissue expander for reconstructive surgery after mastectomy. In a further collaboration with IIMSI (Dr. S. Schiaffino, CSSI, EOC), we elucidate the benefit of autologous fat grafting in hybrid breast reconstruction (i.e., using autologous fat grafting and implants) by radiological and clinical means. Moreover, patient satisfaction after oncoplastic breast surgery (OPBS) in correlation with tumour volume and breast size is investigated using a 3D scanner to test the hypothesis, that overall patient satisfaction is higher after receiving oncoplastic breast surgery (i.e. intraglandular breast tissue rearrangement) after tumorectomy in combination with adjuvant radiation therapy when compared to simple tumorectomy and irradiation. Furthermore, we are investigating the benefit of using local tranexamic acid in different standard plastic surgery procedures, such as reduction mammoplasties, abdominoplasties but also in ablative surgery, i.e. mastectomies, in order to reduce draining times of surgical wounds and eventually hospital stay. In a more “historical and political” project, we sent a nation-wide survey asking the different hospitals and private clinics about their microvascular activity and its evolution in the last two decades, which disclosed a comprehensive offer of microvascular reconstructions spread over the whole country.

Regarding the topic of facial palsy, PD Dr. Schweizer is the local principal investigator

at the EOC in a collaborative project with the renowned Laboratory for Bionic Extremity Reconstruction at the University of Vienna (Prof. Dr. O. Aszmann) and the Department of Plastic Surgery at Cantonal Hospital Aarau (PD Dr. H. Klein). The study aim is to gather detailed qualitative and quantitative information on both nerve and muscle transplants at the time of and after dynamic facial nerve reconstruction, and their histopathological changes over time, with the aim of a better understanding on how the outcome may be improved in this relatively rare entity.

Projects in basic research: Dr. med. E. Limido is completing a research fellowship in Germany during a MD-PhD at the Institute for Clinical & Experimental Surgery of the Saarland University, Homburg/Saar in Germany (Prof. Dr. M.W. Laschke) and do collaborate on projects, assessing the effect of autologous nanofat grafting on healing of complicated wounds and/or irradiated soft tissues in a rodent model. PD Dr. med. R. Schweizer is collaborating with the department of surgical research of the University Hospital Zurich on a project evaluating an in vivo absorbable tissue engineered vascular graft, seeded with induced pluripotent stem cells in a rodent model.

Orthopedics and Traumatology

(Prof. Dr. C. Candrian, Prof. Dr. G. Filardo, PD Dr. M. Delcogliano et al.)

The Ortho-Trauma Clinical Research Team, founded in 2017, is now a well-established reality in the EOC research panorama. The aim of the clinical research team of the Orthopaedic and Traumatology Service is to improve the current knowledge as well as the treatment strategies in the field of orthopaedics and traumatology. In this light, systematic reviews and meta-analyses are performed to synthesise the existing literature, and innovative clinical trials are planned to increase the strength of the current knowledge and explore new promising fields. Moreover, the team takes part in national and international multi-centre studies that involve some of the most important research centres in Europe.



[Click here](#) to see the complete list of publications of the Department of Surgery EOC

The research group is also conceived as an opportunity offered to all residents of the Orthopaedics and Traumatology Service of all the EOC Hospitals, to move their first steps or deepen their knowledge in the field of orthopaedic research.

Urology

(Prof. Dr. A. Gallina, Prof. Dr. N. Fossati)

The Urology service is actively involved in the field of uro-oncology clinical research. In particular, it contributes to an ongoing clinical trial including patients affected by prostate cancer. It is a prospective trial evaluating the diagnostic accuracy of a novel tracer for pre-operative staging with PET/CT scan in men affected by high-risk prostate cancer and candidate for radical prostatectomy plus pelvic lymph node dissection. Furthermore, the Urology service significantly contributes to ambitious translational projects, aimed at identifying novel biomarkers and microbiome signatures for patient outcomes in newly diagnosed prostate cancer. Moreover, the Urology service has become part of two important multi-institutional collaborations investigating the optimal management of patients with prostate cancer recurrence. In 2022, several abstracts from these collaborations have been accepted at the two most important urological international meetings (EAU and AUA), as well as the national Swiss urological congress. Finally, the Urology service renewed the functional service, including the neuro-urology clinic, which will eventually translate in new potential scientific projects regarding chronic pelvic pain and pudendal neuropathy, in collaboration with INSI EOC.

Thoracic surgery

(Prof. Dr. S. Cafarotti, PD Dr. M. Patella et al.)

In 2022 the Department of Thoracic Surgery continued to follow both the international and national pathways of scientific research. We finalized the publication of an important work that came from the joint efforts of several partners in Europe and Canada, with a strong and important message on the multimodality management of locally advanced lung cancer. The study entitled “A risk model to predict the delivery of adjuvant chemotherapy following lung resection in

patients with pathologically positive lymph nodes” was published in Seminars in Thoracic and Cardiovascular Surgery and it received the best clinical publication award in 2022 from the Swiss Thoracic Society (SGT). We also continued our collaboration with the University of Zurich and with several other partners in Switzerland, participating in studies on blood transfusion policy and other non-oncological diseases. Looking at our region, we wanted to investigate and verify in our local reality, the applicability of different dogmas the literature suggests. Our attention focused on the quality offered by our health care system in lung cancer treatment and how it can be improved.

Laboratory Research

[See section chapter III on the Laboratories for Translational Research.](#)

Department of Gynecology and Obstetrics EOC

Prof. Andrea Papadia, MD, PhD
Head of Department

The Department of Gynecology and Obstetrics EOC is part of an international scientific network involving other Swiss and European Universities and Hospitals. Since its foundation in 2019, the Department of Gynecology and Obstetrics has progressively implemented its scientific profile. In few years the output of peer-reviewed original manuscripts has increased dramatically. In 2022 several major prospective studies in the field of gynecology, senology and obstetrics were active or launched at the Department of Gynecology and Obstetrics. Our aim is to allocate every patient into a research database.

Members of the Department of Gynecology and Obstetrics were involved in the following prospective multicentric international trials as national coordinators:

- “Cascade trial” (CArdiovaSCuAr outcomes after major abDominal surgEry):

a cohort study, aiming to characterize rates of post-operative cardiovascular complications following major abdominal surgeries

- “CovidSurg trial”: a cohort study, aiming to assess the outcomes of surgery in patients diagnosed with COVID-19
- “AXSANA trial” (AXillary Surgery After NeoAdjuvant Treatment): A prospective multicenter cohort study to evaluate different surgical methods of axillary staging (sentinel lymph node biopsy, targeted axillary dissection, axillary dissection) in clinically node-positive breast cancer patients undergoing neoadjuvant chemotherapy
- “Melody trial” (MEthods for LOcalization of Different types of breast lesions (EUBREAST 4)): A prospective multicenter cohort study to evaluate different imaging guided methods for localization of non palpable malignant breast lesions

For the following prospective trials members of the Department of Gynecology and Obstetrics joined as local PIs:

- “Go Soar trial”: A prospective multicenter cohort study to evaluate international variation in 30-day post-operative morbidity and mortality following gynecological oncology surgery between very high/high and medium/low human development index country settings
- “Surgery for deep infiltrating endometriosis, symptoms and outcomes”: a multicenter analysis of the surgical outcomes in certified endometriosis centers in Europe
- “A long term prospective Post Market Clinical Follow-up (PMCF) of the GYNECARE TVT ABBREVO® Continence System for the treatment of Stress Urinary Incontinence”
- “Covipreg”: a Registry for pregnant women exposed to SARS-CoV-2 and future emergent pathogens
- “MoCHIV” (Swiss HIV Cohort study): a Swiss cohort study on patients affected by HIV
- “B-Red trial”: a prospective clinical randomized trial assessing the safety and efficacy of a new surgical tool to reduce uterine bleeding at the time of

- hysterotomy during Cesarean section
- “I-Prepare”: International Prospective REgistry on Pre-pectorAl breast REconstruction
- “IRMA”: A multicenter, observational cohort study of women undergoing Immediate breast Reconstruction following Mastectomy

Furthermore, the Department of Gynecology and Obstetrics, IOSI and IOR are currently collaborating on two translational research lines focused on the immune landscape in breast and ovarian cancer patients. For this purpose, cord blood from 190 patients who delivered at the Regional Hospital of Lugano was collected and processed at the Cancer Immunotherapy lab of the IOR. The recruitment is under voluntary donation and it is currently offered to every patient who delivers at the Regional Hospital of Lugano.

Finally, multiple retrospective studies are continuously produced either as single center or in collaboration with some of our research network partners.



Department of Anesthesiology, Emergency and Intensive Care Medicine EOC

Prof. Paolo Merlani, MD
Head of Department

The Department of Anesthesiology, Emergency and Intensive Care Medicine of Ente Ospedaliero Cantonale (DAEICM-EOC) is a functional Department. Each of the four public hospitals of the EOC has his own DAEICM. The four local DAEICMs make up the DAEICM-EOC. Each local DAEICM is headed by a local Director. The DAEICM-EOC coordinates and facilitates the integration and the collaboration among the four local DAEICMs for patients' care and for research purposes. The DAEICM-EOC is responsible for the didactic module "Critical Care Medicine" for the Medical Master School (third semester) of the Università della Svizzera italiana (USI). Furthermore, it supports students for their master's and/or doctoral thesis. In 2022 a Local Research Unit (LRU) for the entire DAEICM-EOC has been established and it is headed by an attending physician of the Department. The main aim of the LRU is to support any researcher/investigator from study conception to manuscript submission in peer review journals. Anesthesiology, Emergency Medicine and Intensive Care Medicine manage their own research projects as both monocentric and multicentric trials in collaboration with other Swiss and foreign research centers. One of the purposes of the LRU of the

DAEICM-EOC is to encourage common projects among the different Services of the Department and especially regarding the transition of care (interdisciplinary and interprofessional projects). Quality-improvements studies and especially process-improvements trials are still scarce in medicine and especially in the acute care setting.

Different topics are currently investigated by the DAEICM-EOC, and thereafter we mention the most relevant:

- Diagnostic accuracy of Point of Care Ultrasound in different clinical settings, especially in Anesthesiology and Intensive Care Medicine
- Cost-effectiveness analysis in Anesthesiology
- Perioperative transfusion's strategies
- Use of different analgetic regimen and analgetic procedures in the post-operative setting
- Treatment strategies for chronic pain syndromes
- Impact of the COVID-19 pandemic on the occurrence of different complications (herpes simplex recurrence) in the ICU population
- Best practice for the care of mild traumatic brain injury in the Emergency Department (ED)
- Communication practices in the ED for outpatients



Different projects are working in progress. Here a selection of the most promising: optimization of the time to transfer for patients from the ED to the ICU and identification of barriers; use of artificial intelligence and machine learning techniques for the early identification of nosocomial infections; strategies to improve communication's skills of ED doctors; impact of delayed transfer of patients from the ward to the ICU.

Other EOC Services



Rehabilitation Clinic EOC (CREOC)

Nicola Schiavone, MD
Medical Director

Rehabilitation Clinic of Ente Ospedaliero Cantonale (CREOC) deals with rehabilitation charge of patients who need to improve their quality of life. In order to achieve the goal, it is used the biopsychosocial model, trying to recover biological factors (genetic, biochemical, ecc.), psychological factors (personality, behaviour, ecc.) and social factors (cultural, familiar, socioeconomics, ecc.) of every single patient. Rehabilitation charge try to reduce pain, improve movement and strength of upper and lower limbs, improve the respiratory capacity and global reconditioning of each patient. CREOC, together with Clinica Hildebrand, belongs to REHA TICINO, the organization that has the cantonal inpatient rehabilitation mandate. Moreover, CREOC offers rehabilitation to every inpatient in all over EOC Hospitals and takes care of outpatients who need rehabilitation.

Inpatient rehabilitation is located in Faido and Novaggio, where the Canton has provided the skeletal muscle mandate, internal medicine – oncology mandate and pneumological mandate. Moreover, for taking care of elderly patients, CREOC applies a geriatric patient care concept. The taking charge of inpatient is based on rehabilitation diagnostic therapeutics pathways (PDTA) that have a focus on the main illness.

Inside CREOC there are groups of study and work that coordinate standardized working procedures, related to evidence-based medicine. The groups are dealing either with rehabilitation diagnostic therapeutics pathways or specific fields as postural rehabilitation, pelvic floor rehabilitation, respiratory rehabilitation and lately rehabilitation after COVID-19 illness.

About research activities, in 2022 CREOC collaborators presented at local congresses an original and effective work, promoted by Swiss Patients Safety Foundation (FSPS), that tries to teach to the nursing team how to deal with risk management, errors and pharmacological process. Concerning with taking care of SARS-CoV-2 affected patients, CREOC collaborators concluded and published a study on the evaluation of the predictive factors of the short-term effects of a multidisciplinary rehabilitation in COVID-19 survivors, developed together with Clinica Hildebrand of Brissago. The goal of this study was to evaluate the short-term effects of a multidisciplinary rehabilitation program, applied to patients affected by COVID -19, after discharge from intensive care and admission to rehabilitation Clinics. Moreover, the study evaluated which are the factors that influence the rehabilitation program.



Nursing Research Centre EOC

Annette Biegger, MSc

Head of EOC Nursing Division

In recent years, the EOC Nursing Department has promoted organisational and cultural improvements in order to advance the quality of care. This has led to changes concerning the organisation and philosophy of care, as well as the electronic documentation of nursing activities. In addition, great attention has been given to the advancement of nurses' knowledge and skills, which resulted in the implementation of the Clinical Nurse Specialist roles. These professional figures are crucial for the development of advanced nursing skills and for the clinical governance of professional activities.

Furthermore, given the increasing complexity of patients and health systems, new nursing roles are emerging at the EOC, such as the advance practice nurse. To support this process, investment has also been made in people with specific research training (PhD) and dedicated offices.

In the medical field, knowledge evolves rapidly: something that was effective yesterday may no longer be effective today. This concept underlines the importance of developing research studies focused on the nursing field, to provide nurses with new evidence and develop new expertise. Polit & Beck (2006) define nursing research as a "systematic inquiry designed to develop knowledge about issues of importance to nurses, including nursing practice, nursing education, and nursing administration".

Nursing research activities, with dedicated resources, were first registered at IOSI in 2010 and are annually documented in the EOC Annual Scientific Report and EOC Nursing Activity Annual Report. In September 2016, the Centre of Competence for Nursing Research (CcRI) was established within the EOC Nursing Division with the following mission: "...to promote the development and quality of nursing care in response to the health needs of the population, promoting innovation and nursing research in all clinical realities of the EOC and maintaining a close link between research (theory) and nursing care (practice)". In the coming years, the plan is to expand the network of colleagues and/or nursing research offices within the EOC sites.

Today, a research office and/or nurse are active across different sites, including IOSI, IPSI, Ospedale Regionale di Bellinzona e Valli, ICCT and Ospedale Regionale di Lugano. At the moment, in each of this hospital/institute there is at least a nurse dedicated to nursing research.

Currently, the Leads of the local nursing research units are: Angela Tolotti at IOSI, Sarah Jayne Liptrott at the Ospedale Regionale di Bellinzona e Valli, Elena Corina Luca at the Ospedale Regionale di Lugano, Colette Balice-Bourgois at the IPSI and Michele Villa at the ICCT. Laura Moser worked as data manager for the research team at the IOSI until December 2022.

The Chief of the Centre of Competence for Nursing Research (CcRI) is Loris Bonetti, in post since September 2021. Laura Steiner also works as research nurse at the CcRI.

Nurses active in research are part of the Nurse Led Research Coordinating Team (CooRI) team, which coordinates EOC research activities and research development plans in the nursing field.

The CooRI has recently been formally recognized as Local Research Unit. The following are the main activities the research network is responsible for:

- To develop training courses on research topics to be offered to all EOC nurses, but especially to clinical nurse specialists. This is critical to create essential synergies with clinical realities and the network of clinical nurse specialist within EOC.
- To promote and support the cultural and scientific production by colleagues (preparation of abstracts, posters, journal articles, research projects, etc.).
- To collaborate with colleagues with managerial roles and colleagues from clinical governance for the collection and/or critical analysis of data, the dissemination of evidence and the development of internal guidelines.
- To develop and implement nursing research projects, using either quantitative, qualitative or mixed methods.

- To develop and maintain collaborations with national and international nursing research associations and networks.

Research questions usually arise from needs identified by nurses working in clinical practice or holding managerial roles and are answered through studies, which can be carried out with internal or external resources and funding. Internal resources are mainly used to conduct 'action-research' studies-planned and developed together with nurses involved in clinical practice and in the context of interest, aimed to explore a specific clinical or professional phenomenon in order to change and improve nursing practice. Such studies are 'context-related' and therefore results can be transferred to similar realities, but usually cannot be generalized. Despite this, such action-research studies are deemed important as they require a proactive role from professional teams and stimulate reflective thinking and critical review of their professional practice. Other studies conducted, especially those externally funded, tend to focus on phenomena of clinical or professional interest of a more general nature.





The following are the research studies currently active, funded by EOC or other funding bodies:

- A mixed-methods study (NuBE Project), in collaboration with the Catholic University of Milan, which aims to identify which relational and communicative behaviours of nurses in oncology promote patients' engagement, in order to develop a reliable psychometric tool to identify such behaviours.
- A pilot project to test a complex intervention delivered by oncology nurses aimed at promoting a healthy lifestyle among cancer patients. The intervention is based on patient engagement theory. The study will be conducted in partnership with the Catholic University of Milan.
- Three studies about the implementation of Advanced Practice Nurse (APN) role respectively with Stroke patients, patients undergoing peritoneal dialysis and in palliative care.
- An observational study on the introduction of sonography in nursing practice in the Urology service of the Ospedale Regionale di Bellinzona e Valli, with the aim of developing professional skills and improving nursing care (appropriateness of bladder catheterisation).
- An investigation of a Mindfulness Based Stress Reduction program was held in the Ospedale Regionale di Bellinzona e Valli, open to participants from all professional categories. The study explores the effect of the program on mindfulness practices, stress and psychological well-being.
- An implementation study on the development of a complex intervention to promote positive workplaces.
- A Delphi study on the adaptation of the Ethical Conflict Nursing Questionnaire-Critical care to acute and chronic care contexts.
- A non-inferiority randomized controlled trial, open and controlled in parallel group on safety and benefits of mobilization at 4 hours compared to 24 hours after implantation of an intracardiac electronic device.
- An observational study on the frequency and predictors of haemodynamic instability during automated changeover of norepinephrine infusion pumps.

- A prospective observational study about active involvement of patients with cancer in clinical research, with the aim to educate patients in order to be directly involved in the performance/discussion/design of clinical studies.

Other studies are being planned in collaboration with colleagues working in the various clinical settings.

In October 2022 the second EOC nursing research congress took place in Lugano, in collaboration with the University of Applied Sciences and Arts of Southern Switzerland (SUPSI). In this edition 40 abstracts from EOC nurses were presented, meaning that nursing research is evident within the EOC. Furthermore, 9 abstracts presented at the conference were from the Bachelor and Master of Science in Nursing theses of SUPSI students. Prof. Sabina DeGeest, a renowned researcher and Chief of the Nursing Faculty at the University of Basel, participated in the congress as keynote speaker, with an enlightening presentation about implementation science. The event was very successful and appreciated by the EOC nurses, with the third edition already planned for spring 2024.

It is also important to underline the great participation of many EOC nurses at the 11th Research day in Ticino (in June 2022), in which a specific parallel section was dedicated to nursing research projects. One of the nursing projects was also presented during the plenary session and two projects were awarded best nursing oral presentations.

Moreover, in February 2022, the collaboration between the EOC Competence Centre for Nursing Research and the SUPSI Competence Centre for Health Practices and Policies was officially signed. This collaboration between clinical and academic fields will be strategic for the development of nursing research and the nursing profession at the EOC and in the Ticino context.

Information and Communications Technology Division (ICT) EOC

Marco Bosetti

Head of EOC Information and Communications Technology Division

The “Information and Communications Technology area” (ICT) is the IT department of EOC. With more than 90 employees, it covers the traditional IT services like infrastructure, operations and support. Moreover, it implemented in-house and maintains the Geco that is the clinical information system of EOC since 2007. For this reason, the ICT has extensive experience in integration and analysis of clinical records.

The ICT is actively involved in research activities mainly through his new dedicated team named Datascience and Research that provides both technical and consultant expertise. Besides internal resources, the ICT strategic development foresees the establishment of collaborations with external organizations. The collaboration with the Swiss AI Lab (IDSIA) aiming at applying cutting edge artificial intelligence techniques on EOC data was established in 2019. Since 2019 the ICT is part of the Swiss Personalized Health Network to develop IT tools in the field of personalized medicine.

The ICT is the central point for extraction and analysis of clinical data supporting EOC researcher and Medical Master School students in teaching activities, scientific publications and writing of funding proposals

The pipeline of running financed projects includes the early identification of ventilator-associated pneumonia using AI (Innosuisse) and the prediction of clinical deterioration in oncological patients (SAKK/AbbVie award).



The ICT is also involved in internal projects aiming at enhance the hospital management like the implementation of an automatic shift planner and the use of AI to manage&optimize hospitalizations supporting multiple use cases (LOS, AKI, PBM,...).

Additionally, the ICT scientific production growth substantially during 2022 thanks to fruitful collaborations with clinical partners. Finally, the suite of research software/ infrastructure solutions developed and

maintained by ICT includes an electronic case report forms to manage clinical trials (5dBase - validated FDA cfr 21 part 11), an event driven direct data entry database to manage both biobanks and surveys (eddie - validated FDA cfr 21 part 11), a clinical data warehouse, tools to anonymize clinical narratives and a high throughput cluster for -Omics/RNA seq data analysis.

Laboratories for Translational Research (LRT) of EOC



Laboratories for Translational Research (LRT) of EOC

Prof. Andrea De Gottardi, MD PhD
Director

The Laboratories for Translational Research of the Ente Ospedaliero Cantonale are dedicated to applied biomedical research by a multidisciplinary approach and aim to improve health and to understand human disease.

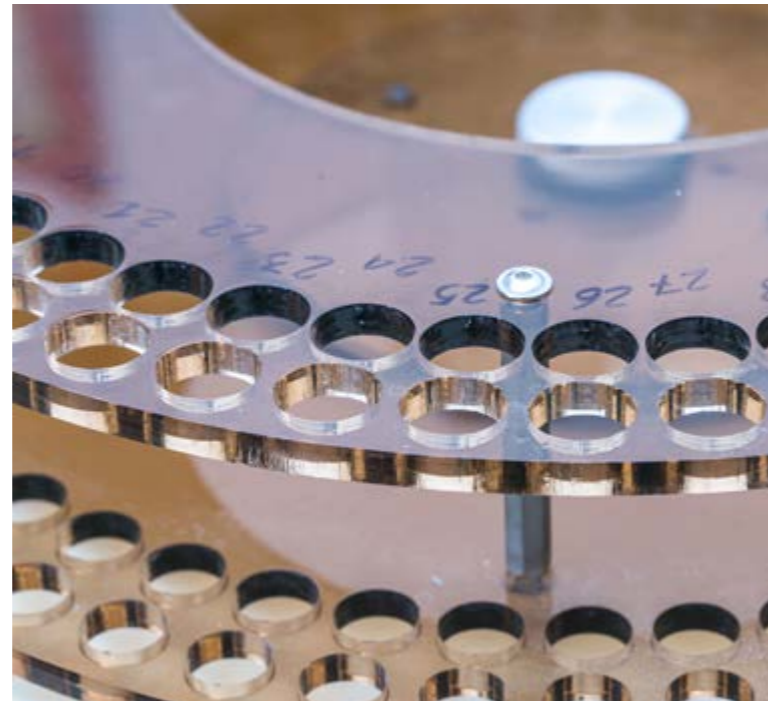
Our research teams are interdisciplinary and include biologists, biochemists, physicians, pharmacologists, technicians and students. We use cutting edge technologies ranging from biomolecules to cellular cultures and organoids, to organs on a chip from animal and human origin.

We are committed to work with animal models only when strictly unavoidable. The administrative team supports all research groups in the management of all administrative and financial tasks. In addition to our focus on translational research, we are dedicated to teaching and training young scientists to foster their career in basic and clinical research.

Open positions for master and doctoral thesis are available.

The Laboratories for Translational Research were born in 2019 from the initiative and support of EOC and are funded by private donations, by charitable organizations, by biotech companies and by foundations for scientific research. In October 2021 the laboratories moved to the new research building located in via Francesco Chiesa 5 in Bellinzona.

More information is available on the new homepage <https://lrteoc.ch/>.



Laboratory for Cardiovascular Theranostics

(Prof. L. Barile)

Role of senescence in cardiovascular diseases

In vitro model of senescence in human cardiomyocytes: this lab is active in studying the role of senescence in cardiovascular diseases. In particular, an in vitro model of premature senescence in human cardiomyocytes (functional cardiac cells) was developed using the Induced Pluripotent Stem Cells (iPS) technology. This model is a unique platform for the study of cellular mechanisms lying behind senescence-associated cardiac disease in human. The platform will be the basis of an important study aiming to evaluate the susceptibility of senescent cardiomyocytes to SARS-CoV-2.

In-vivo stress-induced senescence: the role of senescence is also studied in-vivo in animal model of myocardial infarction. The acute ischemic event induces the formation of fibrotic tissue and the accumulation of senescent cells that release factors that can alter the functionality of heart cells. This may contribute to exacerbate the necrosis process resulting from the acute ischemic event. We aim to studying possible "senolytic" and/or "senostatic" approaches to ameliorate senescence-associated damage in the heart.

Exosomes and Extracellular vesicles as Liquid Biopsy

EV as Biomarkers: recently, our lab has completed a series of studies performing molecular analysis of the biofluid-derived extracellular vesicles (EV). We profiled surface protein and membrane lipids of plasma and serum-derived EV. The EV-based diagnostic test standardized at LCT laboratory has been evaluated as diagnostic tool in acute cardiac rejection following heart transplantation to discriminate different types of rejection as well as for prognosis of severity of rejection. We recently evaluated the lipid composition of circulating EVs and their diagnostic potential after myocardial ischemia. Very recently we used the profiling of EV as prognostic tool for predicting severity in COVID-19 patients.

Role of EV as Mediators of Inflammation: the lab has made important advances in research studying the role of circulating EVs resulting from the inflammatory process following an acute ischemic event of the myocardium. We proved that inflammatory EV secreted by polarized macrophages carrying inflammatory cytokine such as IL-1 α , IL-1 β e have direct cytotoxic effects on heart cells.

Regenerative Medicine Technologies research group

(Prof. M. Moretti)

The Regenerative Medicine Technologies Lab adopts an interdisciplinary approach, at the intersection of engineering, biology and medicine, aiming at regenerating biological tissue substitutes in vitro. Major lab competences are represented by advanced technologies such as biofabrication and microfluidics, enabling tools for the generation of innovative constructs replicating the complexity of in vivo biology. In more detail, our lab generates 3D tissues based on cell co-cultures mainly of human origin, embedded in suitable biomaterials and cultured in engineered environments providing biophysical and mechanical stimuli.

Our lab mainly focuses on musculoskeletal tissues, producing biological replicates with a scale ranging from microfluidic devices to macro-scale constructs of clinically relevant dimensions. The microarchitecture of physiological tissues can be hierarchically replicated, spanning from mimicking the interface between different tissues (e.g. tissues composing the joints or the tendon bony insertion) down to the reproduction of microscale tissue features (e.g. bone trabeculae or microvascular networks). From a translational point of view, smaller scale devices are currently being improved towards high throughput platforms usable for drug screening purposes and testing of personalized therapies, whilst bigger scale constructs could be translated to biological substitutes usable in clinics for the regeneration of diseased tissues. In particular, we are working on reproducing bone, cartilage and skeletal muscle constructs, to improve drug discovery and to personalize available therapies in fields such as bone tumors, osteoarthritis and muscle fibrosis. In the field of bone tumors, we demonstrated that our models were able to reproduce the effects of known anti-tumor drugs on cancer cells better than simple 3D cancer cell models. We are also working on improving the mechanical properties of the support matrix through bioprinting techniques, and on increasing

the throughput of our system to enable drug screening with a higher throughput.

About skeletal muscle, we are studying models able to replicate the main features of fibrotic muscle tissue, to provide advanced tools to researchers searching for new therapies. The currently generated models have shown that they can replicate the organization of muscle fibers, the presence of a vascular network similar to that present in vivo, and better reproduce the altered production of extracellular proteins in pathological conditions as compared to the corresponding 2D models.

In our lab we also develop patient-specific multi-tissue 3D models, such as microfluidic models of an osteoarthritic joint, that allow to compare different possible therapeutic approaches for the single patient and can represent an innovative instrument for the development of personalized therapeutic regimens for osteoarthritis and other orthopedic pathologies.



Gastroenterology and Hepatology research group

(Prof. A. De Gottardi)

Our research interest is to better understand the complex role of the gut-liver axis in chronic liver disease such as fibrosis and cirrhosis associated with portal hypertension and fatty liver disease. Intestinal microbiota is confined to the intestinal lumen by the mucosal and vascular barriers. But in chronic liver disease, changes in the gut barrier and liver dysfunction cause pathological translocation of bacteria and their metabolites to the extra-intestinal organs, that mediate hepatic and systemic inflammatory responses promoting liver injury. The intestinal and mesenteric blood and lymphatic vessels are major communication routes between the gut and the liver. We seek to understand how the alterations of gut mucosa and vascular barriers may affect the progression of advanced chronic liver disease associated with portal hypertension.

We recently demonstrated that splanchnic angiogenesis and lymphangiogenesis are increased under the control of Paneth cells during the development of portal hypertension. Paneth cells are specialized

intestinal innate immune cells that secrete several bactericidal peptides with pro-angiogenic properties. Intestinal-mesenteric lymphatic vessels are the center of dietary lipid uptake and transport within the body, but their role is not known in fatty liver disease. Metabolic dysfunction-associated fatty liver disease (MAFLD) prevalence is increasing quickly around the world, but no effective drug or efficient treatment yet exists. We aim to explore the underlying mechanisms of MAFLD by investigating the role of Paneth cells in the regulation of intestinal-mesenteric lymphatic vessels and lipid metabolism in MAFLD. For this project, we are in collaboration with Prof. P. Jay at the University of Montpellier, France, and Prof. R. Wiest at University hospital of Berne.

For the in vivo investigations, we use different experimental models of chronic liver disease and portal hypertension, combined with in vitro studies of 3D intestinal organoids, angiogenic assays in specific conditions with or without Paneth cells along with cutting-edge molecular biology technologies.

Cellular and Molecular Cardiology research group

(Prof. G. Vassalli)

This research group has investigated the biological role and the therapeutic potential of secreted extracellular vesicles (EV), particularly of nanosized vesicles of endosomal origin called exosomes, in cardiovascular diseases, with a focus on acute myocardial infarction. Studies from the lab showed that EV secreted by human cardiac mesenchymal-derived progenitor cells were endowed with beneficial biological activities, as demonstrated in both in vitro and in vivo myocardial infarction models and chemotherapy-induced cardiotoxicity models. These properties included cytoprotective, pro-angiogenic, anti-fibrotic, and pro-proliferative effects in cardiomyocytes. In collaboration with the Department of Heart Surgery at Zurich University Hospital, we also demonstrated that such EV significantly reduced infarct scar in a preclinical model of acute myocardial infarction in pigs.

Furthermore, CMC is actively studying the role of miRNA carried by cardiomyocyte derived-extracellular vesicles in heart development. Particularly, we aim to understand the physiological signaling that unlock cardiomyocyte proliferation to the development of new therapeutic strategy in myocardial regeneration. This lab has a major expertise in extracellular vesicles (EV) isolation and characterization. In vitro and in vivo model for EV functional evaluation has been developed and optimized in the past years. In vitro EV are tested on cardiomyocyte (from neonatal mouse or human iPS-derived), cardiac fibroblast (mouse and human) and on a human heart organoid (iPS-derived cardiomyocyte and cardiac fibroblast). The use of 3D culture is a cutting edge technique that allow to perform experiments taking into account the synergistic effect of cardiac cells under different stimuli. In vivo acute myocardial infarct models were used for studying the effects of human cardiac mesenchymal-derived progenitor cells on ischemic cardiac injury. An in vivo model of chemotherapy-induced cardiotoxicity was also developed.

Surgical research group

(Prof. G. Iezzi)

The research activity of this group focuses on the interaction between gut microbiota and immune system in gastrointestinal diseases, primarily colorectal cancer (CRC), or in systemic diseases of surgical relevance, such as obesity.

Starting from the analysis of the gut microbiota and the immune contexture in human samples, this group aims at:

- identifying novel biomarkers predictive of long-term response to surgical or medical treatment,
- identifying patients at high risk of recurrence who may benefit of additional therapies,
- developing novel therapeutic concepts based on gut microbiota conditioning through administration of selected probiotics or targeted antibiotics.

The gut microbiota consists of trillions of microorganisms critically shaping immune system development and immune responsiveness, through their effects on host metabolism.

In pathological conditions associated with increased gut permeability, such as cancer or inflammatory diseases, defined species of the gut microbiota translocate from the lumen into the lamina propria of the intestinal mucosa and therefore directly interact with epithelial cells and infiltrating immune cells.

Recent studies of this group in CRC have demonstrated the capacity of bacterial components of the gut microbiota to induce, upon direct contact with tumor cells, expression of chemokines recruiting into the tumor bed immune cells predictive of favorable clinical outcome. Analysis of tumor-associated microbiota in human CRC samples has led to the identification of a group of bacteria positively associated with high immune cell infiltration and prolonged patients' survival. Immunomodulatory properties of identified bacteria and their capacity to promote antitumor effects have been investigated in in vitro and in vivo models. We identified one bacterial species capable of expanding cytotoxic T cells in vitro and to induce anti-tumor effects in

vivo, alone or in combination with immune checkpoint blockade-based immunotherapy. Furthermore, we identified an additional species, belonging to a different bacterial genus, capable to elicit antitumor functions in tumor-associated neutrophils. We plan to test effects of a combined administration of these two bacteria in the next future.

On the other hand, defined bacterial species enriched in CRC tissues have been found to mediate pro-tumorigenic effects. In particular, *Fusobacterium nucleatum*, is associated with a poor response to chemotherapy and unfavorable prognosis. In collaboration with IOSI, a clinical study has recently been initiated, to evaluate the effectiveness of pre-operative antibiotic therapy with metronidazole to reduce tumor colonization by *Fusobacterium nucleatum* in patients with CRC undergoing surgery. A more recent project aims at identifying bacterial species of the gut microbiota promoting healing or anastomotic leakage after colorectal surgery.

In collaboration with the Department of Medicine and the ISFSI of EOC, our group has also evaluated the impact of the gut microbiota composition on immune responses to SARS-CoV-2 in COVID patients and in healthy subjects undergoing vaccination. We identify bacterial genera associated with either good or poor response to anti-SARS-CoV-2 vaccine in healthy subjects. The causative effect of identified bacteria is currently being tested in experimental models, by means of fecal microbial transplantation experiments.





Neurodegenerative Disease research group

(Prof. P. Paganetti)

Protein conformation disorders: abnormal protein conformation, deposition and proteotoxicity are common traits of progressive degenerative disorders of the brain causing relatively frequent diseases but also rare inherited syndroms. The protein Tau is involved in the development of several brain disorders, most prominently Alzheimer's disease, and less frequent progressive tauopathies, including frontotemporal dementia and parkinsonism linked to chromosome 17 caused by mutations in the gene encoding for Tau. The disorders are associated with the brain deposition of tau in fibrillary tangles. When deposited Tau present several characteristics modifications such as hyperphosphorylation and a pathological conformation. How this occurs and causes neuronal dysfunction is poorly understood, but the presence and distribution of tangles correlate well with the cognitive loss observed in patients afflicted by the disorder.

Nephrology research group

(Prof. P. Cippà)

The function of the kidney depends on the precise organization and coordination of more than 30 cell types, forming the functional units called nephrons. A variety of clinical conditions, such as diabetes, drug toxicity, and autoimmune diseases, can damage kidney cells and alter the balance necessary for the organ to properly function. Since the kidney loses the ability to generate new nephrons after birth, the maintenance of renal function after kidney injury depends on the compensatory adaptation of the remaining nephrons, and the ability of the kidney to repair damaged nephrons. Unfortunately, this process is often incomplete and can result in the persistence of altered cells and a dysfunctional tissue architecture, which contribute to the progression of renal disease towards renal failure. In this stage the kidney damage is irreversible, and the patient rely on dialysis or kidney transplant to survive. Our research aims to characterize and better understand the biological processes of kidney repair. The goal is to support functional tissue regeneration processes and prevent the progression to chronic kidney disease.

Parkinson research group

(Prof. G. Melli)

Parkinson's disease (PD) is a chronic progressive neurodegenerative disease associated with intra-neuronal accumulation of aggregated alpha-synuclein (α -Syn) protein, but unfortunately current knowledge still does not address the underlying pathogenic mechanisms. Consequently, PD lacks both an effective causal treatment and definitive early diagnostic criteria. In the recent years a large amount of evidence has shown that peripheral tissues like skin autonomic nerves, gastro-intestinal mucosa, salivary glands show immunoreactivity to pathogenic forms of α -Syn.

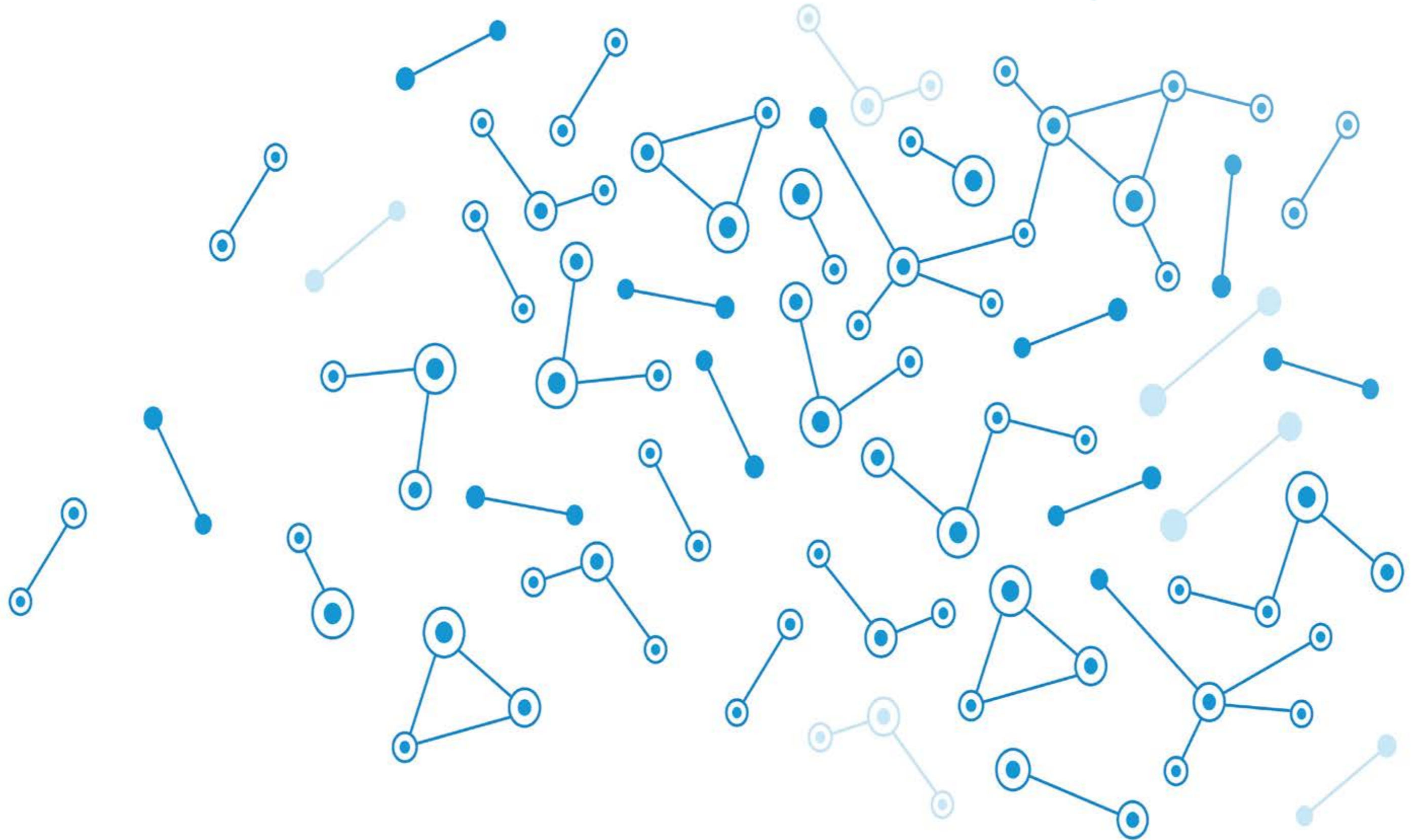
Thus, peripheral tissues, that can be easily biopsied, contain nervous tissue prone to the pathology, and bear potentially important information on disease mechanisms.

Our laboratory has a strong translational attitude and aims to set-up novel pathway biomarkers of disease by analyzing skin biopsy and peripheral fluids via advanced new biotechnology. Our group has published several papers on the detection/ characterization of α -Syn and its pathogenic species in the skin of PD and other neurodegenerative diseases like atypical parkinsonisms plus we demonstrated that

PD patients present a somatosensory small fiber neuropathy that correlates with disease duration and progression.

More recently we published several papers on the specific profile of surface proteins of blood- and CSF-derived extracellular vesicles (EVs), which stratifies patients with PD and atypical parkinsonisms. These EV markers are related to circulating inflammatory and immune cells. We are now profiling prodromal and early phases of disease to identify specific signature and we are investigating how peripheral inflammation and immune dysregulation can contribute to brain neuronal cell death in PD. These projects take advantage of a biobank and clinical database for PD and atypical parkinsonisms that were established at INSI in Lugano since 2015.

Research Services



Clinical Trial Unit EOC

Prof. Alessandro Ceschi, MD, MSc
CTU-EOC Director

The mission of CTU-EOC is to promote and increase clinical research within EOC by offering high quality services for the implementation and conduction of clinical trials in all fields of medicine ensuring the application of national and international regulatory requirements. CTU-EOC celebrated 10 years of operation in 2022, and the number of requests has increased year by year. Thanks to experienced collaborators, support in clinical research is possible for both academic and industry research projects. CTU-EOC has been actively involved in monocentric and multicenter projects in Switzerland and Europe.

During 2022, the CTU-EOC took in charge 66 projects, of which 40 were interventional (ClinO, ClinO-DM). The most requested service was biostatistics, followed by the development of eCRFs, document writing and review, and the submission to Regulatory Authorities. Seventeen projects were promoted by external researchers, of which 5 promoted by the Industry.

Since 2020, the CTU-EOC is a full member of the Swiss Clinical Trial Organization (SCTO), which is a not-for-profit Organization acting as the central cooperation platform for patient-oriented clinical research in Switzerland, and it includes the CTUs of the University hospitals and the cantonal hospital of St. Gallen. CTU-EOC is the coordinator of the SCTO safety platform.

Further to provide the whole range of competences required to support a clinical study, CTU-EOC also provides individual support to students, particularly of the USI Faculty of Biomedical Sciences, who have to perform their master or doctorate thesis.



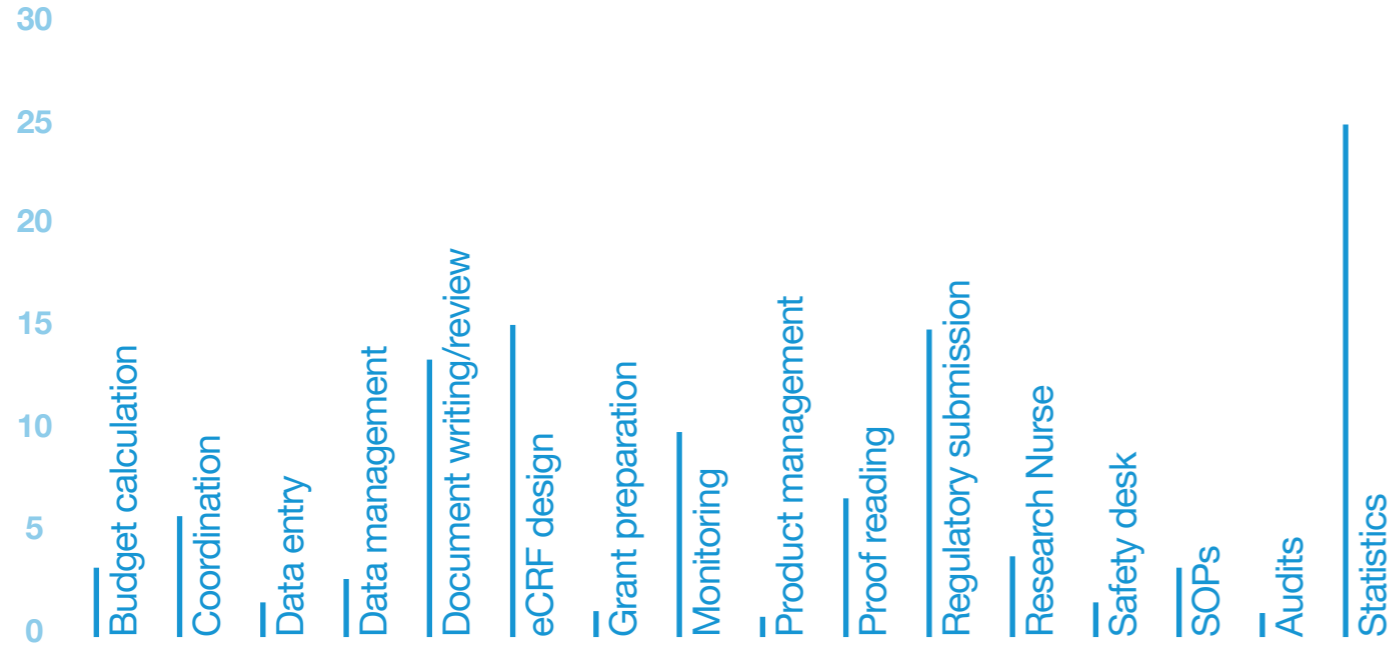


Figure 1.
Services provided
by the CTU-EOC
in 2022

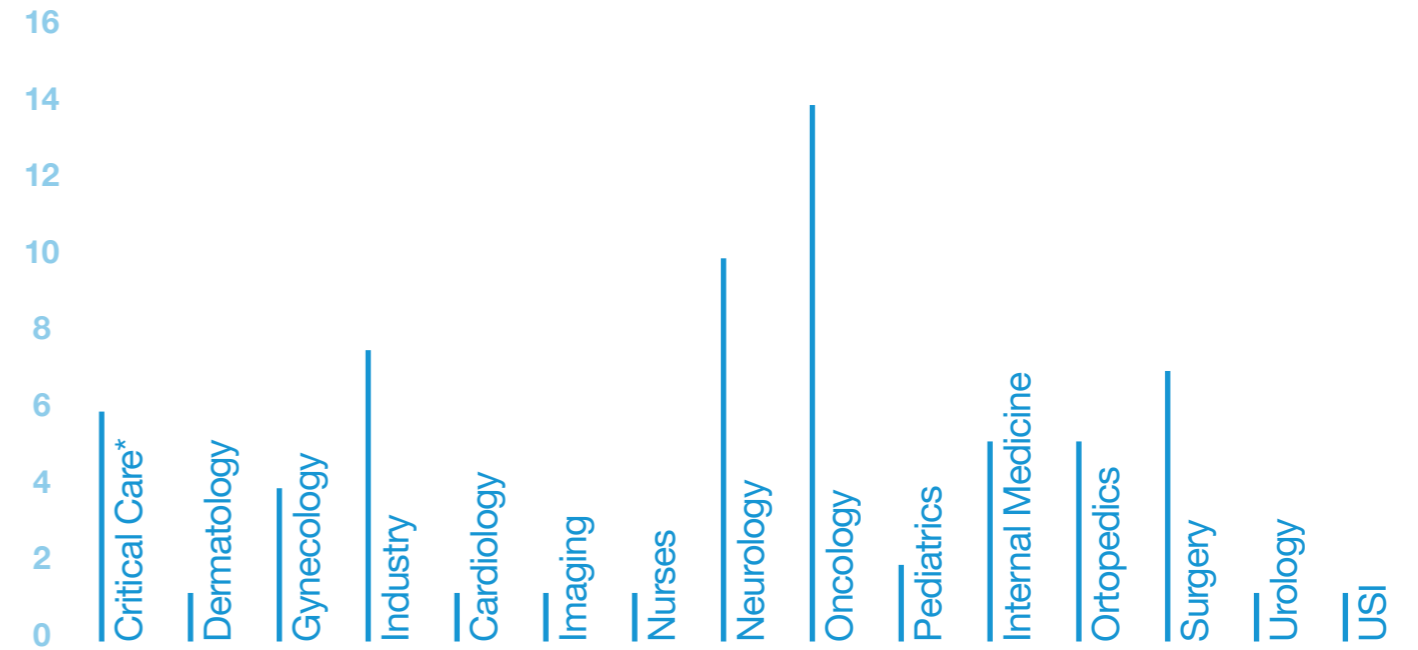


Figure 2.
Collaborations of
the CTU-EOC in
2022

*Anesthesiology, intensive care, emergency department. USI: Università della Svizzera italiana

Education

The educational program of the CTU-EOC points to contribute in developing the next generation of researchers with high quality and dynamic courses. The program is adaptable to respond to needs and requirements of researchers in our institution and among our partners. We offer harmonized and recognized GCP training, up-to-date courses according to new regulatory and legal requirements, and we are always attentive to new challenges to develop innovative programs.

CTU-EOC continue to give courses within the PhD programs of the Faculty of Biomedical Sciences at USI as general skills, that are also open to PhD students of our partner's institutions as ETHZ and USZ. Courses are delivered in English in the spring and in Italian in the fall session yearly. CTU-EOC actively contributes to the SCTO educational platform, ensuring a constant exchange and update in the field at the national level.

Events

CTU-EOC, in collaboration with the Medical education and Research Division (AFRi) of the EOC general directorate and the USI Faculty of Biomedical Sciences, organized the "11a Giornata della Ricerca e dell'Innovazione in Medicina Umana della Svizzera italiana". The participation has constantly increased over the years, both in numbers of abstracts submitted in each category of research (80 for clinical or epidemiological research, 21 for nursing research, and 58 for basic translational or computational research) and in number of attendees (342 in 2022).

More information is available at the following link:

<https://www.ctueoc.ch/giornata-dellaricerca/>

External participants

Internal participants

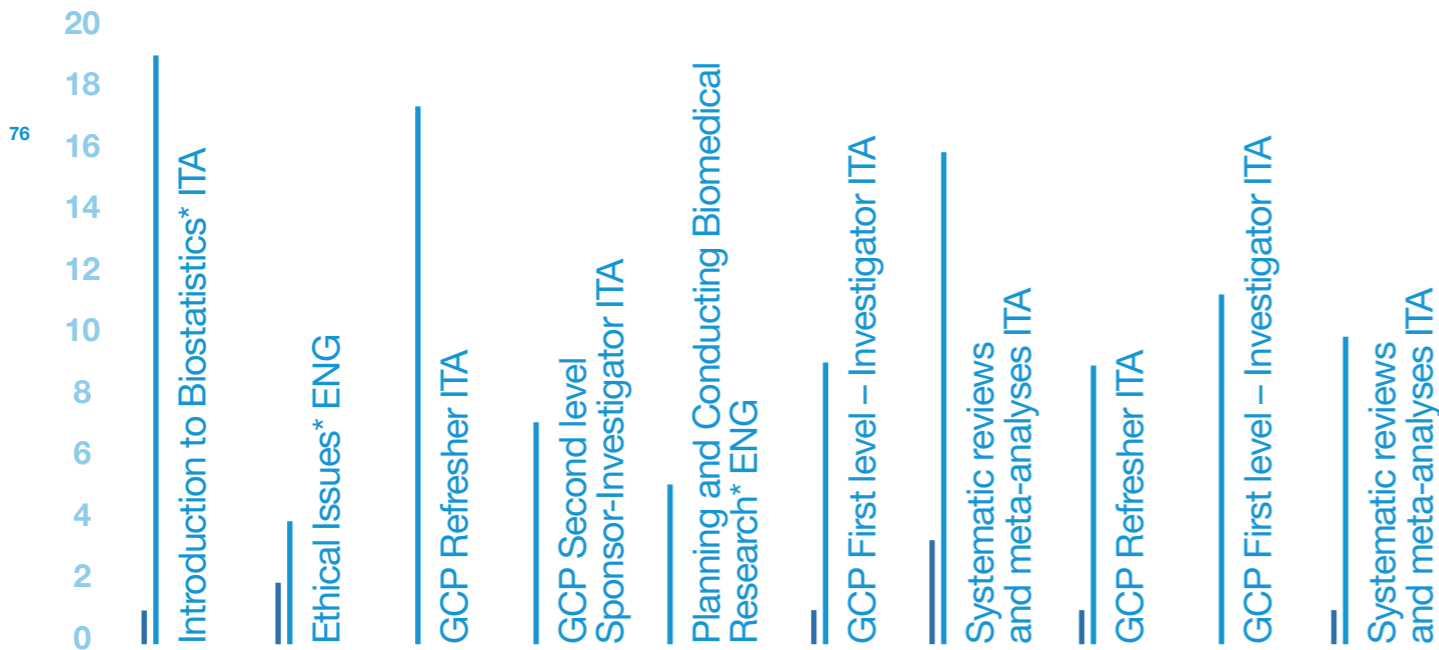


Figure 3.
Courses organized by the CTU-EOC in 2022

*PhD courses - General Skills - open to Internal and External participants



Grants and fundraising Unit

Annual private third-party funding in the form of grants or donations for research projects encourages talented EOC researchers working on innovative projects aimed at ensuring effective, appropriate, and efficient health treatments.

The Grants and Fundraising Unit at AFRI is the reference point for participation in national, European and international research funding (Calls) and the coordination point for the fundraising activities through the EOC-USI Fund for biomedical research.

The Unit also works closely with the Research Service of USI (SRIT) providing the necessary support to EOC researchers affiliated to the USI Faculty of Biomedical Sciences both in the proposal submission phase and for financial reporting.

We provide advice and support to EOC researchers to transform their research ideas into fundable projects. We are very proud of the new collaborations created in 2022 with National funding agencies and other public funding institutions in Switzerland and abroad. There was close to 20 new successfully funded projects, which raised research funds for clinical and translational research studies.

We work closely with our founders and are grateful for the trust from donors and research partners. Together, we create knowledge experiences that contribute to meet the needs and requirements of patients and their families.

For more information about grant management and private third-party funding at EOC, please contact the Grants & Fundraising Unit:

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EOC Biobank

The Biobank-EOC in 2022 continued its activity in the management of biological samples within the framework of clinical studies mainly promoted by EOC researchers.

Currently, there are 5 collections of blood-derived biological specimens under management with more than 41,000 aliquots stored at the facility.

Collaborations with diagnostic companies have also started to introduce new tests to the market or to further improve existing ones. In addition to the importance of new knowledge in clinical medicine, patients can thus also benefit from improvements in diagnostic tests.

The EOC biobank is accessible not only to EOC researchers but to all research groups interested in benefiting from an infrastructure dedicated to the collection and storage of biological samples, as well as their use.

For more information:

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Intellectual property

Intellectual property (IP), and related rights, indicates a system of legal protection to protect creative and inventive activities from commercial abuse and illicit reproduction. The significance of intellectual property is primarily of commercial nature.

EOC promotes the development of scientific knowledge and the enhancement of the results of research activities, also through the transfer of technology related to the intellectual creations of EOC researchers.

All inventions and any other innovation likely to be the subject of an industrial patent (or similar title related to intellectual property) that has a patrimonial value and is susceptible of exclusive rights are governed by the EOC regulation on intellectual property (EOC_D-Cda-023).

The EOC Intellectual Property Commission (IP Commission) is in charge of the implementation of this regulation. This commission is composed by collaborators of the EOC General Directorate. The USI Technology Transfer Manager is also a member of this commission and supports its activities. The IP Commission has the competence on the protection and enhancement of inventions or potentially patentable and / or otherwise protectable inventions. The IP Commission is required to meet every time there is a request for the patentability of an invention or any subject of intellectual property.

General consent for research purposes

In 2021 the AFRi EOC started the introduction of the General Consent for research purposes in the EOC Institutes and Departments.

A large part of research is based on patients' personal health data contained in their medical records, such as laboratory test results, treatments administered and genetic predispositions. All the biological material collected at the hospitals and no longer required for treatment is also very important for research purposes.

Patients can make a contribution to the research by agreeing for their clinical data and/or biological samples to be stored, sent to third parties and reused for research purposes, subject to approval by the competent Ethics Committee. Through the introduction of the General Consent for Research at the EOC hospitals, the patients may now give or not their consent on the use of their personal health data and biological samples for research purposes.

For more information:

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USI Faculties and Institutes

Research activities in biomedical sciences at USI are mostly organized within the Institutes of the Faculty of Biomedical Sciences, but also comprise more interdisciplinary activities in domains of computational sciences, health economics, and health communication in cooperation with the Faculties of Informatics, Economics, and Communication, Culture and Society.



USI Faculty of Biomedical Science

USI Faculty of Biomedical Sciences was established in 2014 with the aim of addressing an important national problem: the dearth of physicians trained in Switzerland. To reach its goal, the Faculty offers from 2020 a Master of Medicine (three years of clinical training), in cooperation with ETH Zurich and University of Basel regarding academics, and with EOC, clinics, and general practitioners regarding clinical training. The Master addresses the new challenges of medical practice by combining clinical and scientific training with communication skills. The Faculty is also offering a joint Master's programme in Cognitive Psychology in Health Communication with Vita-Salute San Raffaele University, Milan.

The Faculty is offering a PhD program in Biomedical Sciences with majors corresponding to its main research areas, i.e. immunology and cell biology, cancer biology and oncology, (human) neurosciences, (human) cardiovascular sciences, drug sciences, public health and computational biomedicine, as well as a Medical Doctor Programme. Moreover, the Faculty organizes an Executive Programme in Sleep Medicine (with the University of Bern), in Gastrointestinal, Gynecological and Prostate Cancer (with the European School of Oncology) and in Patient's Communication (with Fondazione Sasso Corbaro). Finally, the Center of Advanced Studies on Entrepreneurship in Biomedicine (CASE BioMed) adds to the medical and

scientific education at the Faculty by offering advanced programmes in innovation and entrepreneurship.

The Faculty currently comprises more than 20 full, associate and assistant professors, as well as a large number of adjunct, titular professors, and of private lecturers; these mostly include directors and senior scientific staff of the EOC presented in section 1 of this report.



Research in the Faculty is developed in immunology and basic cancer research at the two affiliated institutes IRB and IOR and in the transnational research laboratories of EOC epidemiological and clinical research is performed within the Institute of Public Health at USI and in the institutes and centers at EOC (see chapters 1-3 of this report); finally, research in computational biomedicine is developed at the Euler Institute:

- The Institute for Research in Biomedicine (IRB; www.irb.usi.ch), affiliated and based in the Bellinzona campus. IRB was founded in 2000 with the goal of advancing the study of human immunology, with particular emphasis on the disease-relevant mechanisms of host defense. Since 2010, it is affiliated to USI as part of its Faculty of Biomedical Sciences. It currently comprises 13 research groups about 120 international collaborators.
- The Institute of Oncology Research (IOR; www.ior.usi.ch), affiliated and based in the Bellinzona campus, hosts researchers from all over the world performing basic and translational research in oncology with special focus on cancer

biology, genomics, molecular oncology, experimental therapeutics. Since 2017, IOR is affiliated to USI as part of its Faculty of Biomedical Sciences. The Institute currently comprises 8 research groups and about 70 researchers and PhD students.

- The Institute of Public Health (IPH), see below.
- The Euler Institute (www.euler.usi.ch), common to the Faculties of Informatics and Economics. Euler is USI's central node for interdisciplinary research and the connection between exact sciences and life sciences. Euler applies advanced computational methods to fields such as Life Sciences, Medicine, Physics, Mathematics, and Quantitative Methods. It currently comprises four professors, as well as several adjunct professors and groups leaders and about 45 researchers and PhD students.

Institute of Public Health (IPH)

The Institute of Public Health (IPH) is an inter-faculty Institute and one of the five training and research entities supported by the Faculty of Biomedical Sciences of the Università della Svizzera italiana. Members of the IPH perform research, training and practice, and provide services across a broad range of activities related to public health. The IPH is the 'antenna' in Ticino of the Swiss School of Public Health (SSPH+) and is home to research programs articulated across several research projects. In addition, IPH members provide support and scientific advice on the methodology of epidemiological, biomedical, behavioral, and other areas of research, from the conception, to the conduction, through the implementation of studies with various designs and spanning qualitative and quantitative methodological approaches. IPH is an USI inter-faculty Institute with members from the Faculty of Communication, Culture and Society, the Faculty of Economics, and of Biomedical sciences.

The IPH staff currently includes the Director, 4 tenured Professors, 2 Senior Researchers, and several Postdoctoral Fellows, PhD students, and research Assistants. Their backgrounds range from the medical field, to epidemiology, health economics, health communication, health promotion and behavior, communication, public management and policy, psychology, and marketing, through the social and health sciences.

Members of the IPH conduct a variety of studies in the fields of aging, mental health, ethics, public health, socio-emotional well-being, cognitive development and decline, social media addiction, cognitive and social neuroscience, clinical psychology, psychophysiology, social psychology, human behavior, nutrition and physical activity, vaccine acceptance, social integration, and health psychology. Study designs include epidemiological, population-based studies, experimental studies, qualitative studies (interviews and focus groups), fMRI studies, media content analyses, psychophysiological studies, scoping reviews, systematic reviews, meta-analyses, and RCTs. Target populations include older adults (>65), younger adults (18-35), adolescents (<18), and healthcare providers.

Researchers at the IPH have strong collaborations at the local (e.g., Canton of Ticino, ProSenectute Ticino e Moesano, Alzheimer Ticino, SUPSI), national (all Swiss universities, ETHZ, and EPFL) and international level (University of Vienna, University of Parma, University of Milan, University of Turin, and Loughborough University, University of Pittsburgh Graduate School of Public Health, Imperial College London, University of Valencia, University of Nottingham, University of York, University of Kent, National University of Ireland Galway, and many more), as well as with the World Health Organization, UNICEF, SDC, and many NGOs based in Switzerland and abroad.

Members of the IPH give courses in various USI Bachelor's degrees, and within the USI Master in Medicine, the Master in Communication, Management and Health, and the Master in Cognitive Psychology in Health Communication, Masters of Marketing, Master in Public Management and Policy, Master in Corporate Communication, Master in Digital Fashion. Other graduate trainings include the SSPH+ Summer School in Public Health Policy, Economics, and Management, the Net Megs and more.

The IPH is committed to promote the highest ethical and methodological standards in population sciences and embraces a strong participatory action research approach and community engagement of the public and of all relevant stakeholders. Both research and training at the IPH ultimately aim to improve public health, to inform policy decisions and practices, and to contribute to positively influence individual behaviors and societal practices that promote and protect health, prevent diseases, and favor optimal and equitable care based on needs.

Other USI Faculties and Institutes

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The Institute of Economics (IdEP; <http://www.idep.eco.usi.ch>) supports research and teaching activities in the broad field of economics and public management. The research interests of IdEP members also include health economics and public health. Their research focuses on issues of public interest such as ageing and cognitive decline, mental health, sleep deprivation, and the determinants of the increasing health care costs. IdEP also organizes a Master of advanced studies in health economics and management (NETMEGS). Since 2012 IdEP coordinates the activity of the Center for Economic and Political Research on Aging (CEPRA). The center aims at stimulating empirical research on population aging, pension and health care design and spending, at promoting academic training on this topic, and at disseminating research output to a specialized and a general audience, through workshops, conferences, and public debates. Finally, since 2012 IdEP is part of the national research project Swiss learning health.

Within the Faculty of Communication, Culture and Society, there are specific competences related to health communication and literacy (prof. Schulz, prof. Hannawa); the Faculty also offers a master program in Communication, Management and Health. The Institute of Public Communication also closely cooperates with the Institute of Public Health to conduct research to understand determinants of health-related behaviors and to develop strategies to change behavior, focusing on communication (BeCHANGE research group). This includes topics such as making healthy behaviors 'easy' options for people, communicating complex health messages and increasing vaccine acceptance and uptake.



Peer-reviewed publications



Peer-reviewed publications

A comprehensive computer literature search of PubMed/MEDLINE database was carried out independently by two different collaborators of the EOC Medical education and Research Division. The last database search was performed in March 2023. The string search was built using keywords based on the different EOC sites.

Journal Impact Factor (IF)¹ and Journal IF Quartile² of the scientific journals in which EOC researchers published their peer-reviewed articles were also evaluated using data from the Journal Citations Reports (JCR) 2021.

The scientific work carried out by the EOC researchers, based on the inclusion and exclusion criteria mentioned above, produced **779** peer-reviewed articles. Although the total number is slightly lower than 2021, the EOC scientific production in 2022 - as illustrated in Figure 4 - confirmed the upward trend recorded the previous year.

The following inclusion criteria were used in the selection process:

- peer-reviewed articles listed in PubMed/MEDLINE with at least one co-author affiliated with EOC;
- final publication of the article in a scientific journal issue in 2022. Articles listed in PubMed in 2022 as "Online ahead of print" before their publication in final or print format were therefore excluded from the current analysis.

¹ Journal IF is defined as citations to the journal in the Journal Citations Reports (JCR) year to items published in the previous two years, divided by the total number of scholarly items (also known as citable items - these comprise articles and reviews) published in the journal in the previous two years. A Journal IF of 1.0 means that, on average, the articles published one or two years ago have been cited one time (Source: Clarivate Analytics).

² Scientific journals are assigned to a subject category by JCR and ranked by Journal IF. The Journal IF rank is then transformed in a percentile value. The Journal IF percentile is used to calculate the Journal IF Quartile. Q1: Journal IF percentile ≥ 75 ; Q2: $75 < \text{Journal IF percentile} \leq 50$; Q3: $50 < \text{Journal IF percentile} \leq 25$; Q4: $25 < \text{Journal IF percentile}$ (Source: Clarivate Analytics).

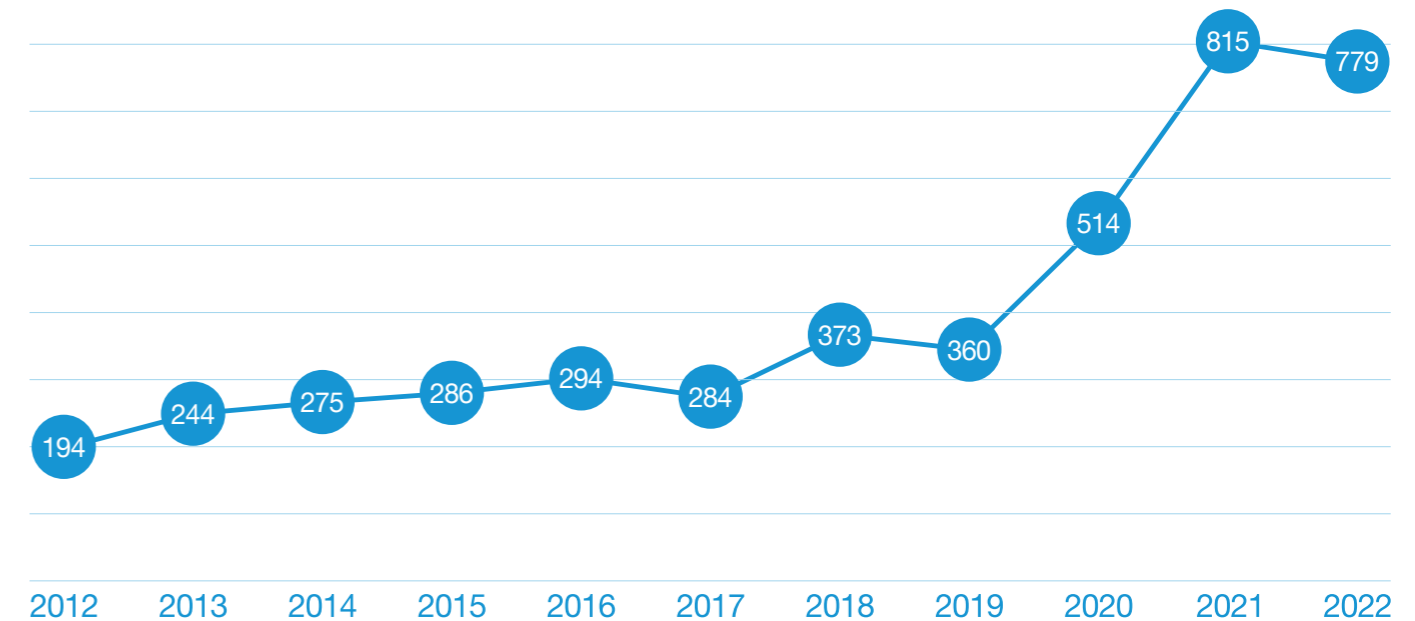
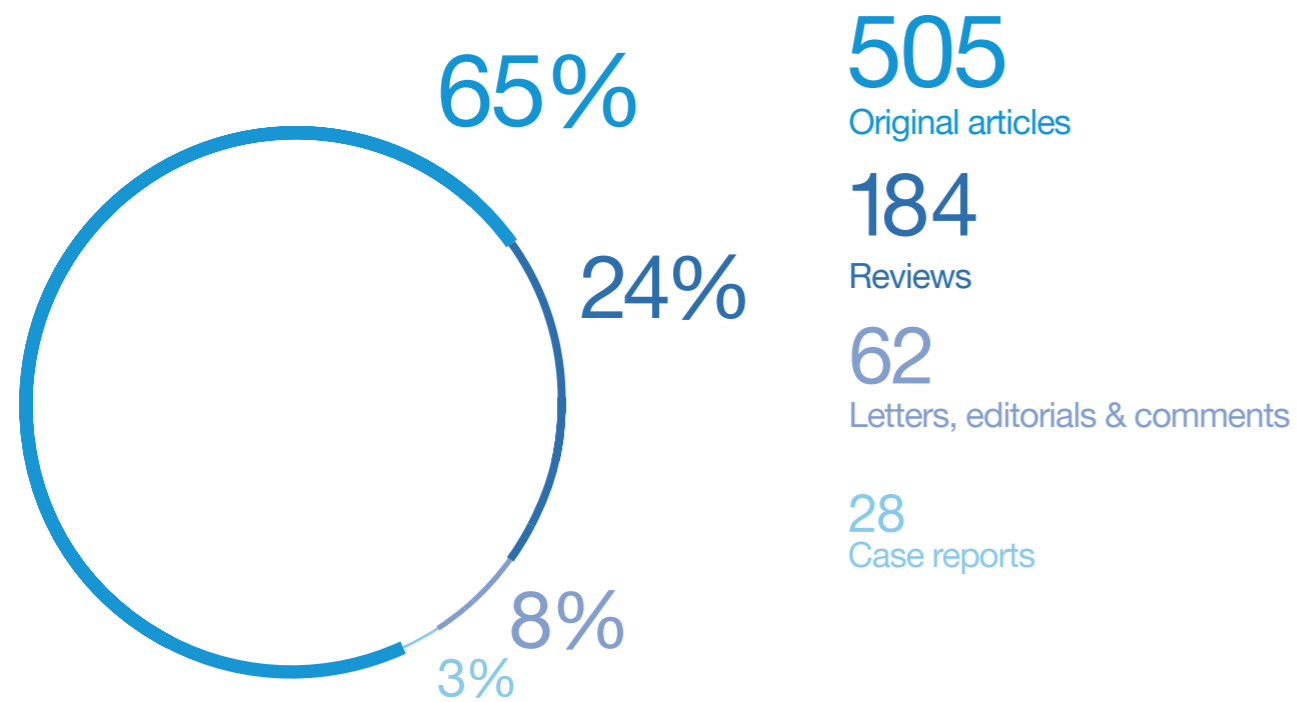


Figure 4.
Number of EOC peer-reviewed articles published by the EOC researchers since 2012

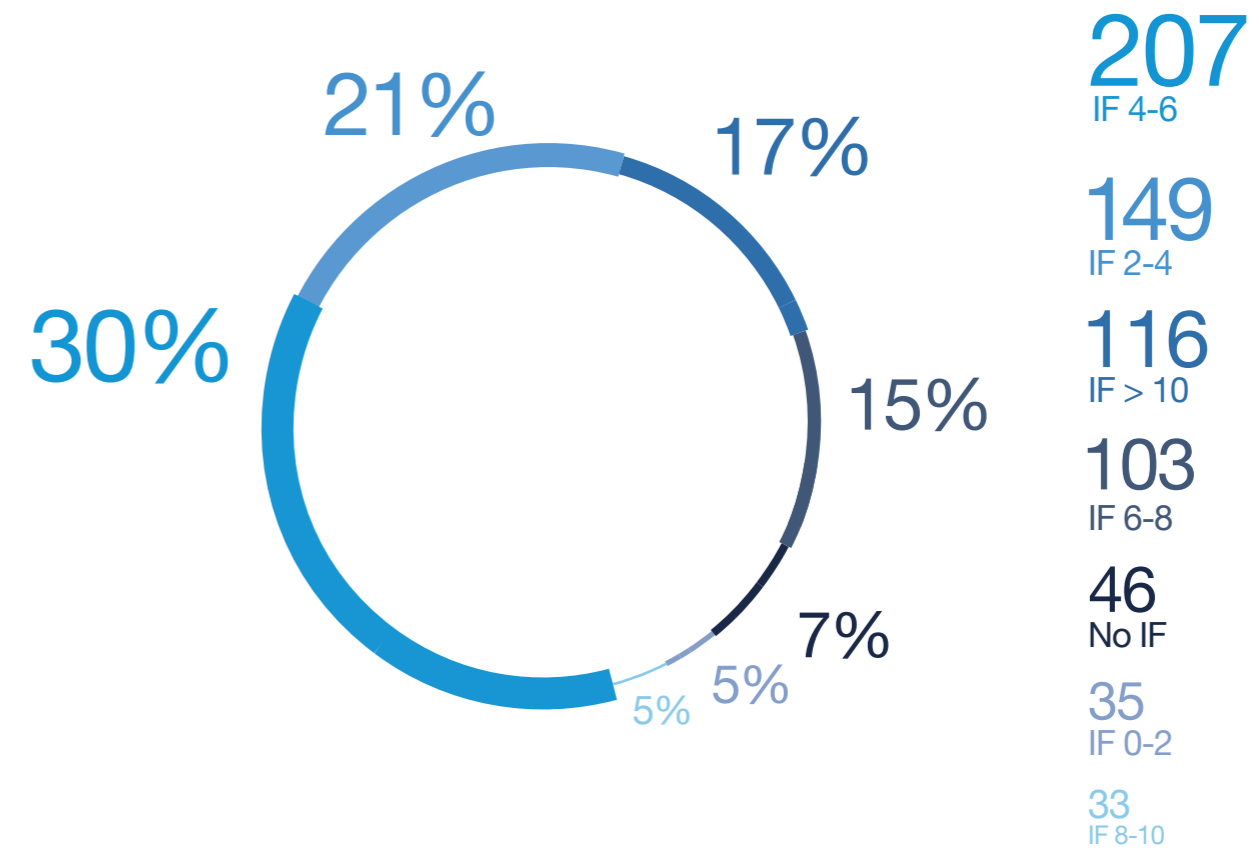
Most EOC peer-reviewed publications in 2022 were original articles or reviews (**689**, 89%) in line with 2021 (87%). Publications in Q1 journals decreased by 7% compared to the previous year, however the total number of articles published in Q1 and Q2 journals (77%) remained at the same high level as in 2021 (74%) confirming the high quality of the EOC scientific outputs. Median IF of original articles or reviews was 5.2, slightly higher than in 2021. Notably, 17% (N=116)

of articles were published in journals with high impact factor (Journal IF ≥ 10). The EOC researcher was the first or last author in 40% (N=310) of original articles or reviews, of which almost 35% were published in Q1 journals and 10% in high impact factor journal. Even in 2022, most publications were the result of an important network of international and national collaborations (93%). Key data are summarized in figures 5-10.



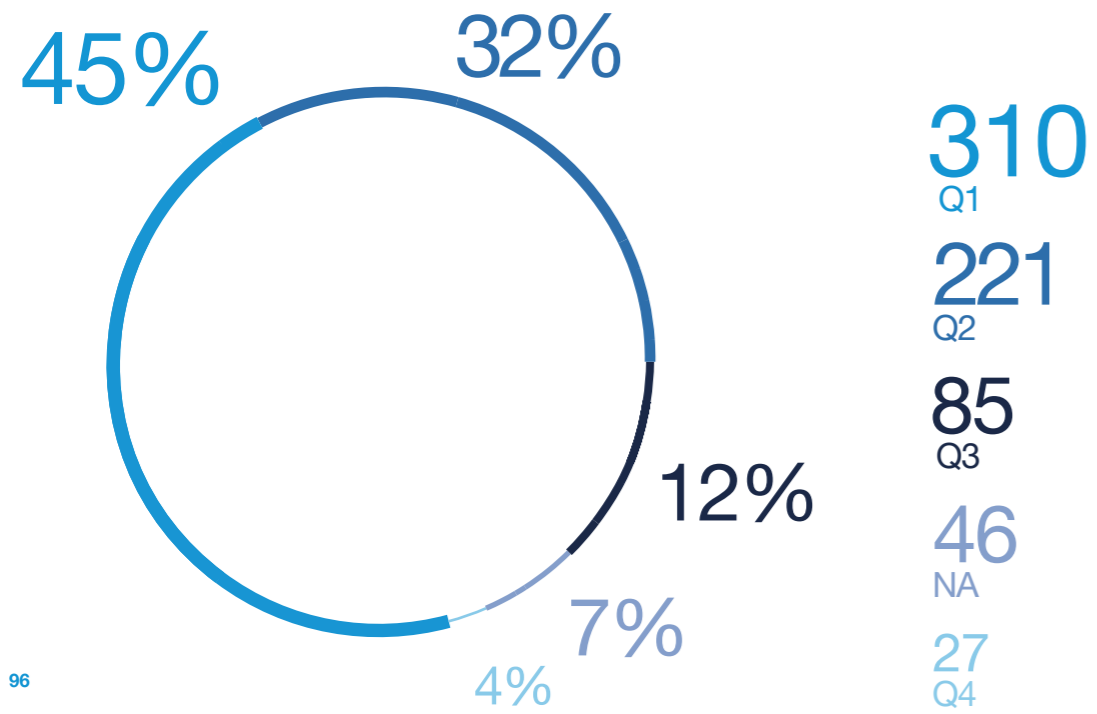
**Figure 5.
Number and type
of articles**

Number and type of the peer-reviewed articles published in 2022 (N = 779). Meta-analyses, surveys, research letters, consensus and delphi-based consensus paper are classified as original articles. Systematic reviews without meta-analyses and protocols are classified as reviews.



**Figure 6.
Impact Factor**

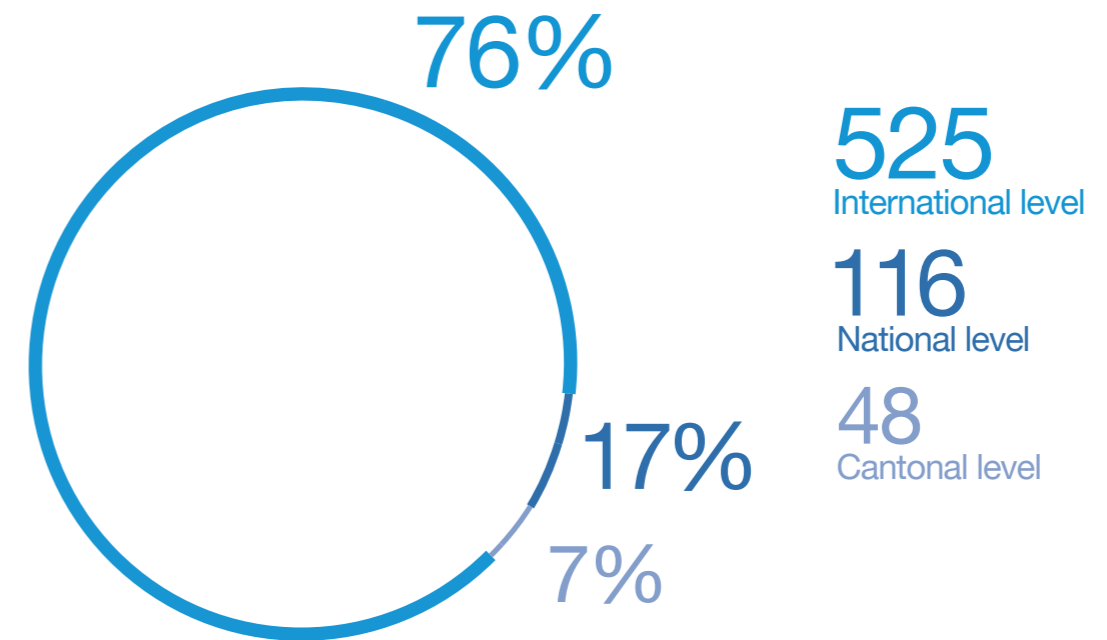
Journal IF of the scientific journals in which peer-reviewed original and review articles were published in 2022 (N = 689).



96

**Figure 7.
JIF Quartile**

Journal IF Quartile of the scientific journals in which peer-reviewed original and review articles were published in 2022 (N = 689). Journals can appear in multiple subject categories; in this case, the highest Journal IF Quartile was selected. Scientific journals without Journal IF are classified as NA.



**Figure 8.
Collaborations**

Type of scientific collaboration (local, national or international) in peer-reviewed original and review articles published in 2022 (N = 689).

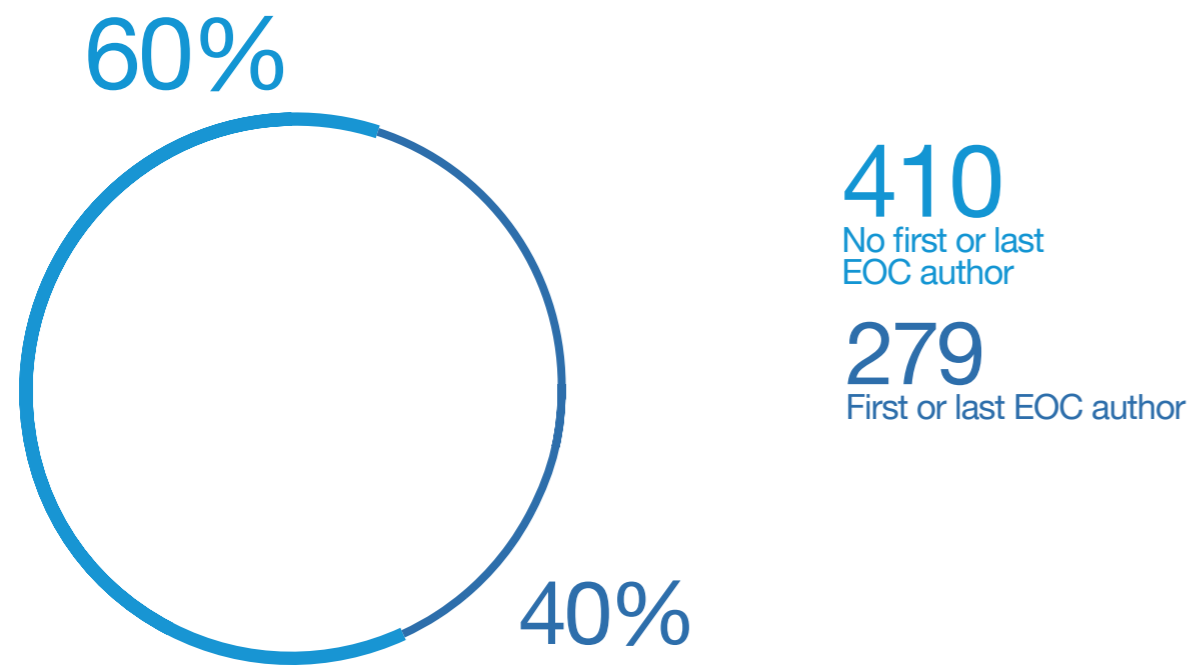


Figure 9.
EOC researcher
as first or last
author

Number of peer-reviewed original and review articles published by EOC researchers as first or last author in 2022 (N = 689).



Original articles Reviews Other

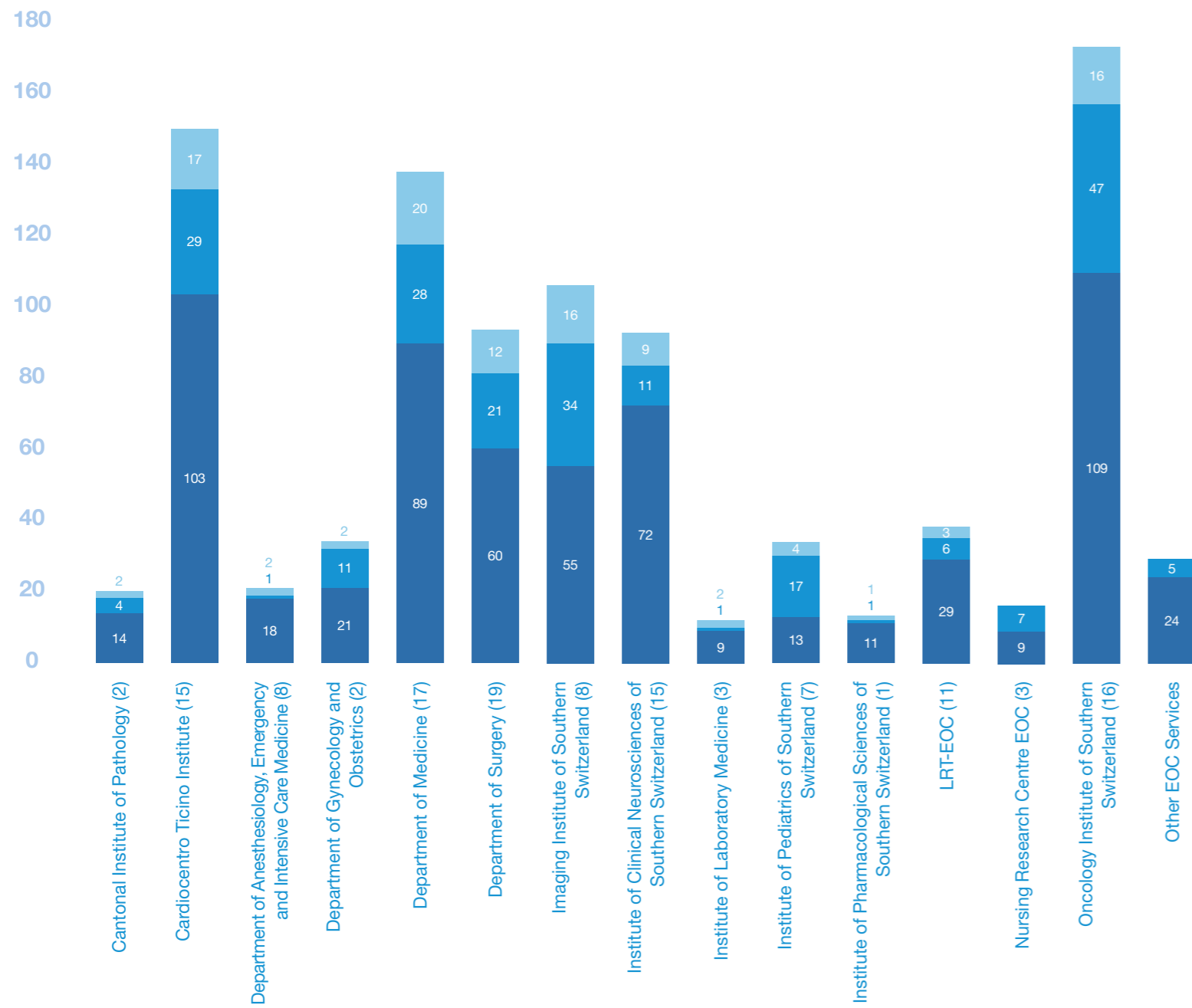


Figure 10.
EOC Institutes and
Departments

Number of peer-reviewed articles published by each EOC Institute and Department in 2022. Publications shared among different EOC Institutes/Departments were counted for each Institute/Department involved. The name of the

Institute/Department with the related number of Professors and PDs are indicated on the horizontal axis. Almost 5% of the EOC scientific publications were from LRT-EOC.



Research projects activated in 2022



SCIENCE 2.0

```
page_select_objects
if(len(bpy.context.page_select_objects) == 1)
    modifier_ob = bpy.context.active_object
    print("one is selected")
    mirror_mod = modifier_ob.modifiers.new("mirror_mirror","MIRROR")
    # mirror_ob.select = False
    # mirror_ob = bpy.context.selected_objects[0]
    # page_select_objects
    mirror_ob.select = False # last ob selected
    mirror_ob.select_set(False) # pop modifier_ob from sel_stack
    print("popped")
#_page_select_objects
modifier_ob = bpy.context.selected_objects[0]
print("Modifier object: "+code.next
```

IOLOGY

Research projects conducted at the EOC hospitals are recorded into a dedicated database. A total of **115** research projects involving persons that require authorization by the Ethics Committee and/or Swissmedic were activated in 2022. As of 31.12.2022, **342** research projects were ongoing at the EOC hospitals.

Figure 11 shows the number of EOC research projects by the category to which are assigned on the basis of the type of effect they have on participants. Most (64%, N=74) were non-clinical trials, almost equally divided between

the sub-categories “research involving persons” (N=39) and “further use of pre-existing health-related data and/or biological material” (N=35). The EOC general consent was used in 46% of the non-clinical trials, including all “further use” research projects. The remaining were clinical trials (N=41) of which over 60% investigating medicinal products. According to available data, EOC researchers conceived and promoted 49 research projects in 2022, which were mostly non-clinical trials (N=43). Almost 45% of the research projects were funded, primarily clinical trials (71% of them funded).

As shown in Figure 12, more than half of the EOC research projects involved more than one participating site internationally (N=39) or nationally (N=25).

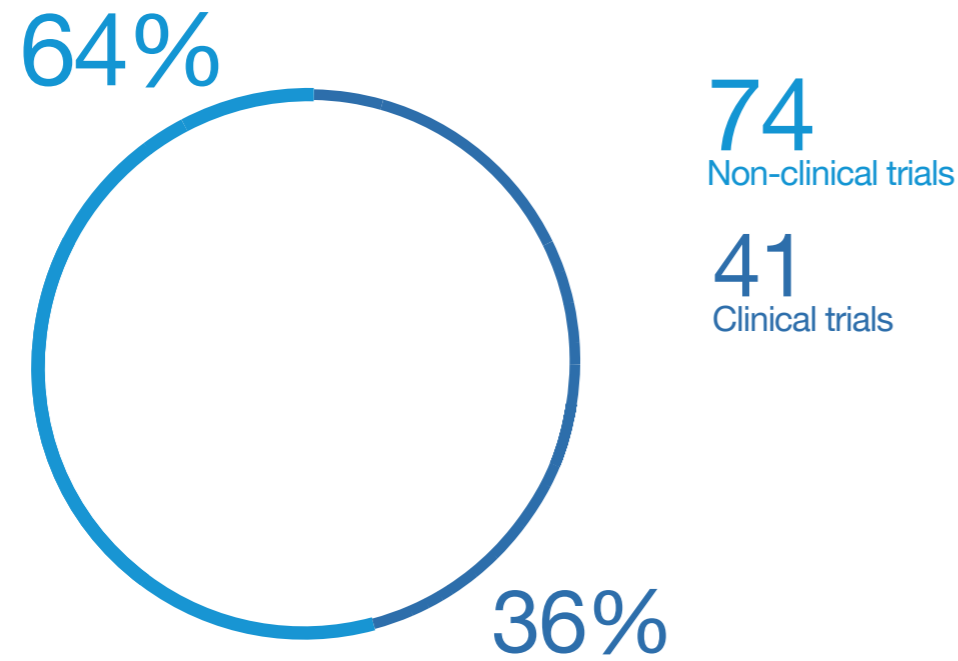


Figure 11.
Number of EOC research projects that require authorization activated in 2022

Number of EOC research projects activated in 2022 (N = 115).

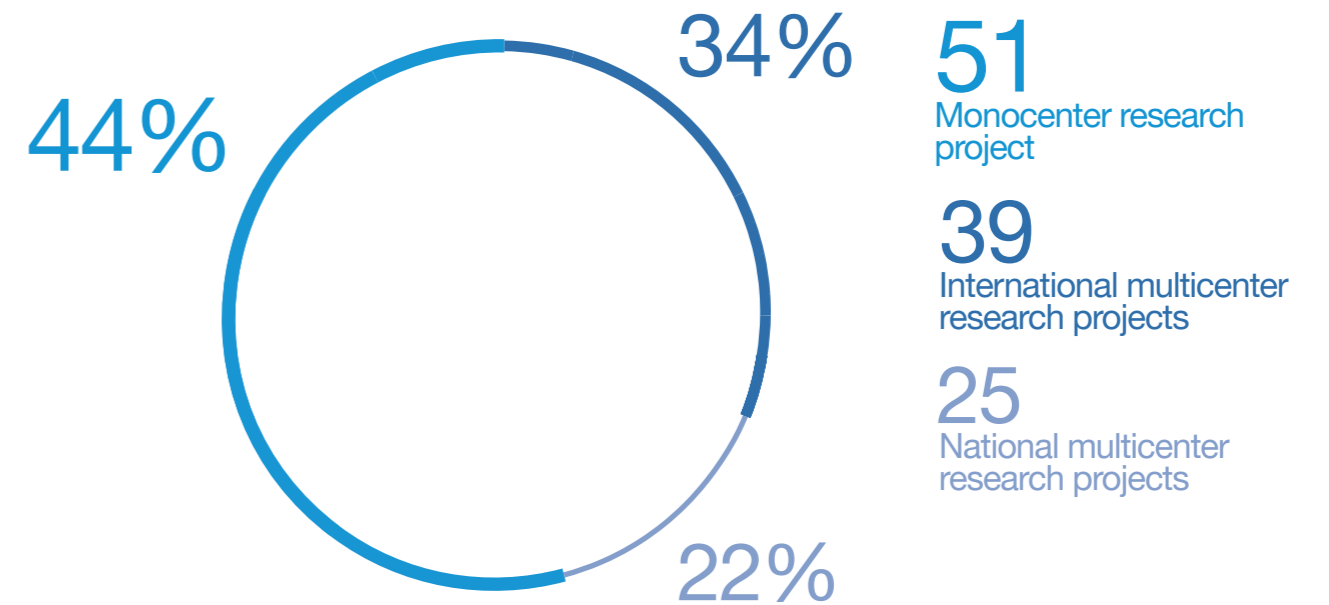
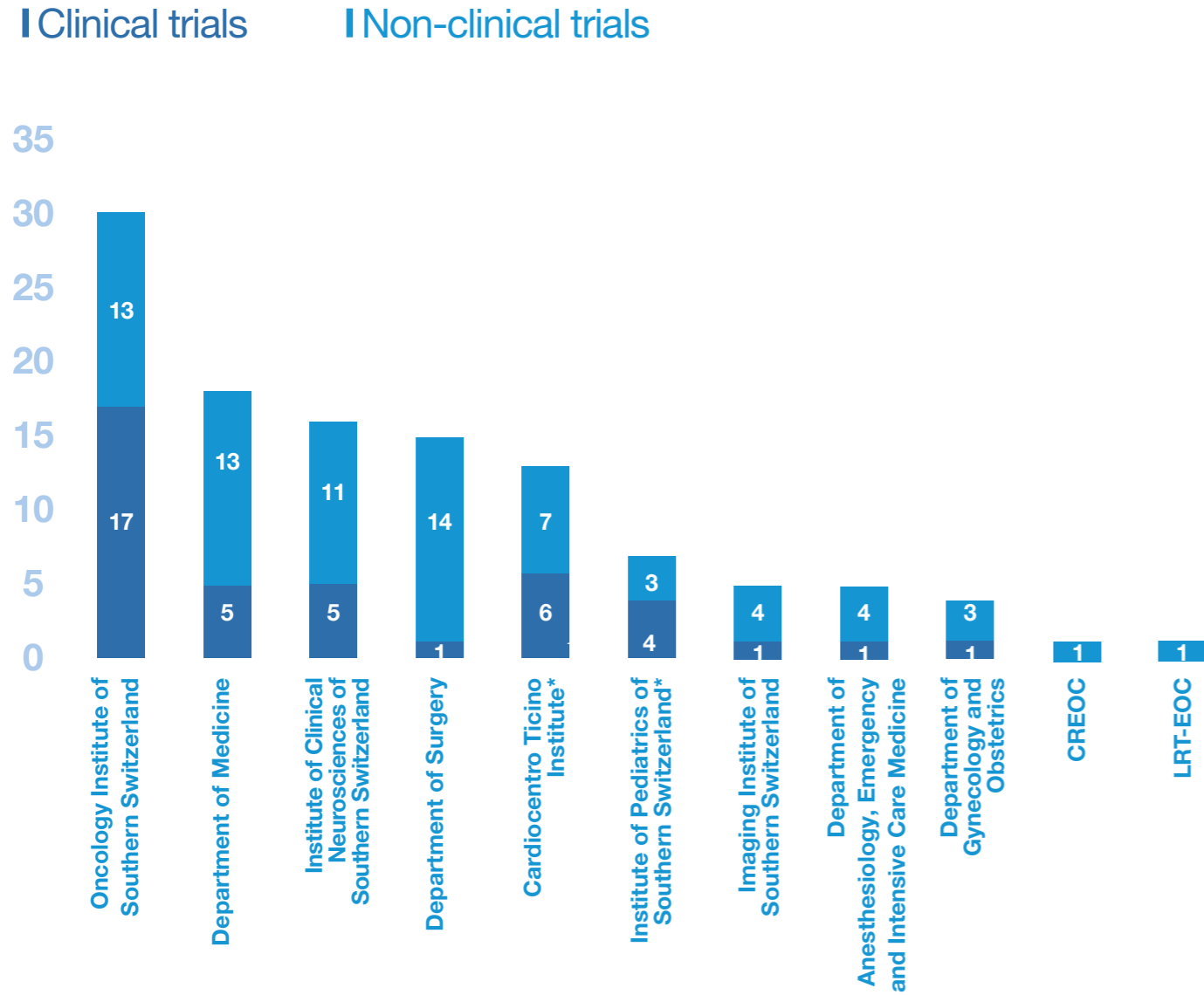


Figure 12.
Number of EOC mono- and multicenter research projects activated in 2022

Number of EOC research projects with single or multiple participating sites activated in 2022 (N = 115).



*Including 1 nursing study

Figure 13.
Type of research projects activated in 2022 by Institute/Department

Number of EOC research projects activated in 2022 by Institute/Department (N = 115). Clinical and non-clinical trials are also shown.



Figure 13 summarizes the number of EOC research projects activated in 2022, by clinical and non-clinical trials and by Institute/Department. The research project was assigned based on the EOC affiliation of the Sponsor-Investigator/Project Leader/Principal investigator, whichever applies.

Although the total number of newly activated research projects at the EOC hospitals has slightly decreased compared to 2021, overall statistics for 2022 are consistent with the data reported in previous years.

Overall, about **CHF 9.5 Mio** of external funding for research activities were obtained by EOC researchers in 2022. The external funding increased compared to the previous year and it was assigned to both clinical research (64%) and translational research (36%).

Among the different EOC Institutes and Departments, IOSI, ICCT, INSI and LRT-EOC achieved the highest external funding for current ongoing research projects.

The funders were:

- **private foundations**
(34% of external funding and main funders of translational research),
- **industry**
(32% of external funding and main funder of clinical research),
- **European and international programs**
(9% of external funding),
- **government agencies**
(12% of external funding),
- **other Swiss programs**
(13% of external funding).



Research Highlights



Overall, EOC research activities in 2022 confirmed the upward trend in productivity and the high quality of the scientific outputs achieved in 2021.

- **779** peer-reviewed scientific articles were published in 2022, most of them as **original or review articles (89%)**.
- Both **clinical** and **translational** research are performed in EOC. The majority of the scientific output of EOC research in 2022 was related to clinical research.
- Peer-reviewed original and review articles were published in scientific journals showing very **heterogeneous Journal Impact Factors (IFs)** with a median value of **5.2**, slightly higher than in 2021. Notably, a total of **116 original articles and reviews (17%)** were published in **high impact factor** journals (Journal IF \geq 10) in 2022.
- Nearly **45%** of peer-reviewed original and review articles was published in scientific journals ranked in the **Journal IF quartile Q1**, confirming the high quality of the EOC scientific production as in previous years.
- Most of the peer-reviewed original and review articles (93%) was the result of **national or international collaborations**.
- The EOC researcher was the **first or last author in 40%** of the original articles or reviews, which means that several research projects were conceived and conducted at EOC.
- **115 new research projects** involving persons – **74 non-clinical** and **41 clinical trials** – were activated in 2022 and **342** research projects were ongoing at the EOC hospitals and labs in 2022.
- About **CHF 9.5 Mio** of external funding for research activities were obtained by EOC researchers in 2022. The external funding increased compared to 2021. IOSI, ICCT, INSI and LRT-EOC achieved the highest external funding for current ongoing research projects.

Complete list of publications



Publications are listed in alphabetical order. EOC authors are highlighted in bold.

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List of Publications for each Institute or Department



Oncology Institute of Southern Switzerland

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

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Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

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Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

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Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

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Institute of Pharmacological Sciences of Southern Switzerland

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Original articles

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Department of Medicine EOC

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Original articles

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Department of Gynecology and Obstetrics EOC

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

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Department of Anesthesiology, Emergency and Intensive Care Medicine EOC

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

- Amati M, Valnegri A, Bressan A, La Regina D, **Tassone C**, **Lo Piccolo A**, Mongelli F, **Saporito A**. Reducing Changeover Time Between Surgeries Through Lean Thinking: An Action Research Project. *Front Med (Lausanne)*. 2022;9:822964. IF 5.058, Q2, PMID [35573003](#)
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Nursing Research Centre
EOC

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

1. Botti S, Cannici C, **Liptrott SJ**, De Cecco V, Rostagno E, Gargiulo G, **Orlando L**, Caime A, Samarani E, Galgano L, Cioce M, Mordini N, Mandelli NE, Tombari L, Errichiello S, Celon N, Lupo R, Rea T, Serra N. Nurses' Physical and Psychological Symptoms During the first COVID-19 Lockdown in Italy: a Nationwide Cross-Sectional Study in Stem Cell Transplantation Setting. *Mediterr J Hematol Infect Dis.* 2022;14:e2022010. IF 3.122, Q3, PMID [35070217](#)
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Information and Communications Technology Division

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Original articles

1. Biasco L, Valotta A, Klersy C, Valgimigli M, Gabutti L, Bruna RD, Pagnamenta A, **Ruinelli L**, Senatore G, Pedrazzini GB. Association among myocardial injury and mortality in Influenza: A prospective cohort study. *Int J Cardiol.* 2022;369:48-53. IF 4.039, Q2, PMID [35944772](#)
2. Epistolio S, Ramelli G, Ottaviano M, Crupi E, Marandino L, Biggiogero M, Maida PA, **Ruinelli L**, Vogl U, Mangan D, Pascale M, Cantù M, Ceschi A, Bernasconi E, Mazzucchelli L, Catapano C, Alimonti A, Garzoni C, Gillessen Sommer S, Stefanini FM, Franzetti-Pellanda A, Frattini M, Pereira Mestre R. P1245 Polymorphic Variants of HSD3B1 Gene Confer Different Outcome in Specific Subgroups of Patients Infected With SARS-CoV-2. *Front Med (Lausanne).* 2021;8:793728. IF 5.058, Q2, PMID [35874037](#)
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5. Trimboli P, Camponovo C, **Ruinelli L**. The dilemma of routine testing for calcitonin thyroid nodule's patients to detect or exclude medullary carcinoma: one single negative test should be valuable as rule-out strategy to avoid further calcitonin measurements over time. *Endocrine.* 2022;77:80-5. IF 3.925, Q3, PMID [35391593](#)

Laboratories for Translational Research

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

1. Bignon Y, **Rinaldi A**, Nadour Z, Poindessous V, Nemazanyy I, Lenoir O, Fohlen B, Weill-Raynal P, Hertig A, Karras A, Galichon P, Naesens M, Anglicheau D, **Cippà PE**, Pallet N. Cell stress response impairs de novo NAD⁺ biosynthesis in the kidney. *JCI Insight.* 2022;7. IF 9.484, Q1, PMID [34793337](#)
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3. **Burrello J**, Caporali E, Gauthier LG, Pianezzi E, **Balbi C**, Rigamonti E, **Bolis S**, **Lazzarini E**, **Biemmi V**, Burrello A, Frigerio R, **Martinetti G**, **Fusi-Schmidhauser T**, **Vassalli G**, Ferrari E, Moccetti T, Gori A, Cretich M, **Melli G**, Monticone S, **Barile L**. Risk stratification of patients with SARS-CoV-2 by tissue factor expression in circulating extracellular vesicles. *Vascul Pharmacol.* 2022;145:106999. IF 5.738, Q1, PMID [35597450](#)
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25. **Vacchi E, Lazzarini E, Pinton S, Chiaro G, Disanto G, Marchi F, Robert T, Staedler C, Galati S, Gobbi C, Barile L, Kaelin-Lang A, Melli G.** Tau protein quantification in skin biopsies differentiates tauopathies from alpha-synucleinopathies. *Brain*. 2022;145:2755-68. IF 15.255, Q1, PMID [35485527](#)
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30. **Bersini S, Francescato R, Moretti M.** Biofabrication of 3D Human Muscle Model with Vascularization and Endomysium. *Methods Mol Biol*. 2022;2373:213-30. IF N/A, PMID [34520015](#)
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36. **Balbi C.** Editorial: Transverse aortic constriction-induced heart failure leads to increased levels of circulating microparticles. *Int J Cardiol*. 2022;348:109-10. IF 4.039, Q2, PMID [34915076](#)
37. **de Marco A, Barile L.** Good reasons for targeting SARS-CoV-2 by engineered extracellular vesicles. *Mol Ther Methods Clin Dev*. 2022;25:41-2. IF 5.849, Q2, PMID [35317050](#)
38. **Lionetti V, Barile L.** Fndc5/irisin-enriched extracellular vesicles: a new hormonal relay in the regular race against vascular ageing. *Eur Heart J*. 2022;43:4596-8. IF 35.855, Q1, PMID [36130321](#)

Clinical Trial Unit EOC

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

1. Bassi J, Giannini O, Silacci-Fregni C, Pertusini L, Hitz P, **Terrot T**, **Franzosi Y**, Muoio F, Saliba C, Meury M, Dellota EA, Jr., Dillen JR, Hernandez P, Czudnochowski N, Cameroni E, Beria N, Ventresca M, Badellino A, Lavorato-Hadjeres S, Lecchi E, Bonora T, Mattiolo M, Trinci G, Garzoni D, Bonforte G, Forni-Ogna V, Giunzioni D, Berwert L, Gupta RK, Ferrari P, **Ceschi A**, Cippà P, Corti D, Lanzavecchia A, Piccoli L. Poor neutralization and rapid decay of antibodies to SARS-CoV-2 variants in vaccinated dialysis patients. *PLoS One*. 2022;17:e0263328. IF 3.752, Q2, PMID [35143540](#)
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6. Leo M, Galante A, **Pagnamenta A**, **Ruinelli L**, Ponziani FR, Gasbarrini A, De Gottardi A. Hepatocellular liver injury in hospitalized patients affected by COVID-19: Presence of different risk factors at different time points. *Dig Liver Dis*. 2022;54:565-71. IF 5.165, Q2, PMID [35093272](#)
7. Low JS, Jerak J, Tortorici MA, McCallum M, Pinto D, Cassotta A, Foglierini M, Mele F, Abdelnabi R, Weynand B, Noack J, Montiel-Ruiz M, Bianchi S, Benigni F, Sprugasci N, Joshi A, Bowen JE, Stewart C, Rexhepaj M, Walls AC, Jarrossay D, Morone D, Paparoditis P, Garzoni C, Ferrari P, **Ceschi A**, Neyts J, Purcell LA, Snell G, Corti D, Lanzavecchia A, Velesler D, Sallusto F. ACE2-binding exposes the SARS-CoV-2 fusion peptide to broadly neutralizing coronavirus antibodies. *Science*. 2022;377:735-42. IF 63.714, Q1, PMID [35857703](#)
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13. Trimboli P, Camponovo C, **Ruinelli L**. The dilemma of routine testing for calcitonin thyroid nodule's patients to detect or exclude medullary carcinoma: one single negative test should be valuable as rule-out strategy to avoid further calcitonin measurements over time. *Endocrine*. 2022;77:80-5. IF 3.925, Q3, PMID [35391593](#)
14. Zaffina C, Wyttenbach R, **Pagnamenta A**, Grasso RF, Biroli M, Del Grande F, Rizzo S. Body composition assessment: comparison of quantitative values between magnetic resonance imaging and computed tomography. *Quant Imaging Med Surg*. 2022;12:1450-66. IF 4.630, Q2, PMID [35111638](#)

Other EOC Services

Publications are listed in alphabetical order for each section: original articles (including meta-analyses, surveys, research letters, consensus and delphi-based consensus paper); reviews (including systematic reviews without meta-analyses, protocols); editorials or letters or comments; case reports. Only EOC authors affiliated with the Institute/Department are highlighted in bold.

Original articles

1. **Amati M, Valnegri A, Bressan A**, La Regina D, Tassone C, Lo Piccolo A, Mongelli F, Saporito A. Reducing Changeover Time Between Surgeries Through Lean Thinking: An Action Research Project. *Front Med (Lausanne)*. 2022;9:822964. IF 5.058, Q2, PMID [35573003](#)
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7. Galiandro F, Agnes S, Moschetta G, Orlandi A, Clarke G, Bria E, Franceschini G, **Treglia G**, Giovinazzo F. Prognostic Factors in Patients with Breast Cancer Liver Metastases Undergoing Liver Resection: Systematic Review and Meta-Analysis. *Cancers (Basel)*. 2022;14:1691 IF 6.575, Q1, PMID [35406462](#)
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- Receptor-Positive Breast Cancer: A Systematic Review and Meta-Analysis. *J Clin Med*. 2022;11:1919. IF 4.964, Q2, PMID [35407526](#)
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