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Foreword

The University of Bern was founded in 1834 and has from the beginning included a Faculty of Medicine. In the last 185 years, many personalities formed and developed our Faculty, including Theodor Kocher, who was awarded the Nobel Prize in 1909 for his work on the thyroid gland. Theodor Kocher was the first surgeon and the first Swiss medical doctor to have received the Prize. Today the Faculty of Medicine offers a full program of study in Medicine and Dental Medicine. In addition, Master studies in Biomedicine, Biomedical Engineering and newly also in Pharmacy have been established. Our faculty is ranked among the top 100 medical institutions world-wide.

High-quality undergraduate teaching and close student - faculty contact remain central to our philosophy. The popularity of our teaching is reflected by the number of medical students enrolled which has increased continuously in the last decade and is the highest in Switzerland. Beginning in September 2018, we accommodated an additional 100 students in the first year of study; the total first year medical class now numbers approximately 340. In the last years, we have worked hard to optimally plan the new content of our teaching programs, as well as resources and logistics, which needed to be adapted to the increased numbers of students. We are happy to say that we continue to offer a problem-oriented study program, which demands a high proportion of well-trained teaching staff. Currently, we are newly planning the master study which requires additional efforts to accommodate all medical students in our clinical departments.

The Faculty of Medicine of Bern provides excellent opportunities for high quality research. Together with the Board of Directors of the Inselspital, our research strategy was updated in 2017. Basic research is carried out in the pre-clinical institutions (Anatomy, Physiology, Biochemistry and Molecular Medicine, Pharmacology, Pathology, and Infectious Diseases). Additional basic research is facilitated by the Department of Biomedical Research (DBMR), which offers all clinics of the Inselspital and UPD excellent opportunities, infrastructure and technical support for their work. Clinical research, including participation in large international trials, is performed with the support of the Clinical Trial Unit (CTU). Currently, we are in the process of developing a new Department of Clinical Research (DCR), which will accommodate the CTU as a service unit. The DCR will also develop its own research and teaching programs. It will be located in the Swiss Institute for Translational and Entrepreneurial Medicine (SITEM), in close proximity to the clinical departments of the Inselspital.

To promote patient-oriented clinical research, an innovative MD-PhD program in Clinical Sciences was started in 2017. Our Faculty gives strong support to the promotion of young researchers and to gender equality. For instance, we offer intramural grant opportunities (e.g. CTU grants, PhD student fellowships, protected research time).

Translational research programs in the fields of biomedical engineering were greatly energized by the creation of the unique Center for Artificial Organs (ARTORG) in 2008. Some ARTORG research groups were re-located in the SITEM. Other groups of the University and several units of the Inselspital also found space in SITEM in 2019 to promote collaborative interactions with partners of the industry. A good example here is the Diabetes Center Bern (DCB), in which new technologies are being developed to combat diabetes. Another strategic step to strengthen Translational Medicine was the decision to build up the Bern Center for Precision Medicine (BCPM) which has begun its work on 1 January 2019. Together with other local platforms (e.g. the unique Bern Liquid Biobank, several omics platforms, the Clinical Genomics Lab) and the University Hospital, this Center carries out research projects to develop the field of Precision Medicine and gives Bern a strong position within the Swiss Personalized Health Network (SPHN) initiative. These efforts require the development of additional expertise in the field of Computational Medicine. For this purpose, we are currently planning to create a new Center for Artificial Intelligence in Medicine (CAIM).

We hope that you will enjoy reading this report, which was prepared to promote familiarity with the breadth of knowledge and collaborations represented within and outside our University. We take this opportunity to thank all our employees and partners for their outstanding commitment which has made 2019 a very successful year for the Faculty of Medicine of Bern.

Hans-Uwe Simon
Dean
(on behalf of the Board of the Faculty of Medicine)
Highlights

January 2019

1.1.2019
In 2019, two internationally renowned research institutes in Bern merged: the Institute for Surgical Technology and Biomechanics and the ARTORG Center for Biomedical Engineering Research. Under the ARTORG brand, they form a national and international beacon for teaching, research and innovation in medical technology, in close connection with the Inselspital, Bern University Hospital, and the translational center sitem-insel AG.

March 2019

1.3.2019 – Diploma Celebration in the Münster of Bern, Prizes & Honours 2018
Faculty awards for the three best dissertations of the year 2018
1st price – Mr Dr. Georgios VASILAKOS for the Dissertation „Superimposition of 3D-digital maxillary dental casts on palatal structures: Method validation and clinical application“ under the direction of PD Dr. N. Gkantidis, Institute of Dental Medicine, Department of Orthodontics and Dentofacial Orthopedics, Bern
2nd price – Mrs Dr. Andrea Sara DULCEY-HUSI for the Dissertation „Feasibility, Safety and Outcome of Recanalisation Treatment in Childhood Stroke“ under the direction of Prof. Dr. M. Steinlin and Dr. S. Bigi, Department of Pediatrics, Inselspital Bern
2nd price – Mr Dr. Jonathan RÖCKEN for the Dissertation „The role of TNIK in non-small cell lung cancer (NSCLC) development and Wnt pathway activation“ under the direction of PD Dr. C. Riether and Dr. E.D. Bührer, Department for BioMedical Research, Research Group for Tumor-Immunology, University of Bern and Dr. M.A. Amrein, Department of Medical Oncology, Inselspital Bern
Eduard-Adolf-Stein-Preis of the University of Bern for an outstanding dissertation Mrs Prof. Dr. med. Lia BALLY
Recognition Prize of the Dental Society SSO For outstanding achievements in the study of dentistry (awarded at the Master's Ceremony on 13 September 2018) Mr Dr. med. dent. Fabrice DULLA
Prize of the Medical Society of the Canton of Bern For the best Federal examinations in human medicine 2018 Best Candidate for CS Examination – Mr Nicola Andrea BLASER Best candidate for the MC-Examination – Mrs Laura Barbara ANDEREGG
CSL Behring Award For Excellence in the Master program Biomedical Sciences Mrs Vera IMBODEN & Mrs Anastasia MILUSEV & Mr Emilio FLINT
Alumni Biomedical Sciences Award For the best Master's thesis in the Master program Biomedical Sciences – Mr Akira NOMURA

April 2019

12.4.2019 – André Schroeder Research Award 2019
The André Schroeder Research Prize 2019 goes to Prof. Dr. Vivianne Chappuis from the Department of Oral Surgery and Stomatology and her co-authors for their study “Osseointegration of ultra-fine titanium with a hydrophilic nano-structured surface: an in vivo examination on mini pigs”.

6.3.2019 – Visit of the Dean to Taiwan
Dean Simon stayed in March 2019 for a few days at the College of Medicine of the National Cheng Kung University in Taiwan. On the occasion of his stay, a Memorandum of Understanding was signed by the representatives of the two universities.

26.3.2019 – World’s first institute accredited for the detection of bacteria
The Institute for Infectious Diseases (IFIK) at the University of Bern is the world’s first accredited laboratory that can identify bacteria using nanopore sequencing.
18.9.2019 – AI-based prognosis in intensive care: decision-relevant patterns identified in EEG of coma patients

A reliable prognosis for coma patients in the intensive care unit is crucial. Improved transparency will boost the use of Artificial Intelligence (AI) to support end-of-life decisions. For the first time, a research team has succeeded in identifying specific patterns in Electro-Encephalogram (EEG) analyses that the deep-learning network uses for making prognosis decisions. This is an important step towards decoding the “black box” of deep-learning processes and thus towards AI transparency.

16.9.2019 – Start of studies 2019

On 16 September, 357 students (human and dental medicine) began their medical studies in the newly created Alhambra auditorium (formerly the Alhambra Cinema). This is already the second study start with 100 additional study places in human medicine since this was decided by the government council to counteract the shortage of doctors in Switzerland. This increase of +100 places makes the Medical Faculty of the University of Bern the largest in Switzerland and strengthens Bern’s position as a medical location.
23.10.2019 – Special cells contribute to regenerate the heart in Zebrafish

It is already known that zebrafish can flexibly regenerate their hearts after injury. An international research group led by Prof. Nadia Mercader of the University of Bern now shows that certain heart muscle cells play a central role in this process. The insights gained could be used to initiate a similar repair process in the human heart.

31.10.2019 – Removing liver tumors safely, noninvasively and efficiently

Many liver tumors have long been difficult or impossible to remove. Since 2015, however, it has been possible to treat these tumors by combining noninvasive surgical techniques, radiological imaging and a navigation system. For the first time, a new study by the University of Bern and Inselpital, Bern University Hospital has impressively demonstrated the success of this technique.

13.11.2019 – Day of BioMedical Research

This year’s Johanna Dürmüller-Bol DBMR Research Award of the Department for BioMedical Research (DBMR) of the University of Bern has been awarded to Maria-Nieves Sanz. She receives the award, endowed with CHF 30’000, for her research on inflammatory processes during heart transplantsations.

The Medical Faculty in Numbers

1’983 students in
2 Bachelor programs
4 Master programs
254 students in
30 CAS/DAS/MAS programs
313 Final Master Degrees
226 Medicine
36 Dental Medicine
28 MSc in Biomedical Sciences
20 MSc in Biomedical Engineering
3 Masters of Medical Education
107 Full Professors
17 Assistant Professors
134 Associate Professors
451 Lecturers
331 Doctorate Degrees
231 Dr. med.
33 Dr. med. dent.
60 from the Graduate School for Cellular and Biomedical Sciences
7 from the Graduate School of Health Sciences
184’436’845 Budget
29’039’492 from the Swiss National Science Fondation
44’834’102 third party funds for research
2’168 Original Articles published
136’878 hours of teaching
Glimpses of the History of the Faculty
Deans of the Medical Faculty

Historical
At the turn of the 20th century, Bern was famous for its quadruple of highly acclaimed medical professors: the surgeon Theodor Kocher (1841-1917), the internist Hermann Sahli (1856-1933), the dermatologist Josef Jadassohn (1863-1936), and the pathologist Theodor Langhans (1839-1915).

Whereas it is rather easy to determine some of the major scientific achievements of Langhans, Jadassohn and Kocher, it is much more difficult to establish the scientific contribution of Sahli. This is not due to a lack of Sahli's originality but due to his ideal of medicine in an era of increasing specialization.

The emergence of specialization
When Sahli, at the early age of 32, was elected as professor and head of the medical clinic in 1888, the buildings on the new hospital ground were just four years old. The whole hospital had a rather simple layout and was constructed around the two main clinics of surgery and medicine. Within a couple of years, this set-up was changed radically. Until 1910, new separate institutes and clinics of pharmacology, otorhinolaryngology, microbiology, dermatology, ophthalmology, radiology and pediatrics had been established. And a whole group of other specialties like neurology, cardiology and pneumology were calling for separate university teaching, professors and clinics.

This was not to Sahli's taste. In a 1914 paper on the relation between science and medicine, he stated that "in today's distractions and peculiarities of specialization, it is the duty of internal medicine to maintain the unity and simplicity of medicine. ... They have recently started to open their offices: heart physicians, lung physicians, nerve physicians, kidney physicians, stomach physicians, bowel physicians. What a brilliant idea! There is no objective justification for such separations. ... The human organism is a unity and it will not go unpunished if diagnosis and therapy are separated in such a manner according to organs."

Sahli bemoaned the tendency that patients would no longer go to their family doctor but directly to the specialist. This would be the reason why patients on the countryside were usually better off than those living in the cities. Such a development would neither be good nor necessary. Sahli stressed that the demands for a doctor's office could still be met with good education and experience. "And this will continue to be the case in hundred years when detailed knowledge will have increased tenfold." Sahli himself devoted the majority of his time to make this happening, to furnish physicians with the knowledge and the tools to become good GPs. He did so on three levels: by teaching, publishing and the invention of new diagnostic tools.

Holistic teaching
In Sahli's teaching, the presentation of the single case took center stage. He prepared each case meticulously and it was usually discussed for several hours. "Each clinical case is considered as a problem and scrutinized as far as possible with all the means of examination and reasoning." This would lead to an discussion of all the areas of knowledge involved, especially physiology and pathology – but also psychological aspects had to be taken into consideration. Even younger students with little clinical knowledge would participate. Sahli was, in fact, a champion of problem-based learning. In his view, there was no simple case, and routine was not something physicians should strive for. Due to the complexity and interconnection of the body there was no need for specialized clinical teaching in e.g. neurology. As many internal diseases were related to or based on neurological pathologies, this could be done within internal medicine. In addition to clinical cases, the theoretical lectures would be restricted to some selected difficult areas like metabolism or pathology of infectious diseases. He stressed that there is no simple medicine, there is only scientific medicine which students have to be immersed in.

A textbook: everything you need to know
True to his words, Sahli covered the whole range of internal medicine and – although an expert on blood pressure and circulation – published on such dissimilar topics as the development of antibodies, neurosis and hysteria, or insulin therapy. He criticized the increasing number of superficial, short-lived and irrelevant journal articles, arguing that World War One had at least one positive effect that not all the doctoral dissertations were printed anymore. As for himself, he announced many of his new findings only in his major work, the textbook on diagnostic methods. Published in seven increasingly voluminous editions from 1894 to 1928 and translated into English, Russian, Spanish and Italian, this is the work Sahli was mainly known for. He expected his students not only to use this massive textbook of almost 3,000 pages but to read it through carefully. In his view, diagnostic methods in his form of presentation, i.e. including physiological and pathological explanations, were "the basis of all our medical knowledge, thinking and skill."

Such a synopsis of modern medical and technical knowledge was necessary more than ever and would help to strengthen the diagnostic skill of the general physician.

Modernizing doctor's office: Sahli's hemometer
Situating diagnostic methods at the center of medicine and of his own scientific and clinical work, Sahli also developed a series of new methods and instruments himself, among other a manometer to measure blood pressure – and most famously – a hemometer to measure hemoglobin. This latter instrument was based on a colorimetric comparison of diluted and dissolved blood with a standardized liquid. Being a simple yet reliable tool it was the worldwide (and often imitated) standard model in the early decades of the 20th century and present in many offices until the late 1950s. Today, it is still widely used in primary health care settings in developing countries. Sahli's aim in producing such an instrument was to allow the physician to keep up with the advancement of medicine but still maintain his independence.

Although Sahli helped to strengthen the coherence of internal medicine, the process of specialization continued relentlessly. His ideal, however, of the unity of medicine and of the importance of GPs is still of great importance.
Deans of the Medical Faculty

1834–1835 Hugo Mohl
1835–1838 Hermann Askan Demme
1838–1843 Wilhelm Philipp Friedrich Vogt
1843–1847 Gabriel Gustav Valentin
1847–1849 Friedrich Miescher
1849–1852 Friedrich Wilhelm Theile
1852–1857 Wilhelm Rau
1857–1860 Wilhelm Philipp Friedrich Vogt
1860–1861 Wilhelm Rau
1861–1865 Gabriel Gustav Valentin
1865–1867 Daniel Jonquière
1867–1869 Christoph Theodor Aeby
1869–1871 Karl Emmert
1871–1874 Georg Albert Lücke / Henri Dor
1874–1876 August Breisky / Th. Kocher
1876–1878 Heinrich Irenäus, Quincke
1878–1880 Theodor Langhans
1880–1882 Ernst Pflüger
1882–1884 Ludwig Licheime
1884–1886 Marcellus Wilhelm von Nencki
1886–1888 Hugo Kromecker
1888–1890 Rudolf Demme
1890–1892 Ernst Pflüger
1892–1894 Hans Strasser
1894–1896 Hermann Sahli
1896–1898 Alexander Tschirch
1898–1900 Theodor Kocher
1900–1902 Arthur Carl Wilhelm Heffter
1902–1904 Theodor Langhans
1904–1906 Josef Jadassohn
1906–1908 August Siegrist
1908–1910 Wilhelm Kolle
1910–1911 Emil Bürgi
1911–1912 Leon Asher
1912–1914 Hans Strasser
1914–1916 Hans Guggisberg
1916–1918 Leon Asher
1918–1920 Carl Wegelin
1920–1921 Fritz de Quervain
1921–1923 Georg Sobennhein
1923–1924 Fritz de Quervain
1924–1926 Friedrich Lüscher
1926–1928 Emil Bürgi
1928–1930 Hans Guggisberg
1930–1932 Carl Wegelin
1932–1934 Walther Frey
1934–1936 Emil Bürgi
1936–1938 Paul Casparis
1938–1940 Jakob Klaesi
1940–1941 Hans Bluntschli
1941–1943 Alexander von Muralt
1943–1945 Eduard Glanzmann
1945–1947 Hans Goldmann
1947–1949 Joseph Dettling
1949–1951 Curt Hallauer
1951–1953 Karl Lenggenhager
1953–1955 Bernhard Walthard
1955–1957 Walther Wilbrandt
1957–1959 Erich Hintzsche
1959–1961 Walter Neuweiler
1961–1963 Hugo Aebi
1963–1965 Adolf Zuppingen
1965–1967 Franz Escher
1967–1969 Johann Kuske
1969–1971 Eugen Läppli
1971–1973 André Schroeder
1973–1978 Jürg Hooler
1978–1981 Beat Roos
1981–1983 Herbert A. Flesch
1983–1985 Harald Reuter
1985–1987 Rudolf Berchtold
1987–1989 Max Hess
1989–1991 Georg Esiner
1991–1993 Alfred H. Geering
1995–1997 Bernhard H. Lauterburg
1999–2004 Emilio Bossi
2004–2008 Martin Täuber
2008–2016 Peter Egli
2016– Hans-Uwe Simon
Organigram
Board of Faculty
Institutional Overview
Structural Development of the Inselspital Area
Dean’s Office
Dean’s Office of Student Affairs
Interview with Bernhard Pulver, Chairman of the Board of Directors of the Insel Gruppe AG
New Professors at the Medical Faculty
Institutional Overview

**Uni Mittelstrasse**
- Institute for Medical Education (IME), Mittelstrasse 43
- Institute of Primary Health Care (IPHC), Mittelstrasse 43
- Institute of Social and Preventive Medicine (ISP), Mittelstrasse 43

**Uni Muesmatt**
- Institute of Anatomy, Baltzerstrasse 2
- Institute of Biochemistry and Molecular Medicine, Bühlstrasse 28
- Institute of Forensic Medicine, Bühlstrasse 20
- Institute for the History of Medicine, Bühlstrasse 26
- Institute of Physiology, Bühlplatz 5
- Library Medicine, Baltzerstrasse 4
- Microscopy Imaging Center (MIC), Freiestrasse 1
- Theodor Kocher Institute (TKI), Freiestrasse 1

**von Roll Area**
- Institute of Complementary and Integrative Medicine (IKIM), Freiburgstrasse 46 & Fabrikstrasse 8

**Insel Area**
- Clinics and Institutes at the University Hospital, Inselspital
- ARTORG Center for Biomedical Engineering Research, Murtenstrasse 50
- Dean’s Office, Office of student’s affairs, Murtenstrasse 11
- Department for BioMedical Research (DBMR), Murtenstrasse 35
- Institute of Complementary and Integrative Medicine (IKIM), Freiburgstrasse 46 & Fabrikstrasse 8
- Institute for Infectious Diseases (ifi), Friedbühlstrasse 51
- Institute of Pathology, Murtenstrasse 31
- Institute of Pharmacology, Inselspital, INO-F
- Learning Center, Murtenstrasse 17
- School of Dental Medicine, Freiburgstrasse 7
- Swiss Institute for Translational and Entrepreneurial Medicine (sitem), Freiburgstrasse 18
- University Cancer Center, Freiburgstrasse 10
- University Neurocenter, Freiburgstrasse 16

**Teaching facilities**
- UniAlhambra
- UniZiegler

**UPD**
- University Psychiatric Services, Bolligenstrasse 111
Structural Development of the Inselspital Area

With the construction of a new training building on panel 03 of the Inselspital and a research and training building on panel 07, the first significant expansion and modernization of the spatial infrastructure is taking place and the foundation stone for the concentration of University medical training on the area of the Inselspital is being laid.

The central location of the organizational units of the Faculty of Medicine of the University of Bern on the Inselspital contributes on the one hand to strengthening the medical location of Bern, and on the other hand it promotes exchange in research and teaching and contributes to a practice-oriented learning and teaching environment.

Panel 03: the new base camp for medical students

In panel 03, the teaching and learning infrastructure for basic medical training for years 1-6 (Bachelor and Master) will be accommodated in the future. The main use of this training infrastructure will be teaching and seminar infrastructures (lecture halls, seminar and small group rooms), which will also be used for examinations, learning workstations for individual learning and learning in groups, a larger range of catering facilities, the Medical Faculty library and various organizational units responsible for the operation and management of the new training building.

The feasibility study for panel 03 was completed at the end of 2019, the next step is now the architectural competition. The commissioning of the new medical training building is planned for 2029.

Murtenstrasse 24-28: A medical research and service building

North of the Inselspital site, between Murtenstrasse and the railway line, and adjacent to the Inselparking, a modern laboratory building for the Medical Faculty is being constructed. From autumn 2021 onwards, it will serve as the site visit for the Institute of Forensic Medicine IRM, which is currently spread over seven locations. As one of four main sites, it will complement the research areas of the Department of Biomedical Research DBMR. In addition to accommodating research groups in large laboratories, the new building will allow the central services of the DBMR, the CoreFacilities, to be concentrated in one location.

The construction work is currently concentrating mainly on the interior fittings, while planning for the ELO project (Facility and Relocation Logistics) has started in parallel.

Replacement building for the Dental clinics (ZMK)

As already reported last year, a new replacement building for the zmk Bern is being constructed. The current location, the building at Freiburgstrasse 7, is already significantly overcrowded and will continue to reach its capacity limits as the number of students, patients and staff increases as well as the upcoming digitalisation. With the move of the research department (Dental Research Center, DRC) into the sitem-insel building, space has been created to optimize the zmk. An improvement in the space situation will provide short-term relief for users of the property at Freiburgstrasse 7. The selective conversion is expected to begin in summer 2020 and will take about one year.

The zmk Bern’s vision for the replacement building is to bring together the dental clinics and thus form the new service building on panel 07. The selective conversion is expected to begin in summer 2020 and will take about one year.

Panel 07: Construction of a research and training center on the Inselspital

The new research and training building in construction area 07 will house a modern laboratory infrastructure for research, seminar rooms and parts of the practical training laboratories. The new building will bring together a large part of the medical institutes (including all precinicsians from the Musematt) at one location, lead to better utilization of the infrastructure and create synergies with the Inselspital. The competition projects have been submitted and are in the preliminary examination stage.

Laying of the foundation stone for the main building of the Inselspital

The foundation stone was laid with a dignified ceremony on 2 July 2019. This represents a further milestone in the “Inselspital Masterplan” and the “Inselspital Areal III development plan”, which was adopted by a large majority of Bernese voters in 2015. The aim is to modernise and harmonise the Inselspital area. From 2023, the new building will replace, among other things, the high-rise bed block and thus form the new heart of the Inselspital.

With its digital planning and hospital building certified as Minergie-P-Eco, the complex new building is a true pioneering project. In addition to shorter distances, which should enable even more efficient processes, the well-being of the patients was the focus of the planning. In addition to central contact points and larger treatment rooms, they will also benefit from a pleasant atmosphere. A lot of daylight, views and bright and friendly materials were taken into account.
Interview with Bernhard Pulver
Chairman of the Board of Directors of the Insel Gruppe AG

Dr. iur. Bernhard Pulver is Chairman of the Board of Directors of the Insel Gruppe AG since 2019. From 1999 to 2006 he was a member of the Grand Council of the Canton of Bern and from 2006 to 2018 he was a member of the government of the Canton of Bern. From 2010 - 2011 he was President of the Swiss University Conference (SUK) and from 2015 - 2018 Vice-President of the Swiss University Conference (SHK). Since 2019 he is also the lecturer for Political Governance in the Master in Public Management and Policy (Master PMP) at the University of Bern.

Mr. Pulver, you are the Chairman of the Board of Directors of the Insel Gruppe AG since 1 February 2019. You know the Inselspital well from your previous work as a member of the government council. How did you notice this change? What experiences have you had in this first year?

I had a great start! I was very positively received and the positive expectations on my part were exceeded. I had a great start! I was very positively received and I see the enthusiasm in the clinics.

The enthusiasm in the clinics is certainly one of the main reasons that this contributes to a higher quality of care. The active involvement in the front line of research is a major difference compared to standard treatment. Exact, and this combination enables patients to receive treatment at the highest level. It is also our task to find interfaces where this interlocking can lead to a crunch in the gears. We find this, for example, in the dual employment of a medical doctor who is simultaneously employed by the University as a lecturer and researcher and by the Insel Gruppe as a service provider. This can lead to high stresses. However, I see this combination primarily as an opportunity from which we can benefit.

Now you have mainly talked about the opportunities and the friction surface. When the hospital is viewed as a business enterprise, the opportunities are emphasized, but do you also see dangers?

Swiss politics dictates that a hospital has to function like a business today. Hospital financing with DRGs (Diagnosis Related Groups) provides us with case-oriented income and we have to finance our costs and all investments – IT, construction and equipment – ourselves. In my opinion, the prevailing basic statement in Swiss politics today that the healthcare system is too expensive and that hospitals must therefore function like a business is in principle too general, but this basic assumption has had a huge impact on Swiss healthcare policy over the last 20 years. The balancing act is to achieve the necessary profit margin in an institution that must by no means function according to purely economic logic. After all, we are not a manufacturing company. It is about people and health, it is about people’s resources with relationships, prevention, aftercare. This is much more complex than simply a case that can be evaluated industrially. This is where I see the greater tension than in the combination of teaching, research and services. If we want to work in an entrepreneurial way and at the same time try to retain the human element, we could succeed in the “balancing act”.

If we now once again build a bridge to teaching and research in this environment. One statement I often hear is: “Research is something I do in my leisure time”. With this balancing act, how would you see future cooperation in terms of teaching and research? How do you think that should work?

When I visit clinics, they always present their research to me. I have not perceived this research activity as an activity in their free time. If you want to interpret this statement provocatively and take into account the long working hours of doctors, then it would indeed already be leisure time. For my feeling, the Insel Gruppe with its performance is not conceivable without this interlocking with teaching and research. We would not be where we are.

The key word is sustainability, which is on everyone’s lips. How do you interpret sustainability and ecology in hospital and research operations? We make great efforts in the area of sustainability.

Now you have mainly talked about the opportunities and the friction surface. When the hospital is viewed as a business enterprise, the opportunities are emphasized, but do you also see dangers?

Swiss politics dictates that a hospital has to function like a business today. Hospital financing with DRGs (Diagnosis Related Groups) provides us with case-oriented income and we have to finance our costs and all investments – IT, construction and equipment – ourselves. In my opinion, the prevailing basic statement in Swiss politics today that the healthcare system is too expensive and that hospitals must therefore function like a business is in principle too general, but this basic assumption has had a huge impact on Swiss healthcare policy over the last 20 years. The balancing act is to achieve the necessary profit margin in an institution that must by no means function according to purely economic logic. After all, we are not a manufacturing company. It is about people and health, it is about people’s resources with relationships, prevention, aftercare. This is much more complex than simply a case that can be evaluated industrially. This is where I see the greater tension than in the combination of teaching, research and services. If we want to work in an entrepreneurial way and at the same time try to retain the human element, we could succeed in the “balancing act”.

If we now once again build a bridge to teaching and research in this environment. One statement I often hear is: “Research is something I do in my leisure time”. With this balancing act, how would you see future cooperation in terms of teaching and research? How do you think that should work?

When I visit clinics, they always present their research to me. I have not perceived this research activity as an activity in their free time. If you want to interpret this statement provocatively and take into account the long working hours of doctors, then it would indeed already be leisure time. For my feeling, the Insel Gruppe with its performance is not conceivable without this interlocking with teaching and research. We would not be where we are.

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The key word is sustainability, which is on everyone’s lips. How do you interpret sustainability and ecology in hospital and research operations? We make great efforts in the area of sustainability.
And now there is still a new generation at the very beginning, among the students. The education of the students is actually our core task. From your position as board member, what is your message to the students? My message is that we are an exciting place to work. Working in our hospital is a highly meaningful task. And: We are a future-oriented industry - it is worthwhile to enter medical studies! With the “+100” project, we have increased the number of places in medical education by 100 students per academic year and have once again expanded pharmacy studies to full-time studies. This was an important concern for me at the time as Director of Education.

If we go one step further now, is that the same message for assistant physicians and senior physicians? For me it is the same message. It is not an ordinary job. It is a challenging job. Certainly, nobody becomes a doctor because they can have an easy job. I myself have great respect for this job because it involves dealing with people in difficult situations every day and demands an incredible amount of responsibility and flexibility.

This is probably why the willingness to perform is so high. Yes, that is enormous and at the same time certainly also a problem. When I ask the doctors how they can be on duty for so long on weekends, I hear that it is simply a calling for them as well. And when the work is finished, it often becomes even more interesting for them: now the patient is in the room, now they want to see how he is doing - and they don’t want to go home yet. I still don’t have an answer today as to how doctors can better manage their working hours.

That is a very difficult question. Maybe in a few years I’ll have an answer... (laughs)

We have already addressed the topic of the medical location two or three times. The Center for Precision Medicine started at the beginning of 2019, and the next major topic will be digitalisation and artificial intelligence. How do you see the cooperation between the University and the Inselspital in this area? Politically, the population, the Inselspital, the University and Bern want to strengthen this medical location. The Canton has stated in its 2030 commitment that it wants to be the number one medical location. The Insel Gruppe’s vision states: “We want to become one of the world’s leading hospital groups.”

At the moment we are in the process of building a leading medical location. With the +100 students we have expanded the Medical Faculty. With the siminsel, a translational research center has been created at the same time as the new Center for Precision Medicine. The ARTORIK already exists and we have the Insel Data Science Center. A Center for Design and Health is coming, which will explore the potential of design in the broadest sense for medicine. Or the topic of artificial intelligence - our position in research into artificial intelligence in medicine in Switzerland is good, probably even leading the way in Switzerland.

My vision for 2050 is that the campus will continue to expand beyond today’s perimeter, occupy more space, we will have even more start-up companies, and that even more research will take place. Digitalisation will be a core issue. As one of the leading hospitals worldwide, the Insel Gruppe has now defined this as a vision. Threads are being woven and ideas developed. We have a base where we are already excellent, and this is where we are building and want to improve.

This is also noticeable outside of medicine. It is incredibly great what was unanimously approved at the session in the Grand Council regarding the medical site1. As a former member of the government, I can say here that there have been few things that were approved so unanimously. Now I hope that we can still do a lot with it.

The area of digitalisation will be our next big topic. Within the next years, we plan to implement it in a structural form. From the University’s point of view, of course, the question remains open as to how firmly this should be integrated into the services of the Insel or how tightly it could be cross-linked. We are in the fortunate situation that we, as a University Hospital, have excellent cooperation with the Medical Faculty and the University. In artificial intelligence, knowledge is gained from an enormous amount of data. Here we can make progress by promoting research in the whole field of medicine, digitalisation and artificial intelligence. Here we as a medical campus - together with the canton - should also give financial support to start-ups. These will bring in innovation from the background of the companies. We are currently at the very beginning of discussions, in the preliminary phase with the question of financing. The University has the know-how about the interesting projects and should be involved in the selection of the start-ups.

You have said several times that the Inselspital is one of the leading hospitals and the largest medical location in Switzerland. We have achieved something in terms of power - is it also the best? Where are we already world class? I think we are way ahead in terms of knowledge and innovation, especially in terms of recently acquired developments. Take the Theodor Kocher Prize 2018, which Mrs. Kathleen Seidel received with her “radar system” for brain tumor surgery. This technique is now used in 40 countries! This is a great example, and I could go on and on.

If I may quote your statement on Dies Academicus: “The potential for shaping policy is often underestimated.” In your new role as Chairman of the Board of Directors, what do you want from politics? Politics can initiate economic and social developments. The Internet, for example, didn’t just come into being, but massive government research and funding was used, for example by the military in America. My expectations of politicians are to trigger further developments with the aim of becoming the number one medical location. Let me give you an example from another area: if Switzerland were to decide that no new cars with combustion engines would be registered after 2030, this would trigger something: Research, development and it will be good for the economy. The same applies to the medical location. This unanimous vote from the Grand Council sends signals to the economy.

May I finally ask you 1-2 personal questions? Do you possibly have an anecdote from your activity as Chairman of the Board of Directors? I have experienced many exciting moments. In Pneumology, for example, I was able to insert an endoscope into a doll and was shown how to get a candy out of the lungs in this way. Fortunately, it was a doll and not a human being! An impressive moment for me was when a professor showed me his clinic - it was Prof. Graff from Neuroradiology - and it started to flash on the screen during the visit as an indication that an emergency is coming in. He explained to me that a person with a stroke would arrive in 12 to 15 minutes and he would treat it. When I asked him if he had to leave immediately, he replied that we could finish the conversation. He then continued talking to me for the next 7 or 8 minutes in peace. Afterwards he went quietly to his mission. At such moments - there were several such moments during my visits - I am always speechless.

But also the visit to the finishing of the patient menus in the central kitchen was very impressive for me.

Your job is challenging and probably also very time consuming. There is certainly also a balance that you sometimes seek. The nice thing is that I don’t have to make so many operational decisions and therefore have a balance in time that is right for me. In my life, I have seldom had such a good time, purely from a professional point of view, as I have now here with such an exciting task and at the same time the time autonomy that I have. Professionally, I am very happy at the moment. Nevertheless, I need a balance, and for me this is in the form of going to a walk in the fresh air, at the Aare river, simply out and about, and in contrast to that at home, reading a book.

Thank you very much for the great interview, I am looking forward to the exciting future developments!

The interview was conducted by Dr. Lukas Stalder, head of the Dean’s Office.
Andrew Chan grew up in Hamburg and completed his studies in human medicine at the University of Hamburg in 1995. After his doctorate at the Institute of Clinical Chemistry of the University of Hamburg and his training as a specialist in neurology at the University of Würzburg, he habilitated in 2006 at the University of Göttingen on microglial immunobiology. Until 2016 he worked as senior physician and associate professor at the Ruhr-University Bochum, until 2016 he started his work in Bern.

Andrew Chan’s research focuses on neuroimmunological diseases, including multiple sclerosis. The research group focuses on mechanisms and markers for the optimization of immune therapies using different translational approaches.

Raphael Sznitman joined the ARTORG Center in 2014 as head of the Ophthalmic Technology Laboratory research group. Previously, he was a postdoctoral fellow at the Computer Vision Laboratory of the École Polytechnique Fédérale in Lausanne. He holds a Bachelor’s degree in Cognitive Science from the University of British Columbia (Canada) and a PhD in Computer Science from Johns Hopkins University (USA). As part of his research, Sznitman will establish the Artificial Intelligence in Medical Imaging group at ARTORG to develop applications in biomedical imaging, diagnostics, intervention, surgery and therapeutic decision making. Sznitman has successfully developed clinical applications in ophthalmology through artificial intelligence and machine learning, some of which are already in use at the Inselspital Eye Clinic.

Vivianne Chappuis grew up in Winterberg in the canton of Zurich. She completed her studies in dentistry at the University of Zurich in 1997 and subsequently worked in private practice for 4 years. In 2004, she completed her specialization as a dental specialist for oral surgery. During her two-year research stay from 2009-2011 at the Harvard School of Dental Medicine in Boston (USA) she investigated the molecular basis of bone regeneration. The project was supported by the renowned Musculoskeletal Transplant Foundation (USA). She received a Certificate of Advanced Studies in Entrepreneurship in 2014 and a degree in Business Administration in 2017. In 2018 and 2019 she was awarded the André Schroeder Research Prize in Clinical and Preclinical Research. As an oral surgeon, she is involved in teaching and research, education and training at national and international courses and congresses with live operations, workshops and lectures.

Her research focuses on translational research in the field of soft tissue and bone regeneration. It combines clinical and experimental research methods with modern digital 3D imaging with the aim of optimizing treatment concepts for patients. As a service institute and competence center, its clinic is the point of contact for dentists, physicians and patients with oral surgical, stomatological and radiological problems in the oral cavity.

The Executive Board of the University has elected Lukas Hunziker as Associate Professor of Heart Failure with effect from 1 November 2019. He succeeded Paul Mohacs. At the same time, he is the Chief Physician and Head of the Centre for Heart Failure at the University Cardiology Clinic of the Inselspital.

Lukas Hunziker (49) grew up in the Netherlands, Germany and Switzerland and graduated in human medicine from the University of Basel in 1995. After completing his doctorate, Lukas Hunziker completed the postgraduate course for basic immunological research at the Institute for Experimental Immunology at the University of Zurich. After a stay abroad at University College London, he habilitated at the University of Basel in 2006 in the field of internal medicine/immunology. In addition to his training as a FMH specialist in internal medicine, pneumology and intensive care medicine, he began training in invasive cardiology in Essen and at the University Clinic for Cardiology in Bern in 2013 and received a cardiology specialist in 2015. In 2016, he received his postdoctoral lecturing qualification in cardiology at the University of Bern. Since 2016 he has been a senior physician and since 2017 head of the Center for Heart Failure at the University Clinic for Cardiology of the Inselspital.

Lukas Hunziker’s research focuses on mechanical circulatory support and device based therapy/intervention of heart failure.
Prof. Campus graduated from dentistry a long time ago in Italy and spent some time in Jonköping (Sweden). Under the supervision of Prof. Goran Koch he began to work in the field of preventive medicine and oral epidemiology which was also the focused topic of his doctorate. Since 1996 he worked at the University of Sassari as Assistant Professor and then as Associate Professor of Public Health Medicine, Preventive Cariology and Paediatric Dentistry.

Prof. Campus is coordinator of the Italian WHO Collaborating Centre for Epidemiology and Community Medicine. For the WHO Collaborating Centre, his role is to develop and coordinate actions to assess the oral health of the public in Italy and other countries such as Mexico and Burkina Faso. He is also working to develop strategies and recommend measures to promote healthy behaviour from childhood onwards, including reduced sugar consumption, stricter tobacco controls and affordable fluoride toothpaste.

The plans of Prof. Campus are to evaluate and develop new strategies to collect epidemiological data and then generate reliable data for prevention measures. His opinion is that prevention will be the key to future dentistry.

Lorenz Räber grew up in Gelfingen, Canton Lucerne, and graduated from the University of Bern in 2004 with a doctorate in human medicine. After completing his training in internal medicine and cardiology, Lorenz Räber received a three-year scholarship from the Swiss National Science Foundation as part of the special program in University Medicine. From 2011 to 2012 he stayed at the Erasmus University Rotterdam and obtained a PhD in the field of intracoronary imaging. His habilitation followed in 2014 and his associate professorship in 2016. Clinically, he completed his training in interventional cardiology in 2013. In the same year, he took over the position of senior physician and from August 2015 the position of senior physician with head of the Department of Coronary Heart Disease at the Inselspital.

Lorenz Räber’s research focuses on the development, diagnosis and treatment of coronary heart disease, in particular with the aid of new methods such as optical coherence tomography or infrared spectroscopy. He is the author of over 180 publications and numerous book chapters as well as editor of the European Heart Journal.
School of Human Medicine
School of Dental Medicine
Local Student Committee of Bernese Medical Students
Start of studies 2019: „Firsties“ are coming!
Teacher of the Year 2018: Stefan Kuchen
New Auditorium Alhambra
New Medicine Library
Bachelor and Master Program in Pharmacy
Master Program in Biomedical Sciences
Master of Science in Biomedical Engineering
Physicians and medical doctors have been trained at the Medical Faculty in Bern for more than 200 years. The Bern Faculty is currently the second largest institution for the education of medical professionals in Switzerland.

Profile
- Training of physicians and medical doctors for more than 200 years
- 2nd largest institution of Switzerland for the education of medical professionals
- Comprehensive and balanced training concept
- High practical relevance and patient-orientated training
- Increasing the number of study places by 100 for the second time
- Bachelor program based on the educational approach of problem-based learning
- Master program offers subject-specific and practice-oriented study model with a high emphasis on bedside teaching in hospitals and in general practices
- Comprehensive training at the Inselspital in more than 50 disciplines
- Learning objectives according to PROFILES, which forms the basis of the Federal examinations

Start of studies in the new UniAlhambra
On 16 September 397 medical students, whereof 320 new and 28 repeaters in human medicine and 35 new and 14 repeaters in dental medicine started the first year of studies in the former cinema Alhambra, which has been transformed into UniAlhambra.

New information system platforms for teaching
In summer 2019, the “Studmed” site of the Medical Faculty retired after 20 years of use. The student office had to learn how to program a new IT system called Kernsystem Lehre (KSL). Students and teachers had to switch to the ILIAS application. The data migration and the use of the two platforms meant a lot of work, but the start was almost trouble-free.

ILIAS - the new teaching website

Clinical Skills Training (CST) in UniZiegler
The concept of Clinical Skills Training was completely revised. Many course modules now take place in the Renferhaus of the former Zieglerspital. The building has been renovated and there are now 4 seminar rooms, a simulator room, 41 examination rooms and a learning centre available. The OSCE examinations are also held in UniZiegler.

Federal exam Human Medicine 2019
Multiple choice exam: 230 candidates 230 passed 0% missing Clinical skills exam: 229 candidates 224 passed 2.2% missing

School of Human Medicine
Murtenstrasse 11, 3007 Bern
https://www.medizin.unibe.ch/studium/index_ger.html
The zmk bern is one of the few schools of dental medicine in the world offering a fully integrated interdisciplinary synoptic program.

The course of studies in dental medicine is based on the detailed national curriculum for dental medicine. Evidence-based treatment concepts ensure the high quality of dental care in Switzerland. The Bachelor and the Master programs in Dental Medicine at the University of Bern were accredited in December 2018 by the Rectors’ Conference of the Swiss Universities based on the recommendation of the Swiss Center of Accreditation and Quality Assurance in Higher Education.

After completing the master program, candidates are tested on their ability to correctly and independently perform interdisciplinary treatment on patients as well as on their theoretical and analytical competence. Furthermore, they have to present a Master thesis that can consist of either:

- an academic discussion of a topic relevant to dentistry
- an extended case study involving at least one case and relevant literature (evidence-based)
- a report of academic research (e.g. clinical trial protocol)
- a fully elaborated e-learning case.

Swiss national examination in dental medicine

Upon completion of the Master degree, candidates are required to pass the Swiss National Examination in dental medicine. The numbers of students and the male/female ratios are listed in the table below.

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Optimizations according to accreditation requirements

In 2018, the Accreditation Council issued the accreditation of the University of Bern’s Dentistry program with two conditions. In the accreditation report it was stated: “The zmk Bern must convert elective internships of the course into compulsory courses in such a way that all students are taught knowledge in complementary medicine, so that they are able to provide appropriate information about the risks and limits of complementary medicine methods in the expected patient discussions”. Fortunately, it was possible to implement this first edition from the autumn semester 2019 onwards. In the timetable of the 4th academic year a double lesson “complementary medicine” was introduced for the first time. This double lesson was successfully conducted by Prof. Dr. Ursula Wolf, Director of the IKIM (Institute for Complementary and Integrative Medicine).

The digitalisation of planning and treatment, which has been discussed for some time, was actively tackled. In the 3rd and 4th year of study, a joint module “digital impression” with theoretical and practical instruction was introduced by the clinics for tooth preservation and reconstructive dentistry and gerodontology. In the 5th academic year, the cross-clinic lecture series “Digital Reconstructive Technologies + Implant Dentistry” was established. In order to apply the newly acquired theoretical knowledge in the practical courses, an intraoral scanner, the corresponding grinding unit and the sintering furnace was purchased for the synopsis course. The second edition consisted of introducing a digital medical history in the student clinic. The deadline for fulfilling this requirement is the end of 2020. The tender documents for the new hospital information system (HIS) were prepared in 2019. The requirements for the new HIS were recorded in all clinics and student courses and were prepared in the specifications and requirements catalogue.

Bachelor Information Days at the University of Bern

Every year at the beginning of December, high school students in search of the ideal course of study have the opportunity to get to know various courses of study at the University of Bern within the framework of the “Bachelor Information Days” - On the occasion of these “taster days” the zmk bern also puts together a comprehensive program. In addition to general explanations about the study of dentistry and presentations from the various clinics, interested parties can get involved themselves by means of a learning circuit.

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Local Student Committee of Bernese Medical Students

The fsmb (Fachschaft Medizin der Universität Bern) is created by students for students, provides a link between the students and the faculty, contributes to the optimization of studies, has annual spokespersons in all study years, who are contact persons for the students and represent the academic year in the board of the student committee, acts as a mouthpiece for the interests of medical students in Bern, after a common reflection of a concern in the focus group with the annual spokesperson, an official feedback with proposals for solutions is submitted to the corresponding committee with the concern for open dialogue and cooperation, represents the interests of medical students in Bern in educational matters in various commissions (representation and right of codetermination), offers several assistance services for the study and events besides the study: Clinical language courses for French and Spanish, ski and sports weekends, stethoscope sales, the "MediLift", book exchange, a welcome aperitif for master students from Fribourg, fondue dinner, etc. (see homepage), its board is elected at the annual general meeting, interested students are welcome!

Focus on psychiatry: Swiss Medical Students’ Convention (SMSC) March 2019 in Bern

When suddenly 400 students with sleeping bags and sleeping mats under their arms, talking wildly in French, German, Italian and Rhaeto-Romance, streamed from Bern’s main railway station towards the VonKoll area, very few would have suspected this to be the future generation of Swiss doctors. Medical students from all over Switzerland came to Bern from 29 – 31 March 2019 to participate in the Swiss Medical Students’ Convention. This national congress takes place once a semester in rotation at the various medical faculties in Switzerland. This year’s organizing committee, consisting of around 20 committed medical students from Bern, had chosen psychiatry as its topic. Mr Graf’s lecture entitled “the therapy of the pedophile x” was particularly memorable. In one hour the speaker succeeded in describing the “personal microcosm” as a flock of birds and questioned the “free will”. In case studies, he showed us how this microcosm can lead to behaviour that is generally described as repressive. As a psychiatrist, however, it is important not to look at patients’ behaviour through the distorting mirror of one’s own perception and history and thus directly categorize patients’ behaviour. Rather, the goal should be for the patients themselves to realize where and how their thoughts and behaviour patterns have led to criminal acts. At two further lectures and numerous workshops, for example on the topic “Children from families with a mentally ill parent”, the participants of the SMSC received a lot of input on the topic of psychiatry during three days. On behalf of the entire organizing committee, we would like to thank the speakers, the home service of the VonKoll and Musmatt area, the Dean’s Office and the numerous sponsors for their support. All this would not have been possible without the commitment of many people, whose mention would unfortunately go beyond this framework.

National mental health campaign

In connection with the topic of the SMSC in Bern, swisma together with the VSS (Verband Studierendenschaften Schweiz) has launched a national campaign on the mental health of students in Switzerland. In the spring of 2020, swisma will conduct a national survey of all Swiss students, which will serve as an assessment of the current situation in order to introduce and promote evidence-based preventive measures to promote mental health at Swiss universities. In connection with this campaign, fsmb is organizing the mental health december in December 2019.

Before the intensive learning and examination phase, we want to give students the opportunity to deal with the topic of mental health. On the one hand with lectures on efficient learning techniques and mental health, but also with social events such as joint yoga exercises, games evenings in the Wokermensa and move nights in the former cinema Alhambra, which is now the Auditorium for Medical students of the first two years.

Medical Students international

As a founding member of IFMSA (International Federation of Medical Students’ Associations) in 1951, swisma (Swiss Medical Students’ Association) regularly takes part in the international meetings of medical students. These include the General Assemblies (GAs), which take place twice a year in March and August, and the European Regional Meeting (EuRegMe), which is held once a year in April. The Executive Board of swisma assembles a committee of motivated students for each conference, with representatives from all Swiss faculties.

EuRegMe April 2019 in St. Petersburg

Three Bernese medical students were selected for the EuRegMe in St. Petersburg from 20-24 April 2019: Given, Esma and Sebastian. During the congress, there were a variety of training opportunities, project presentations and workshops, where we were able to further educate ourselves together with the other international participants from all kinds of European countries. Through networking and the exchange of ideas, the new generation of doctors was inspired and our leadership and advocacy skills were developed. Gwen took part in a leadership training session that provided her with tools to explore her leadership style in a whole new way. On the Standing Committee for Human Rights and Peace, Sebastian learned about a Danish project to promote the health of refugees. We do not yet have any comparable projects in Switzerland. Esma visited the Standing Committee on Research Exchange and emphasized the opportunity of IFMSA meetings to inspire students: "IFMSA is an incredibly large community and every active person in it is committed to improving the lives of medical students. It’s important to be aware of the purpose and importance of the work you do." The complete swisma report on IFMSA EuRegMe 2019 Russia as well as an article on an alternative approach to travel (instead of using an airplane) “Hitchhiking to Russia” can be found at www.fsmb.ch.

March Meeting 2019 in Portoroz

One month before the EuRegMe the March Meeting of IFMSA took place in Slovenia. Four medical students from Bern were allowed to attend: Felice, Noemi, Alexander and Daniel. About 1000 medical students from all over the world came together in Slovenia. The delegation from Berne reported on the special feeling when borders are overcome and thus a cooperation at eye level and international cooperation becomes possible. In the Standing Committee for Public Health, they learned about antibiotic resistance, diabetes and malaria and the associated problems in different countries and continents.

Sebastian Albermann
President of the Local Student Committee of Bernese medical students
Since 16 September there have been new faces among the medical students in Bern - 355 firsties, or first-year students, to be exact. Here are a few insights into the life of the freshly baked students.

Why exactly did you come to Bern?
Yuk Yi: Actually Bern was not my first choice. In the beginning I was a bit negative, but now I think it’s pretty cool. The location is great because the Alhambra University is right next to the train station, and the city is very beautiful. I think it’s great that we have a lot of lecture-free time, so you can organize everything yourself, but you also need a lot of self-discipline. That’s what I’m still learning (laughs).

What was the first thing that crossed your mind when you first stepped into the dissecting room?
Maria: I was really happy! Everything was still covered and they made a huge secret of it until the preparations were uncovered. Then for the first time we really had something medical, before we only had chemistry, physics and biology. In the anatomy internship we really noticed: Okay, we’re studying medicine.
Elia: Shit, it’s cold in here!
Seraina: I was really surprised that we now have effective human preparations there. I hadn’t thought about that at all.
Yuk Yi: I was very much looking forward to it and had already talked to others about it. That’s why it wasn’t bad for me at all.

Is there something that’s bothering you about med school?
Aurèle: So far it’s a lot of repetition and very dry, I’m not used to that from studying biology before. But as soon as it goes into medicine, I find it very interesting.
Yuk Yi: The few WCs in the Alhambra! One spends the whole break waiting in front of the toilet and misses the beginning of the lecture because of that.
Maria: For commuters it is not easy that there are a lot of train delays and cancellations. And if you are late, you have to sit on the floor. And sometimes it’s quite cold in the lecture hall.

Which course has been the most exciting so far and why?
Elia: The anatomy course because it was the most exciting from a medical point of view. In addition, it best combined the theory learned so far with practice.

Do you find the concept with a lot of self-study and few lectures combined with learning groups and internships useful?
Benjamin: I think that solving tasks in learning groups is a good approach. So far, however, I have always had the impression that the lectures and learning groups are not coordinated with each other. One solved tasks in the learning groups and only weeks later a lecture was added, or vice versa. In biology and veterinary medicine last year, the tasks were effectively coordinated with the lectures, which is not the case here.
The interviews have been conducted by Florin Kalberer (1st year’s student).
Dr. Stefan Kuchen: “I will probably learn more than you.”

Teacher of the Year 2018: Stefan Kuchen

Dr. Stefan Kuchen was selected as Teacher of the Year 2018 and honored at the graduation ceremony in March 2019.

With the words of the German psychologist Wolfgang Prinz: “We know a lot but understand little” the first immunology lecture began in the 3rd year of human medicine. The first part is probably especially true in the time before the exams, the second part is also true, but it also claims its validity in the time before and after the exams - so actually during the whole semester. In order to counteract this, we were introduced to the traffic light system. Slides marked green contain the concept and are important for understanding and orientation. Red dots mark slides with important and exam-relevant content. And grey slides are for illustration purposes and do not contain any learning material.

When I asked him what the teaching meant to him, Mr. Kuchen replied that there was the psychological aspect for him that it was nice to teach interested students. He was also aware that not everyone would have the same enthusiasm for his subject, but that it would be all the nicer if he felt the students’ interest. On the other hand, it has a content aspect that can be particularly well understood when one has attended the Immunology in-depth seminar with Mr. Kuchen.

Mr. Kuchen sees the speed of medical development as a great challenge for the future. Both therapy and diagnostics are subject to rapid progress. Medical findings and knowledge are constantly increasing in quantity and complexity. But what always remains in this change is the patient with his needs, today as well as in the past. These have not changed and in order to do justice to them as new doctors, it is important never to lose sight of this. And in the flood of facts, mechanisms and principles to go back to the brain, to structure oneself as well as possible and not to lose orientation. Namely to want to understand and not only to know. This corrected version of the initial quotation is also based on the lectures of the Teacher of the Year.

The interview was conducted by Recha Suter (4th year student), former secretary of the Local Student Committee of Bernese medical students.

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The spatial infrastructure of the University of Berne is facing major challenges as a result of the strengthening of Bern as a medical location and the associated increase in the number of students in human medicine as part of the +100 project. The former Alhambra cinema in Bern was rented and converted into the UniAlhambra in order to accommodate all medical students of the same year in the lectures. The schedule for the implementation of the measures was “sporty” the conversion time was just under 3 months.

In the short time it was possible to preserve the character of the former cinema through targeted structural interventions and still create the transformation to a modern classroom. The “UniAlhambra”, the new auditorium of the Medical Faculty, has 364 seats and has been in full operation since autumn semester 2019.
New Medicine Library

The department library Bühlplatz (FBB) has become a pure medicine library.

By early concentration of the natural science subjects in the library Muesmatt as well as the development of the roll rack arrangement, place for the Medical Faculty and their student increase (project +100) was created in the library at the Baltzentrasse.

Thanks to the few but targeted measures, the medicine library is now shining in a new light and has been available to students, staff and visitors since the beginning of the 2019 autumn semester as a meeting place and place of learning.

Even among medical students, there is no “THE perfect learning workplace”, which is why the creation of a diversified range of learning locations was the focus. Thus, in the course of the project, the various learning landscapes were developed together with the students, which can finally be found in the new library of medicine. In addition, the redesign of the back wall and the clear zoning provided an appropriate and dignified setting for the open-book collection, in which it is a pleasure to browse through the bookshelves.
Profile
• Upon completion of the Master of Science in Pharmacy, the Federal Exam in Pharmacy has to be passed to have the right to work as a pharmacist.
• In close collaboration with institutes of the Medical Faculty (most prominently the BIHAM), clinics of the Inselspital, and practising pharmacists we aim at developing a high-quality, patient-oriented education of our future generations of pharmacists.

Start of the new 3rd year Bachelor in September 2019
In the first half of 2019, all vacant positions could be filled and lecturers recruited, and the detailed curriculum of the new 3rd year was finalized to complete the Bachelor program in Pharmaceutical Sciences at the Department of Chemistry and Biochemistry (DCB) at the Natural Sciences Faculty. The new full professors are Prof. Paola Luciani (DCB) for Pharmaceutical Technology, Prof. Francesca Paradisi (DCB) for Pharmaceutical Chemistry, and Prof. Jürg Gertsch (IBMM, Medical Faculty) for Pharmaceutical Biology. In September 2019, 31 students were enrolled in the new 3rd year.

Detailed planning for the start of the new Master in 2020 is ongoing.
We are on track to launch the Master course in Pharmacy in September 2020. The contents of the curriculum have been finalized, vacant positions for professors, lecturers and administrative support are advertised, the regulatory documents are under revision, and the budget 2020 has been approved.

Bachelor and Master Program in Pharmacy

Two new professorships for the Master course
The University has created and will finance a unique and innovative Professorship in Primary Healthcare in Pharmacy and Medicine at the BIHAM. This professorship will be shared by a pharmacist and a physician who together will develop a teaching and research program in interprofessional collaboration in primary healthcare. The election process is ongoing.

pharmaSuisse has agreed to provide initial funding for an Assistant Professorship in Clinical Pharmacy. The structure and election committee will commence the process in early 2020.

Bachelor and Master Program in Pharmacy

Administration Office Bachelor
Freiestrasse 3, 3012 Bern
Administration Office Master
Muntenstrasse 11, 3008 Bern

The new Brochure is now available at www.philnat.unibe.ch.
Swiss Academic Institutions and Research Companies have been faced with considerable problems when recruiting young biomedical scientists for many years. Having the goal to ease this situation, educational opportunities in life sciences and biomedicine have multiplied in recent years. Among these efforts, the Biomedical Science Studies of the Universities of Fribourg (Bachelor) and Bern (Consecutive Master) have played a pioneering role. Established in 2006, the harmonized curriculum of the two universities remains unique in Switzerland as it is focused on exposing the students to a translational teaching environment involving both basic and medical sciences. To achieve this goal, the first study year is dedicated to form a firm foundation in natural science while, in the second year, students share the curriculum of medical students thereby acquiring a comprehensive knowledge in human anatomy, physiology and biochemistry. During the third and the fourth year, students participate in a systematic introduction to the pathophysiology of all organ systems with lecturers being associated with basic research institutions, the university hospital and pharmaceutical companies. The Master Thesis is conducted in a laboratory of choice and includes collaborative projects with the industry. Based on the specific design of the curriculum, graduates in biomedical sciences have an exquisite and broad portfolio of both knowledge and skills at the interface between basic sciences and clinical research that endorses them to engage successfully in basic, translational and clinical research including emerging new research fields in human medicine like artificial intelligence and precision medicine.

**Profile**
- Direct admission with a BSc in Biomedical Sciences of the University of Fribourg or a Bachelor in Human Medicine
- 1 1/2 year full time study program
- Human pathophysiology is lectured by basic researchers and clinicians
- Courses include practical work in research laboratories and training in modern experimental techniques
- 2 three-week laboratory internships in research fields chosen by the students
- Opportunities for conducting the master thesis in the industry
- Broad portfolio of systematic knowledge and skills at the interface between basic sciences and clinical research
- Graduates are in high demand for doctoral positions in academia and for research positions in the industry

**Figures**
- Diplomas since 2011: 178
- Presently enrolled students: 34
- Gender (f/m): 3/1
- 22 graduates in biomedical sciences in 2019

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**CSL Behring Prize 2019**
The CSL Behring Prize 2019 for the best Master degrees went to:
1st rank: Imboden Vera *5.82*
2nd rank: Milusev Anastasia *5.70*
3rd rank: Flint Emilio *5.62*

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**Alumni Prize 2019**
The prize for the best Master Thesis 2019 sponsored by the Alumni Organization went to Akira Nomura for his study on “Deciphering the importance of caveolin-1 for cardiac electrophysiology.”

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**ABS Animal Science and Ethics Day**
One-day event with talks, keynote lecture and workshops on animal science and ethics that drew 70 participants from the Universities of Bern, Zurich, Lausanne, Geneva and Neuchatel.
The master's program in Biomedical Engineering is being presented at the opening of the new Sitem-Insel building in August of last year.

Master of Science in Biomedical Engineering

The master's program in biomedical engineering is a full-time study program offered in cooperation with the Bern University of Applied Sciences. It aims at training multidisciplinary engineers to deliver scientifically-founded, sustainable and cost-effective solutions for biomedical problems in academia, medical care and industry.

Profile
• Admission with BSc (FH/HES/SUP/Univ/ETH)
• International program in English
• Affiliated to a leading medical faculty hospital (Inselspital)
• Two-year full-time program but compatible with 40% working time
• Oriented towards clinical applications
• Attractive, central location

Figures
• 148 students enrolled in academic year 2019/20
• 59 regular and 10 exchange or guest students new in 2019
• 25% of 2019 new students female
• 28 biomedical engineers graduated in 2019
• with excellent career perspectives

Swiss Engineering Master Thesis Awards 2019
Jan Stapelfeldt received the award for innovation for his thesis “Smartphone-Based Perimetry for Fast and Low-Cost Visual Field Acquisition”. Michael Rebsamen, was winner of the award for basic science for his work “Fast and Accurate Human Brain Morphometry Estimation with Deep Learning”. The awards were presented by Valentin Herbez from Swiss Engineering.

The Biomedical Engineering Day
The Biomedical Engineering Day is our annual BME career day and networking event. About 350 participants followed a broad and balanced speakers’ program, which included highlights such as a live surgery by Prof. Siebenrock and a science slam. In addition, all research groups from the ARTORG Center and the HuCe, Bern University of Applied Sciences, as well as Swiss MedTech companies and research institutes presented themselves at a large exhibition with 29 booths.

RMS Award 2019
Michael Rebsamen received the RMS Award 2019 from Philippe Zysset, Program Director Master Biomedical Engineering. Each year, the Robert Mathys Stiftung (RMS), an independent service laboratory and research institute, offers it to the BME student with the highest average grade.

Sitem-Insel Open Doors
This summer, the Master’s program in Biomedical Engineering moved to Sitem-Insel, the Swiss institute for translational and entrepreneurial medicine, located next to the university hospital. Sitem Open Doors at the end of August gave us the opportunity to increase public awareness for our unique program.

Retirement Professor Lutz Nolte
In January, Professor Lutz Nolte, the founding father of our master’s program in biomedical engineering, retired after 25 years of distinguished service as a faculty member at the University of Bern. Many colleagues and friends from Switzerland and all over the world joined us for a symposium in honour of his extraordinary work.
Promotion of young academics

- stem-insel School for Translation and Entrepreneurship in Medicine
- CAS, DAS and MAS Degree Programs
- PhD Education
- Grants
- Commission for Equality
Symposium «Bringing Innovation to the Patient»
The new lecture hall at sitem-insel was crowded when people from industry, public offices and academia got together at the sitem-insel School Symposium «Bringing Innovation to the Patient». Keynote speaker Dr. Hans-Juergen Wildau, Biotronik, Berlin gave a fascinating talk on "Intelligent Digital Health".

sitem-insel Open Doors
On the occasion of the sitem-insel open doors, guests were introduced to the concept of the translational process in a playful way. In a ladder game, the players virtually brought a product from the idea to the patient. At the various posts of the game, the players became familiar with the individual stages of the translational process and were able to learn more about its challenges and opportunities by drawing cards.

Medical Device Regulatory Affairs and Quality Assurance (MAS / DAS)
The introduction of the new Medical Device (MDR) and In-vitro Diagnostic Medical Device (IVDR) regulations has created an enormous demand for further training in the Swiss medical technology industry. From September 2020, the sitem-insel School plans to offer advanced studies to train specialists in "Regulatory Affairs". Graduates of the program will be able to carry out the approval of medical devices in cooperation with certification authorities ("Notified Bodies"). The course is currently in planning.

Translational Medicine and Biomedical Entrepreneurship (MAS / DAS / CAS)
7 start-ups and 6 companies in the incubator phase are trained in the sitem-insel School personalized "Translational Medicine and Biomedical Entrepreneurship" continuing education, which includes topics such as R&D, quality management, regulatory affairs, clinical trials, biomedical entrepreneurship and others.
For more information see https://sitem-insel.ch/tmbe

First national phageSuisse Symposium
phageSuisse has organized on 28.11.2019 its first symposium in collaboration with sitem-insel School and could welcome over 50 participants at the sitem-insel building. The need for novel and innovative phage treatments for resistant bacterial infections and the challenges in bringing them to the patients were discussed.

sitem-insel School for Translation and Entrepreneurship in Medicine
sitem-insel School aims to promote researchers and train specialists in the fields of translational medicine and biomedical entrepreneurship. Lecturers and supervisors of the school are representatives from research and development-oriented private companies, scientists from universities, clinicians, collaborators from regulatory agencies, and financial experts.

Profile
The sitem-insel school
• aims to teach students practice-oriented high-quality knowledge
• aims to facilitate a strong network between people involved in the translational process
• offers personalized continuing education using blended learning (presence lectures & e-learning).

Figures
• Three continuing education programs:
  – Translational Medicine and Biomedical Entrepreneurship (MAS/DAS/CAS)
  – Artificial Intelligence in Medical Imaging (CAS, since Sept. 2019)
  – Medical Device Regulatory Affairs and Quality Assurance (MAS/DAS in planning, starting September 2020)
• 7 start-ups and 6 companies in the incubator phase participate in the sitem-insel School continuing education
• The sitem-insel continuing educations offer excellent career perspectives

Symposium "Artificial Intelligence in Medical Imaging"
The CAS “Artificial Intelligence in Medical Imaging” started with an international symposium at sitem-insel on 9 September 2019. In this CAS, physicians are brought together with data scientists and specialists in AI to help shape digital technologies at the forefront. For more information see https://sitem-insel.ch/ai.

sitem-insel School
Freiburgstrasse 3
www.sitem-insel.ch/school
CAS, DAS and MAS Degree Programs

The Medical Faculty offers over 30 programs of advanced studies. All programs are held by teachers of the faculty who transmit practice-oriented and state of the art knowledge. They are addressed to professionals with a tertiary education.

Existing advanced study programs
- Experimental and Translational Nephrology (CAS/DAS ETN Unibe)
- Hepatology (CAS/DAS HEP Unibe)
- Sleep, Consciousness and Related Disorders (CAS, DAS, MAS SCD Unibe USI)
- Clinical Nutrition (CAS)
- Spiritual Care (CAS SpC Unibe)
- Interprofessional Specialist Palliative Care (CAS PallCare Unibe)
- Swiss Exercise Therapy in Sports and Medicine (CAS/DAS SwissETSM Unibe)
- Swiss Cardiovascular and Diabetes Therapy (CAS SwissCDT Unibe)
- Exercise and sports therapy for mental illness (CAS PSY Unibe)
- Exercise and sports therapy in orthopedics, rheumatology and traumatology (CAS ORT Unibe)
- Public Health (CAS/DAS/MAS)
- Leadership in Health Care Organisations (CAS LHCO Unibe)
- Clinical Research in Health Care Organisations (CAS CRHO Unibe)
- Managing Medicine in Health Care Organisations (CAS MMHC Unibe)
- Translational Medicine and Biomedical Entrepreneurship (CAS/DAS/MAS Unibe)
- Master of Medical Education (MME Unibe)
- MAS in Leading Learning Health Care Organisations (MAS LLHC Unibe)
- Master of Advanced Studies in Implant Dentistry (MAS IMPUnibe)
- Master of Advanced Studies in Orthodontics and Dentofacial Orthopedics (MAS ORTHO Unibe)
- Master of Advanced Studies in Cardiology, Endodontology and Pediatric Dentistry (MAS REST Unibe)
- Master of Advanced Studies in Oral and Implant Surgery (MAS ORALSURG and IMPSURG Unibe)
- Master of Advanced Studies in Periodontology and Implant Dentistry (MAS PER and IMP Unibe)
- Master of Advanced Studies in Reconstructive and Implant Dentistry (MAS REC and IMP Unibe)

All study programs can be found on the website: www.medizin.unibe.ch/wiederbildung/cas___das___mas/index_ger.html

New advanced study programs

The main objectives of the Certificate of Advanced Studies (CAS) in Artificial Intelligence in Medical Imaging are to equip medical doctors (MDs) with the necessary skills to take a leading role in the AI-driven transformation of medicine. The course program was developed for medical professionals and provides them with the necessary knowledge and skills to translate medical problems into data science problems. By working on an AI project of their own, the participants will know and apply the fundamental concepts of Artificial Intelligence and can thus judge the feasibility and adequacy of proposed AI solutions.

The CAS program in gender-specific medicine starts in May 2020 and is a part-time university continuing education program, jointly offered by the Universities of Bern and Zürich. It is aimed at people with a Master’s degree in medicine or a related field who wish to deepen their knowledge of gender differences in medicine in order to shape their work according to the latest evidence. It teaches the concepts of so-called gender medicine and the effects of biologically and socio-culturally shaped gender on health and health care.

In the individual modules, the significance of gender in the various medical disciplines is shown. Tools, concepts and ideas are presented on how both health care and medical research can do justice to the sexes.

CAS in Artificial Intelligence in Medical Imaging (CAS AIMI Unibe) for Outpatient Neurology

CAS in Sex and Gender Specific Medicine
PhD Education

The Medical Faculty is associated with two graduate schools offering structured doctoral programs: the Graduate School for Cellular and Biomedical Sciences (GCB) and the Graduate School for Health Sciences (GHS).

Both graduate schools offer research-oriented curricula with a wide choice of courses and special workshops tailored to the individual interests and needs of PhD candidates. The focus is on the research projects for which the students get the best possible support from an expert supervisory committee in their specific research area. The emphasis is on high-quality training in research methods and study design. The PhD programs direct the candidates towards independent scientific work and enable them to assume scientific responsibility.

Profile of the Graduate School of Cellular and Biomedical sciences GCB

The GCB offers training in experimental research with state-of-the-art methods in molecular life sciences, biomedical sciences and biomedical engineering. Research areas include Cell Biology, Biochemistry, Molecular Biology, Immunology, Genetics, Biomedical Sciences, Epidemiology as well as Tissue Engineering and Computer-Assisted Surgery. There are two areas of specialization:
- Cutting-Edge Microscopy
- Stem Cell Research in Regenerative Medicine

The requirement for the program is a master’s degree in Biology, Molecular Biology, Immunology, Genetics, Biomedical Sciences, Epidemiology and related fields depending on the project.

There are five expert committees for the PhD program structure:
- Biological Systems
- Biomedical Sciences
- Cell Biology
- Molecular Biology/Biochemistry
- Biomedical Engineering

Each candidate is supported by a thesis committee consisting of a supervisor, a co-advisor and a member of an expert committee as a mentor. The curricula of the MD-PhD program for medical doctors focuses on basic sciences, but the PhD candidates can spend 20% of their time in the clinic. Graduates receive one of the following titles:
- MD-PhD (Doctor of Medicine and Philosophy)
- DDS-PhD (Doctor of Dentistry and Philosophy)
- DVM-PhD (Doctor of Veterinary Medicine and Philosophy)

The MD-PhD studies are supported by competing scholarships of the SNSF and SAMW.

Profile of the Graduate School of Health Sciences GHS

The GHS offers a research-oriented curricula on psychological and physiological factors that determine the health of individuals and groups in their social contexts and physical environments.

The requirement for the program is a master’s degree in Psychology, Medicine, Biomedicine, Epidemiology, Geography, Sport Science or other fields depending on the respective research project.

There are three expert committees for the PhD program structure:
- FKI: Preventive and Social Medicine, Public Health, Medical Education, Psychology, Rehabilitation and Patient-Related Studies
- FKII: Clinical Sciences

Each candidate is supported by a thesis committee consisting of thesis advisor, co-referee and a member of an expert committee as mentor (only in FKII). Awarded PhD title is:
- PhD in Health Sciences (specialist area) Curricula for medical doctors
- PhD in Clinical Sciences: 50:50 model: Patient-oriented research and clinical career.
- Graduates receive the title: PhD in Health Sciences (Clinical Specialty)

Starting 2019, the GCB & GHS have moved to the Mittelstrasse 43 and have new coordinators: Monica Schaller and Tullia Padovani have taken over the coordination from Marlene Wolf, who retired at the end of January 2019.

Highlights, important events at GCB

- New promotion regulation effective from 1 February 2020 for ALL students.
- The following are the new mentors at GCB:
  - Prof. Selina Aybek
  - PD Michaela Medova PhD
  - PD Dr. Andrea Schueren Stehli
  - Prof. Hildegard Tanner
  - Best PhD degree: Annina Bauer, Vetsuisse Bern

Highlights, important events at GHS

- New GHS brochure
- 50% students increase from 2018 to 2019
- The newFKII, Clinical sciences raises significant interest thanks to the 50:50 model
- GHS Symposium in Gunten
- The following are the new expert committee members at GHS:
  - Prof. Jennifer Inauen
  - Prof. Stephanie Schmidt
  - Dr. Cinzia del Giovane

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Starting 1 February 2020 for ALL students.
Grants

**Graduate School in Health Science (GHS)**
- 3 options: Health Sciences, Cognitive Neurosciences and Clinical Sciences
- Career stage: young MD's (open to other health professionals, psychologists, biologists, engineers)
- Duration: 3 – 5 years
- Number of calls per year: for Clinical Sciences 1x per year, otherwise continuously
- Funding source: Medical Faculty

**Graduate School for Cellular and Biomedical Sciences (GCB)**
- Career stage: biologists, engineers, MedVet, MD's
- Duration: 3 – 4 years
- Number of calls per year: continuously
- Funding source: Medical Faculty

**CTU Grants**
- Career stage: MD's before Habilitation
- Duration: 1.5 year
- Number of calls per year: 2x
- Funding source: University Hospital

**Uni Bern Initiator Grant**
- Career stage: postdocs and young PI
- Number of calls per year: 2x
- Funding source: University
- Additional information: open to all disciplines; support for proposals for third-party funded career schemes (SNF mobility fellowships, Ambizione, SNSF-professorship; EU Marie-Curie-S. fellowships, ERC grants, etc)

**Numerous Foundations**
- Career stage: during BSc or MSc
- Duration: 1 – 5 years
- Number of calls per year: 1x
- Funding source: Foundations
- Additional information: open for students registered at the University of Bern in academic years 2 - 6

**Travel Grants**
- Career stage: during BSc or MSc
- Duration: 1 – 6 years
- Number of calls per year: continuously
- Funding source: Dean’s office
- Additional information: open for students registered at the University of Bern in academic years 1 - 6

**Protected Research Time**
- Career stage: Doctoral Students at the Graduate School for Health Sciences, Senior Physicians
- Duration: 2 – 4 years
- Number of calls per year: continuously
- Funding source: Dean’s office
- Additional information: open for doctoral students at the GHS or Senior Physicians with an academic degree, working at least 6 months at the Insel hospital, UPD or Service Institute of the University. >40 years old and not habilitated
- Women are explicitly encouraged to apply
Commission for Equality

The Medical Faculty Commission for Equality (Innerfakultäre Kommission für die Gleichstellung, IFKG) supports the Medical Faculty in realizing measures ensuring effective equality of women and men. Together with the decision-makers of the institutes and clinics, the IFKG aims to improve conditions that facilitate the decision for women to pursue an academic career.

The immediate aim of the Commission for Equality is an increase of the number of women in leadership positions in the Medical Faculty. Among other measures, the IFKG aims to illustrate career options that improve working conditions such that they allow both women and men to combine their family live with their academic career pathway.

Profile
- Mentoring 4 Women
- Ensuring implementation of equal opportunities in search committees and in the work environment of the Medical Faculty
- Contact point for equal opportunity and diversity topics
- Contact point in case of discrimination
- External partners: Coordinator of the Office for Equality of the University of Zürich; Coordinator of the Office for Equality of the University Hospital of Basel

Successful implementation of the Equal Opportunities Action Plan 2016–2019

All measures outlined in the Gender Equality Action Plan 2016–2019 were successfully implemented. We would like to mention the following points in particular:
- Implementation of gender balance in courses discussed in problem-based learning with medical students during their first years of education
- Initialization of quarterly strategic meetings between the University Bern, the Board of the Medical Faculty, the Board of the Inselspital Bern and the Medical Faculty Commission for Equality
- Equality checklist for search committees
- Implementation of consideration of unconscious biases as well as the DORA regulation in recruitment procedures
- Implementation of novel measures for the promotion of young talents
- Further measures are taken to enhance the number of women in academic careers

Workshop in the «Haus der Universität»

The Medical Faculty Commission for Equality organized a workshop in June with female physicians pursuing an academic career at the Inselspital. The specific aim was to identify with this target group the obstacles prohibiting their academic career advancement. The following 3 focal points to improve attractiveness of an academic career at the Inselspital.

- Protected research time, networking and structuring possibilities
- Mentoring from male physicians pursuing an academic career
- Further trainings, speeches and equality-related topics

The mentees can furthermore benefit from a vast list of available mentors. We have continuously increased the number of mentees from mentee programs of the other faculties of the University of Bern. The mentees can furthermore benefit from a vast list of available mentors. We have continuously increased the number of mentees from mentee programs of the other faculties of the University of Bern.

M4W – Mentoring for Women

We continuously work on improving our Mentoring for Women (M4W) program. We have introduced an initial interview with the applying mentee to optimize finding a suitable mentor. We have continuously increased the list of available mentors. All mentees are regularly informed about interesting further trainings, speeches and equality-related topics and events. The mentees can furthermore benefit from a vast faculty cooperation and can attend interesting courses from mentee programs of the other faculties of the University of Bern.

The M4W program is highly appreciated as underscored by select statements from our mentees:
- For me this is a very helpful program and provides a great opportunity to have an independent advisor. For the moment, I cannot provide any further ideas for improvement, as I am very happy with the program.
- This is exactly why programs like the M4W are important. This is about moral support and the role model function of the mentor.
- My experience continues to be extremely positive, I feel extremely supported and always very well advised. The conversations with my mentor are helpful, trending and enriching for me on a human level.

Quarterly Conversations

Every three months, the Vicerector Quality from the University of Bern, Members of the Board of the Faculty of Medicine and the Inselspital meet together with the President of the Medical Faculty Equality Committee to improve equal opportunities in the Medical Faculty and at the Inselspital.

Several important activities have already resulted from these meetings. Last June the workshop with female physicians pursuing an academic career at the Inselspital took place. The possibility of setting up a day nursery at the Inselspital has been identified and is now being examined. The idea of creating a networking platform for female physicians pursuing an academic career at the Inselspital is also followed up.

We are pleased about the positive development in the proportion of women in academic careers

Commission for Equality
Murtenstrasse 11, 3008 Bern
www.medizin.unibe.ch/ueber_uns/gleichstellung/index_ger.html
Centers and Platforms

ARTORG Center for Biomedical Engineering Research
Bern Center for Precision Medicine (BCPM)
Department for BioMedical Research (DBMR)
Microscopy Imaging Center (MIC)
Swiss Institute für Translational and Entrepreneurial Medicine (sitem-insel)
University Cancer Center (UCI)
University Neurocenter
Clinical Trial Unit Bern (CTU)
ARTORG Center for Biomedical Engineering Research

Murtenstrasse 50, 3008 Bern
www.artorg.unibe.ch

Prof. Raphael Sznitman
ARTORG

Prof. Stefan Weibel
IGT

Prof. Philipp Zysset
MBB

Prof. Tobias Naf
GER

Prof. Oliver Guentet
ODC

Prof. Dominik Obrist
CVE

Prof. Stevanu Muijlakakis
AIHTI

Profile

- Interdisciplinary medical technology research hub between the University of Bern, Inselspital, Bern University Hospital, and Industry
- 11 independent research groups, 5 of them at sitem-insel
- Teaching profile: The center mainly contributes to the Masters program in Biomedical Engineering. The postgraduate program currently has 55 full-time PhD students at the interface between technical and medical disciplines.
- Key collaboration partners: Inselspital; UPD; BFH; Imperial College London, UK; University of Nottingham, UK; University of Oxford, UK; Rutgers University, USA; University of British Columbia, Canada; Helmholtz Center, Germany; Universitätsklinikum Freiburg, Germany; University of Stuttgart, Germany; Vienna University of Technology, Austria; EMPA, of Oxford, UK; Rutgers University, USA; University of British Columbia, Canada; Helmholtz Center, Germany;

Grants

- Swiss National Science Foundation, SNSF (grant No. 163800; 165510; 173130; 176007; 176498; 182966; 182975; 183584; 185265; 188140; 188823; 190521)
- SNSF Sinergia: 183584; CRSII5_189972 / 1; SNSF BRIDGE: 176498; 184179; 183584; 185365; 188140; 180822; 190521)
- Swiss National Science Foundation, SNSF (grant No. 163800; 165510; 173130; 176007; 176498; 182966; 182975; 183584; 185265; 188140; 188823; 190521)
- SNSF Sinergia: 183584; CRSII5_189972 / 1; SNSF BRIDGE: 176498; 184179
- Innosuisse (grant No. 27813.1 PFL-LS; 29936.1 IP-ENG; 31010.1 IP-LS; 32213.1 IP-ICT; 33668.1 IP-SBM; 33780.1 IP-LS; 37855.1 IP-LS; 39409.1 SUIN-LS; 41236.1 IP-LS)
- Innosuisse (grant No. 27813.1 PFL-LS; 29936.1 IP-ENG; 31010.1 IP-LS; 32213.1 IP-ICT; 33668.1 IP-SBM; 33780.1 IP-LS; 37855.1 IP-LS; 39409.1 SUIN-LS; 41236.1 IP-LS)

Highlights

Second-generation Lung-on-Chip

The Organs-on-Chip Technologies group developed the second-generation lung-on-chip that mimics the lung alveolar environment in an unprecedented way. An ultra-thin biological and stretchable membrane, made of collagen and elastin, two proteins found in the lung ECM, was created by surface tension using a gold mesh, and used to culture patient’s cells (picture). Zamprogno et al., bioRxiv 608919, 2019.

Immunostaining of patient cell cultures on a second-generation lung-on-chip (Picture: Pauline Zamprogno)

Micro-Meso Scale Transition of Bone Strength

Osteogenesis imperfecta (OI) is a genetic disorder causing bone fragility, whose biomechanical origin remains unclear. Micro-pillar compression is an innovative technique characterizing the mechanical properties of bone at the ELM level. The Musculoskeletal Biomechanics group demonstrated that the micro-compressive property of adult OI bone is not inferior to those of healthy controls.

Indermayer et al., ASBMR 2019, Orlando, FL, USA.

Quasi static compression test of two representative micro-pillars (Picture: Michael Indermayer)

Mechanical Heart Valves and Turbulent Blood Flow

Mechanical heart valves have been linked to the production of unphysiological turbulent blood flow. As a first step toward a model for laminar-turbulent transition in this flow, the Cardiovascular Engineering group identified an absolute instability in the impinging leading-edge vortices which develop between the valve leaflets.


A pocket of absolute instability between the leaflets of a mechanical heart valve causes turbulent flow in ascending aorta.

Ophthalmologists today largely use Optical Coherence Tomography (OCT) to monitor chronic eye conditions, leading to an enormous amount of image data created. The Artificial Intelligence in Medical Imaging laboratory has developed a machine learning method that identifies a wide range of common retinal biomarkers from OCT scans fast, reliably and consistently for the use of physicians.


Automated detection and localisation of key biomarkers in OCT scans with human-level performance through an AI system.

Expert-level Automated Biomarker Identification

Endovascular therapy in patients suffering from peripheral arterial disease shows high rates of restenosis. The Computational Bioengineering group developed personalized computational fluid simulations to understand the mechanisms responsible for this poor clinical outcome. Our models showed that flow parameters can be better predictors of restenosis than conventional clinical measures.


Distribution of wall shear stress (TAWSS) in a stented artery with arterial kinking during leg flexion, the atheroprone mechanical heart valve causes turbulent flow in ascending aorta

Prediction of Restenosis based on Hemodynamical Markers
The Bern Center for Precision Medicine (BCPM) was established as an interfaculty center in 2019, based on the initiative and support of the canton, the university, and the university hospital (Inselspital). The BCPM is composed of the management board, headed by Prof. Mark Rubin; committees, such as the scientific review committee headed by Prof. Nadia Mercader Huber; the administration (currently two people, which makes the BCPM the smallest center of the Bern University); the academic members (the number is steadily growing, currently we count 65 members); most of its members come from the Medical Faculty.

Structure

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Profile

- Reporting back to the University board and the rector as well as to the University Hospital board
- Offers a platform for the researchers of both the University and Inselspital who are active in the field of precision medicine, with the goal to promote and further the establishment of precision medicine in Bern
- Aims to develop new medications and methods, increases the quality of treatment for patients, and relieves the healthcare system with more efficient therapies; in addition, it strengthens the national and international position of the University of Bern and the Inselspital in the area of precision medicine
- Active in research, education, training and networking. In its first year of existence, it funded a total of 15 research projects selected from two separate project calls. The first call aimed at supporting the use of modern genomics in precision medicine. The second call aimed at improving the reproducibility and sustainability in PM research, and at improving the collaboration and connection among the various research groups and platforms of the Inselspital and the University of Bern. Further plans include the establishment of educational programs in the domain of precision medicine, in cooperation with the existing graduate schools. Promotion and outreach goals have also been tackled via the establishment of the Website (www.bcpm.unibe.ch), a Twitter channel (@PMBern), and the organization of various workshops and conferences, some of which gained international participation.

Grants

- Funds from the University
- Support of the canton and the Inselspital (invested in financing the Next Generation Sequencing infrastructure) These funds will support the first four years of BCPM activity (2019-2022)

Highlights

In the first year of its existence, the center was able to align the technical and knowledge platforms active in precision medicine in a network (a comprehensive list can be found on the Website), and it reached out to the important external partners such as the Swiss Personalized Health Network (SPHN) in Bern, the Data Coordination Center (DCC) in Basel, other universities such as the ETH Zurich, and other hospitals such as the cantonal hospitals of Geneva, Lausanne or Basel-Landschaft.

BCPM Inauguration

On 20 May 2019, the official inauguration of the BCPM took place in Bern, in the presence of Mrs Christine Häslé, Director of Education of the Canton of Bern, Prof. Christian Leumann, rector of the University of Bern, Dr. Uwe E. Jocham, President and CEO of the Insel Gruppe, Dr. Bernhard Pulver, President of the Board of the Insel Gruppe and Prof. Matthias Egger, President of the SNF. One of the highlights was the announcement of the first seven BCPM research projects.

2 pilot project calls (February and July)

First project call: The primary goal of this call was to support innovative translation-al research projects that address key challenge areas in Precision Medicine, with a special focus on the usage of Next Generation Sequencing (NGS). 7 projects were selected.

Second project call: Two topics were defined by the BCPM members for this call. Topic 1: Reproducibility and sustainability in Biomedical Research. Topic 2: Integrating Data from Multiple Platforms for Precision Medicine. 8 projects were selected.

The winners of both BCPM pilot project rounds

Precision Medicine seminars

The first precision medicine seminar was organized in 2018 with the participation of Prof. Dr. William Sellers from the Broad Institute of MIT and Harvard, USA. In 2019, two seminars were organized, the first one with Prof. Mark Bender Gerstein (co-director of the Yale Computational Biology and Bioinformatics program), the second one with Prof. Jean-Pierre Hubaux (ETH Lausanne, head of the laboratory in data security, invested in data security for personalized health issues).

General assembly and member workshops

In 2019, the BCPM gathered its members and stakeholders on three occasions: A first general assembly was held on 6 March, resulting in a first get-together. The inauguration took place on 20 May it showed the political buy-in. A strategy workshop was held on 3 June it formed an important basis for the subsequent project calls, and the research profile as a whole.

First General Assembly of the BCPM

From left to right: Christian Leumann, Mark Rubin, Christine Häslé, Uwe E. Jocham, Bernhard Pulver, Matthias Egger.
Highlights

Dr. Maria Nieves Sanz Garcia

Johanna Dürmüller-Bol DBMR Research Award 2019

The Research Prize was awarded to Dr. Maria Nieves Sanz Garcia, Department of Cardiovascular Surgery, Inselspital, Bern University Hospital and Research Group Cardiovascular Surgery, Department for BioMedical Research, for the project “Mitochondrial preservation and mitochondrial damage-associated molecular patterns (mtDAMPs) in DCD heart transplantation”.

Day of BioMedical Research 2019

Research Prize funded by Johanna Dürmüller-Bol Fondation, Poster Prizes of the DBMR, Best Publication 2018.

The Prize winners

Johanna Dürmüller-Bol DBMR Research Award 2019

Dr. Maria Nieves Sanz Garcia

Grants

• Department of Defense (DoD), Prostate Cancer Research Program, Impact Award, The role of CRIPTO signaling in lethal prostate cancer, PI: Marianna Kruithof-de Julio
• Department of Defense (DoD), Peer Reviewed Orthopaedic Research Program, Applied Research Award, In Vivo and Ex Vivo Models to Study Ischemia / Reperfusion Injury, Endothelial Cell Protection and Limb Preservation in a Prolonged Field Care Scenario, PI: Robert Rieben
• SNF Sinergia, Hijacking Transcription-Coupled DNA Repair for Cancer Therapy, Phd - Shana Sturla (ETH), Mark A. Rubin (Univ. Bern), Orlando Schärer (UNIST)
• Project P3RT, Pioneer Projects (3rd call), Characterization of Oncogenic Protein Isoforms as Targets for Myelodysplastic Syndromes, PI: Nicolas Bonadies
• Project SNF, Project (Div III), Tracing prostate cancer heterogeneity, PI: Marianna Kruithof-de Julio
• Project SNF, R’Equip, Ion mobility enabled ion fractionation in the gas phase for deeper prote-ome characterization, PI: Manfred Heller

Profile

• Research department of the Faculty of Medicine of the University of Bern
• Mission: to provide the researchers of the Inselspital with the best possible environment and infrastructures
• Aim: to bridge laboratory-based and biomedical patient-oriented clinical research
• Most of the research groups host masters, PhD, MD-PhD-students, and post-doctoral fellows who are integrated in the research projects of the individual groups
• Organizing progress reports, seminars and conferences that can be credited by the PhD-students
• All group leaders of the department participate in teaching activities

Structure

The role of the Department for BioMedical Research (DBMR) is to provide its 47 research groups with optimal infrastructure and scientific support. The vast majority (43) of these groups are from clinics of the Inselspital, Bern University Hospital. The remainders (4) are internal DBMR groups involved in the scientific support and coordination of equipment and infrastructure on a daily basis. Equally important, the DBMR is responsible for operating Technology and Animal Core Facilities. Furthermore, the groups of the department are supported by central services responsible for administration, informatics, technical support and bioinformatics.

Department for BioMedical Research (DBMR)

University of Bern, Murtenstrasse 35, 3008 Bern
www.dbmr.unibe.ch

Prof. Mark Rubin
Director DBMR

Prof. Robert Rieben
Directorate Coordinator Murtenstrasse 40/50 Augenklinik

Prof. Willy Holzatter
Directorate Coordinator M.E.M.-Haus Pathologie

PD Dr. Manfred Heller
Coordinator Kinderklinik

Prof. Peter Vormuthen
Coordinator SITEM

PD Dr. Monique Vogel
Coordinator Sahli-Haus 1+2
Microscopy Imaging Center (MIC)

c/o Theodor Kocher Institute, Freiestrasse 1, 3012 Bern
www.mic.unibe.ch

Structure

The MIC was founded in 2005 and has since developed into the University of Bern’s center of excellence for high-end microscopy in the life sciences. Experts in microscopy from the Medical, Science and Vetsuisse Faculties of the University of Bern and from the University of Fribourg have joined forces to support high-end microscopy and imaging analysis. In 2019 the MIC Commission, which is the directive panel, involved 32 representatives. Currently, the MIC oversees 60 instruments including 48 light microscopes, 9 electron microscopes, two atomic force microscopy systems and one micro computed tomography (micro-CT) instrument. In 2019, this equipment was used by 341 researchers.

Profile

• Mission. Expert knowledge and technical support. Implementation of new technologies. Central access to microscopy, teaching, events and services.
• Teaching and events. Lecture series on Advanced Microscopy, MIC workshops and MIC trainings. PhD program Cutting Edge Microscopy (CEM). In 2019, the MIC organized a 3-day summer school, a 2-day visit of the European Molecular Biology Laboratory (EMBL), Heidelberg, Germany, a Mini-Symposium on Multi-Dimensional Image Processing, the MIC Research Day and the MIC Symposium on Machine Learning in Imaging.
• Portfolio. From live imaging of whole organisms to organs, cells and subcellular dynamics. High throughput setups. From 2D to 5D and from the mesoscopic scale to the ultrastructural level. Light microscopy, electron microscopy, microcomputed tomography and atomic force microscopy.
• Service. Software for image analysis, image data handling and processing. Sample preparation. Introduction into microscope operation. Bimonthly newsletter for visibility and information about microscopy related activities in Bern, Switzerland and abroad.
• Internal activities. The MIC Commission convened at four regular meetings and one retreat.
• External partners. Swiss Society for Optics and Microscopy (SSOM). Life Sciences Switzerland (LS2), Intersection Microscopy. Scientific Volume Imaging b.v. (SVI). Swiss Microscopy and Imaging Core Facility Network. Science IT Support Unit of the University of Bern

Highlights

MIC Research Day on 3 July 2019

This yearly event promotes networking of researchers of the University of Bern using microscopy. The keynote speaker Prof. Nicolas Thomas explained advanced imaging for material sampling on Mars. PD Dr. Fabian Blank and Dr. Guillaume Witz prepared a well-balanced and highly informative program which attracted more than 230 participants from Switzerland and abroad. Topics covered included most recent advances in the application of machine learning to various imaging modalities such as ultrasound, electron microscopy and light microscopy.

MIC Symposium on 29 November 2019

In 2019, MIC has selected the hot topic “Machine Learning in Imaging” as the theme of the MIC symposium. The scientific committee formed by Prof. Inti Zlobek, Prof. Raphael Sznitman, Prof. Maurizio Reyes and Dr. Guillaume Witz prepared a well-balanced and highly informative program which attracted more than 230 participants from Switzerland and abroad. Topics covered included most recent advances in the application of machine learning to various imaging modalities such as ultrasound, electron microscopy and light microscopy.

Research highlight Medical Faculty


Research highlight Vetsuisse Faculty


Research highlight Science Faculty

Programmable microfluidic circuits to stimulate living cells with pulses of growth factors applied at different frequencies and concentrations. Live cell imaging using genetically encoded biosensors to measure the activation dynamics of the MAP kinase ERK at the single cell level. Measuring ERK activity dynamics in response to dynamic perturbations and growth factors is combined with mathematical modelling. Such a quantitative and predictive model of MAPK signaling provides a novel understanding on how to target this important pathway in health and disease. Research group of Prof. Olivier Pertz. Published in Embopress 2019, doi.org/10.15252/msb.20198947.
Swiss Institute for Translational and Entrepreneurial Medicine (sitem-insel)

Freiburgstrasse 3, 3010 Bern
www.sitem-insel.ch

Highlights
Promoting Services
sitem-insel Promoting Services provides a network of regulatory agencies that are located within the building and beyond. This network allows translation specialists to access information and know-how such as IP, finance, regulatory, and more in a timely and tailored manner. In addition, regular events such as the MDR Support Panel is dedicated to preparing SMEs in the medical device industry for the upcoming changes in medical device regulation.

Research Platforms are Open and Running
Starting in May 2019, several research platforms have started operating within sitem-insel AG’s new building. At a capacity utilization of almost 95%, approximately 60% of the space is dedicated to medical device technologies, approximately 30% to medicinal and biotech products, and 10% to diagnostics. The blurring of the boundaries between medicinal products, medical devices, and other services makes sense particularly against the background of the ever-closer integration of technologies and people developing and ultimately using them.

Grants, received as center/platform
• HORIZON 2020 (H2020-SC1-2017): Pushing the boundaries of Cost and Outcome analysis of Medical Technologies (COMED)
• NRP74 SMARTER HEALTHCARE: Projecting the impact of health policy changes for Swiss patients with chronic conditions using simulation modeling
• DENTAL MINISCALER, Innosuisse project
• ONE-STEP BALLOON CATHETER, Innosuisse project
• COMPUTER ASSISTED PLANNING FOR ROTATOR CUFF REPAIR, Innosuisse project
• Analysis and simulation of the distal forearm stab. during pro-supination for improved surgical Planning, SNF project
• Innovative encapsulation thin films for neural implants, Innosqueche
• European Medical Device (MDR) Regulations – Experts for Industry, ZUW University of Bern

Profile
• National center of excellence for translational medicine and entrepreneurship
• Non-profit oriented public private partnership
• Mission: to create and foster an enhanced environment for translational medicine in Switzerland
• Fosters the process of translational medicine by reducing silos building between the public and the private sector, between industries, between scientific disciplines, and within individual people
• Translational research platforms for Clinical Anatomy Training & Research and Biosafety Center
• Further tenants in the new building: parts of the Insel Group and the University, private companies and public organizations
• Network consisting of more than 100 partners such as Berner Fachhochschule BFH, Hamburg Center for Health

• Shareholders: Berner Fachhochschule BFH, Cathilf Schweitz AG, CSL Behning AG, Diabetes Center Berne Foundation, Energie Wasser Bern (ewb), Inselspolit-Stiftung, Laboratorium und Grosse Apathische Dr. G. Richelet Holding AG, Universität Bern, VCCI Stiftung zur Förderung der medizinischen und translationalen Forschung, Zur Rose Group AG (in alphabetical order)

The new building at the door of the Insel Campus

• NETWORK & COMMUNICATION
• SCHOOL
• ENABLING FACILITIES
• PROMOTING SERVICES
• CENTRALIZED SERVICES

sitem-insel school
The sitem-insel school offers university education to professionals in medical devices and life science industry. The school’s Continuing Professional Development (CPD) courses are taught by university faculty and private-sector lecturers. In September 2019, the new study program “Artificial Intelligence in Medical Imaging” started with an international symposium at sitem-insel.

Grants, received as center/platform
• HORIZON 2020 (H2020-SC1-2017): Pushing the boundaries of Cost and Outcome analysis of Medical Technologies (COMED)
• NRP74 SMARTER HEALTHCARE: Projecting the impact of health policy changes for Swiss patients with chronic conditions using simulation modeling
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From 28 August to 30 August 2019, sitem-insel AG has held opening festivities for its new building on the Insel Campus. The program featured several highlights such as:

- Press conference with representatives from the Ministry of Economic Affairs of the Canton of Bern, the architectural team, and sitem-insel AG’s management
- Ceremonial occasion with invited guests from politics, the business and industry community, as well as public authorities
- Open house with more than 2500 visitors

Official opening of the new sitem-insel AG building
University Cancer Center (UCI)

Inselspital, Bern University Hospital, Freiburgstrasse 10, 3010 Bern
www.tumorzentrum.insel.ch

Profile
- Prevention, diagnosis, treatment and follow-up of cancer
- Aim: being the reference cancer center within the Canton of Bern and being among the top cancer centers of excellence in Switzerland
- Coordinates activities and services relating to cancer within the Inselspital and offers outside services to institutions, physicians, patients and their relatives
- Strong commitment to basic and translational cancer research with active clinical trials across all cancer diagnoses

Highlights

Launch of a cancer survivor clinic

Highlights of 2019 were the launch of a cancer survivor clinic to offer advice to patients and their family physicians as a cooperation of the Departments of Internal Medicine and Pediatrics together with the University Cancer Center Inselspital. As cancers are being increasingly cured, many pediatric and young adult patients survive into their late adulthood. Cancer survivors are often faced with multiple health problems resulting from late effects of their cancer treatment, which are often unrecognized and/or undertreated. Parallel to the Cancer Survivor Clinic, we offer regular expert meetings to facilitate networking and develop guidelines for the care of such patients. Another vulnerable population with cancer are the old and very old. As many patients live well into their late eighties and early nineties, such patients often have similar expectations in respect to their cancer treatment as younger persons. Meeting these expectations poses a particular challenge. In 2019, we launched an expert group to facilitate interdisciplinary and interprofessional exchange and develop guidelines for the care of such patients.

Structure

The structure of the UCI is outlined in figure 1. Twelve organ-specific cancer centers constitute the core of the UCI – supported by interdisciplinary services and expert groups focusing on special aspects of cancer care.

Outreach

The UCI has close ties with the University of Bern, the cancer registry of the Canton of Bern as well as with national and international research organizations. Referring institutions and practicing physicians can participate in one of 16 organ-specific interdisciplinary tumor boards that offer expert advice on newly diagnosed cancer patients as well as on individual difficult-to-treat cases. In addition, outpatient clinics offer expert cancer care at each of the organ-specific centers for second-opinion.
University Neurocenter
Inselspital, 3010 Bern
www.insl.ch/das-inselspital/universitaeres-neurozentrum-bern

Structure
The Neurocenter Bern, representing the clinical and scientific cooperation of the University clinic for Neurology, Neurosurgery and Neuroradiology as well as Neuropsychiatrics and Psychiatry has been founded in 2012. With 400 beds, >1400 collaborators, >60’000 ambulatory patients, >8000 stationary patients, and >3500 operations/interventions/sciographies each year and a budget of > 350 Mio CHF it is the largest of its kind in Switzerland, and leading in Europe.

Profile
Research areas with strong transdisciplinary approaches/interactions within the Neurocenter include the following:
- stroke* (4 out of 5 departments), including complex neurovascular diseases
- sleep-wake-epilepsy*, including epilepsy surgery (4/5)
- Advanced neuroimaging, intraoperative imaging, post-processing and clinical implementation of machine learning (5/5)
- movement disorders, including functional neurosurgery for Parkinson’s disease, neurological and psychiatric disorders (5/5)
- neurorehabilitation including computer-assisted systems and robotics (4/5)
- neuroimmunology* (3/5)
- neurodegeneration*/dementia (3/5)
- neurorehabilitation, augmented reality, navigation associated developments, targeted procedures and 3D printing and simulation techniques (2/5)
- systems neuroscience and psychopathology (3/5)
*in these areas clinical/human and experimental/animal approaches are used

Grants
- Swiss National Science Foundation
- Neurology: 3 grants awarded in 2019: 185362, 188761, 189077. 18 running grants: 166827; 169975; 175615; 172952; 179629; 179436; 179895; 169789; 169379; 173081; 170060; 179667; 172676; 179277; 177520; 179565; 175984
- Neurosurgery: SINERGIA Project “Predict and Monitor Epilepsy After a First Seizure: The Swiss-First Study”
- SNF SPARK “Using Deep Neural Networks to Bridge Clinical and Quantitative Analysis of Intracranial EEG in Epilepsy”
- SNF Grant “Imagine” (USZ, USH, Inselspital, USB, EOC, CHUV, HES-SO.)
- MR-research project “The Brainfit-Study, Krebsstiftung (4708-02-2019)
- INNOSOUSSE (3 grants awarded in 2019: project no. 30664.1 IP-LS; 35565.1 IP-LS; 32213.1 IP-JCT)
- "Suprafascial vancomycin powder for prevention of surgical site infections after instrumented posterior spinal fusion: A randomized controlled phase-II trial" TU-Research Grant and research grant form the Gottfried & Julia Bangerter-Rhyner Foundation
- “MAGNEURYSM: Real-time Augmented Visualisation of Vessel Perfusion with Automated Video Analysis” Inselspital CTU-Forschungsgrant
- "A proteomics discovery approach to identify candidate biomarkers of athebroclerotic plaque instability in endarterectomy specimens of patients with carotid disease." Schweizer Herzstiftung

Highlights
The Neuro Clinical Trial Unit
The Neuro Clinical Trial Unit (NCTU) is a platform of the Neurocenter Bern to support planning and realisation of clinical research in neurosciences. The NCTU works in close collaboration with the CTU Bern and the “Department für Lehre und Forschung” and has substantiated expertise in the coordination of multicenter national and international clinical trials. Currently, the Neurocenter Bern is conducting over 80 clinical studies; including three large-scale multicenter investigator initiated clinical trials managed by the NCTU Bern:
- ELAN: Early versus Late infusion of direct oral Anticoagulants in post-ischaemic stroke patients with atrial fibrillation (ELAN)
  www.elan-trial.ch
- SWITCH: Swiss trial of decompressive craniectomy versus best medical treatment of spontaneous supra-tentorial intracerebral hemorrhage (SWITCH)
  www.switch-trial.ch
- SWIFT DIRECT: Solitaire™ With the Intention For Thrombectomy Plus Intravenous t-PA Versus DIRECT Solitaire™ Stent-retriever Thrombectomy in Acute Anterior Circumflex Stroke (SWIFT DIRECT)
  https://www.swift-direct.ch/

Opening of the Magnetic Resonance Research Center at the sitem-insel AG – The Translational Imaging Center
The Translational Imaging Center is one of the core facilities of sitem-insel AG. This imaging platform enables translational research ranging from the chemical analysis of samples to model systems and organs to whole body clinical and ultra-high field (UHF) magnetic resonance imaging (MRI), which are integrally linked to the clinical environment and the clinical research of the University of Bern and the Inselspital. Magnetic resonance technology, in particular UHF, opens up a non-invasive examination option for investigating the structure and function of biological tissue. With the paradigm shift from cohort to personalised medicine, it marks a cornerstone for diagnostics and therapy monitoring and the research of new biomarkers.
The established research parameter including a 3 Tesla and a novel UHF 7 Tesla MRI in direct proximity to the Inselspital will foster clinical science in Bern. The MRI-research has been an common interest of the Neurocenter in the past and will allow us to integrate and extend our research even further in the future.

NeuroTec
NeuroTec is an innovative platform - located in the new stem building - that enables efficient interactions between health and technology specialists from science and industry. The goal of NeuroTec is to promote new, flexible and cost-efficient technologies improving outpatient diagnostics and therapy in neurology. A core infrastructure of NeuroTec is the “Neuro-Loft” - an instrumented apartment, in which new methods and devices can be tested under real-world conditions.

https://www.swift-direct.ch/
Clinical Trial Unit Bern (CTU)

Mittelstrasse 43, 3012 Bern & sitem-insel, Freiburgstrasse 3, 3010 Bern

www.ctu.unibe.ch

PD Dr. Sven Tschopp
Director

Prof. Urs Fischer
Deputy Director

Christina Huf
Head Quality Management

Murial Helmers
Head Data Management

Dr. Andreas Limacher
Head Statistics and Methodology

Dr. Felix Renten
Head Monitoring and Regulatory Affairs

Structure

The CTU Bern is an institute of the Faculty of Medicine, works mainly in five areas and is structured accordingly:

- project coordination and management
- statistical analysis and study design
- data management
- monitoring and regulatory affairs
- clinical investigation

The CTU staff is supported by the institutional quality management of CTU to improve processes.

Consulting and project support are one of the main objectives of CTU Bern.

In 2019, CTU Bern conducted more than 330 consultings, mainly for researchers from Inselspital Bern, and supported more than 100 projects.

Profile

- To provide the scientific, technical, and computing expertise needed to conduct patient-oriented clinical research at all stages, from conception to completion and dissemination
- Support is provided in a modular fashion and ranges from advice and general support to full development of the design and conduct of clinical studies
- Staff contributes to undergraduate teaching for 3rd and 6th year medical and biomedical sciences students
- Postgraduate teaching activities involve contribution to the Certificate of Advanced Studies course in Clinical Research (University of Bern) and the Certificate of Advanced Studies for Clinical Research Coordinators offered by the University of Applied Sciences Bern
- Research ethics and Good Clinical Practice (GCP) training form an important part of our teaching activities and are supplemented by special courses such as software trainings or a course on writing a study protocol and the monthly CTU lecture
- Research partners encompass all clinical departments and institutes of the Faculty of Medicine, members of the Swiss Clinical Trial Organization CTU network, and several Swiss cantonal and private hospitals

Grants, received as center/platform

- CTU Bern supports the conduct of numerous projects financed by grants as partner

Highlights

Clinical Investigation moved to sitem-insel building

On 27 August it finally happened – The Clinical Investigation (CI) division of CTU Bern moved into its new facilities of stem-insel. The new environment offers fully equipped state-of-the-art examination and treatment rooms. Smaller offices have been set up as a “touchbase” to allow presence of CTU staff located at Mittelstrasse 43. They offer the possibility to promote exchange between the CTU Bern and Inselspital investigators. Altogether, this move offers the possibility to occupy an important place on the stem-insel agora of scientific exchange.

ISO 9001:2015 certification and audits

In the first quarter of 2019, CTU Bern had its quality management system (QMS) externally evaluated and assessed. Since April 2019, CTU Bern has successfully been ISO 9001: 2015 certified. Although an ISO certification is not a regulatory requirement, external review and feedback offer the opportunity to improve processes and to achieve higher quality. In summer 2019, CTU Bern was successfully audited as part of a clinical trial in which CTU Bern takes on major responsibilities. This external audit further assured CTU Bern, that internal processes are well established, generally aligned with study sponsor requirements and up to date in regard to regulatory expectations.

SCTO/SERI platform statistics and methodology

In 2017, the Swiss network of clinical trial units received a four year grant from the State Secretariat for Education, Research, and Innovation to foster collaboration and improve the clinical research environment for multicentre clinical trials. Within this grant, CTU Bern is responsible for the statistics and methodology platform.

This year, CTU Bern hosted two events where statisticians of most CTUs as well as SAKK participated. In spring, a workshop on competing risk and multistate models was held with an internationally renowned faculty. In autumn, a workshop was held in Biel where quality aspects within the statistical workflow of a clinical study were discussed with a focus on improving the automatic and reproducible reporting of statistical information in publication-ready summary tables.

Department of Clinical Research: An outlook to 2020

So far, CTU Bern has been the only unit at the Faculty of Medicine dedicated to support academic clinical research activities. Complexity of clinical research has been growing steadily and faculty has acknowledged the importance of such research projects. Therefore, CTU Bern will move under a larger umbrella in 2020: the Department of Clinical Research. This new department will strengthen the clinical research activities at the Faculty of Medicine especially at Inselspital. It will be led by a full professor of clinical research. The department will be responsible for clinical research but also for education and especially promotion of young clinical researchers. CTU Bern will remain with its offices at Mittelstrasse 43. The other parts of the department will be located at the sitem-insel building on the Inselspital campus. A major focus there will be clinical investigation which will run an outpatient clinic for clinical research.
Institute at the University of Bern

Institute of Anatomy
Institute of Biochemistry and Molecular Medicine (IBMM)
Institute for the History of Medicine (IMG)
Institute of Pharmacology (PKI)
Institute of Physiology
Institute of Social and Preventive Medicine (ISPM)
Theodor Kocher Institute (TKI)
Institute for Infectious Diseases (IFIK)
Institute of Pathology
Institute of Forensic Medicine (IRM)
Institute of Dental Medicine (ZMK)
Institute of Primary Health Care (BIHAM)
Institute of Complementary and Integrative Medicine (IKIM)
Institute for Medical Education (IML)
Cell membrane structure and microvesicle shedding capacity determines differential survival of immune cells

We show here that myeloid cells are more resistant to cell death by pneumolysin (PLY), a pore-forming toxin from S. pneumoniae, as compared to lymphoid cells. Our results suggest that, immune cells important early in infections, i.e. myeloid cells, strongly benefit from an efficient cellular repair mechanism to ensure survival after PLY attack (Yu L et al., FASEB J 2019, in press).

Cellular Mechanisms of Cardiac Regeneration

(A) This year, the group of N. Mercader explored the role of macrophages during regeneration. They found a subset of macrophages with a pro-regenerative gene signature that has an important role during heart regeneration (Sanz-Morejón, García-Redondo et al., 2019 Cell Reports).

(B) Furthermore, the group identified a small subset of cardiomyocytes that expands in response to injury at a higher degree than the rest of cardiomyocytes and plays a fundamental role during myocardium regrowth (Sande et al., 2019 Cell Reports).

Pneumolysin and pneumococcal capsule influence taxis and motility of microglia

We show that microglia possess the capacity for a very agile response towards bacterial pathogens, but key pathogenic factors, such as pneumococcal capsule and pneumolysin, are able to abolish this response by interference with microglial taxis and motility. Hupp S. et al., J of Neuroinflammation (2019) 16:105.

Institute of Anatomy

Balztenstrasse 2, 3012 Bern
www.ana.unibe.ch

MicroangiCT of a minipig eye

First International Workshop on MicroangiCT

Institute of Anatomy (Prof. V. Djonov) and MicroCT Group (PD Dr. R. Hlushchuk) with the support of MIC (Prof. B. Zuber) have successfully conducted the first international workshop on “MicroangiCT: Vascular imaging using a contrast-enhanced microCT approach”. It will become a yearly event held in September. In 2021 it will be supported by COMULIS COST Action (CA17121).

Highlights

- Teaching: The Institute of Anatomy is responsible for the comprehensive teaching of medical students (Human and Veterinary Medicine, Dentistry) and students of related paramedical fields (Biomedical Sciences, Pharmacy, Biomedical Engineering, Physiotherapy, Osteopathy). In addition, postgraduate education and advanced surgical training is provided. We also educate postgraduate medical students (Master/MD) as well as PhD students at the GCB, including the interfaculty Cutting Edge Microscopy PhD Specialisation Course (responsible: Prof. N. Mercader Huber)
- 9 research groups with various research topics: microbeam radiation therapy; cellular damage control and liposomal nanotracers against bacterial toxins; cardiac development and regeneration; ultrastructure and function of synapses, protein structure; pulmonary development and function of cilia; neuron inflammation; microCT-based imaging
- External research partners: Australia: Australian Synchrotron, University of Melbourne, University of Sydney, Monash University, Finland: Biomedicum Center Helsinki, France: IGMCB, CNRS, ESIR, University Hospital-Henri Mondor, Institut du Fer à Moulin. Germany: MR-HUK, University of Giessen, DZNE, MPIB, University of Münster. Israel: Hebrew University. Latvia: University of Latvia. Spain: CNIC, Sweden: Lund University, Switzerland: Idiap Research Institute, Anapath Services AG, University of Basel, Institut Straumann AG, University of Zurich, PSI, Lauscco SA, ETH Zürich, CHUV UK: University of Birmingham
Institute of Biochemistry and Molecular Medicine (IBMM)

Bühlstrasse 28, 2012 Bern
www.ibmm.unibe.ch

Deputy Director
Prof. Christin Albrecht

Pro-drug: Gertsch

Prof. Dr. Martin Lochner
Co-Director

Grants
• Swiss National Science Foundation (NCCR TransCure (Leading House), NCCR Molecular Systems Engineering (NCCR TransCure: Excellence in Membrane Transport Research)
• D-BSSE & D-CHAB, ETH Zürich, Switzerland, Swiss Institute of Bioinformatics, University of Geneva, School of Medicine, University of Nottingham, UK, Department of Pharmacology, University of Cambridge, UK, Novartis Institutes of Biomedical Research, Basel, Switzerland, Max-Planck-Institute, Munich, Germany, Laboratory of Molecular Electron Microscopy, The Rockefeller University, New York, USA, PeterMac Institute, Melbourne, Australia, NIH, NAAA, Rockville, USA, Institute for Research in Biomedicine (IRB Barcelona), University of Barcelona, Spain

Highlights
• Basic and specialized teaching in biochemistry, molecular and pharmaceutical biology, and molecular medicine, teaching activities in the faculties of Medicine, Natural Science and Vetsuisse to pre- and postgraduate students (GCB)
• Nine research groups
• Interdisciplinary research on structure, function, and pharmacology of membrane proteins (transporters, ion channels, and receptors), with a strong emphasis on the roles of these membrane proteins in human diseases such as cancer, neuropsychiatric and cardiac disorders, pre-eclampsia and pathogen infection
• Aim: Unravel molecular working mechanisms of selected membrane proteins, discover new therapeutic targets
• Leading House of NCCR TransCure - Excellence in Membrane Transport Research
• SNSF project grants No. 31003A_179215, 31003A_189220, 31003A_170923, 31003A_170385, 31003A_163359, 31003A_162581, 31003A_176175, 31003A_169355, 31003A_189220
• SNSF-Sinergia grants No. CRSII5_183481, CSRII5_180326/1, CRSII5_170923, 310030_184980, 31003A_169355, 310030_189220
• SNSF project grants No. 31003A_179215, 31003A_162581, 31003A_176175, 31003A_163359, 31003A_169355, 31003A_189220
• Swiss National Science Foundation (NCCR TransCure (Leading House), NCCR Molecular Systems Engineering (NCCR TransCure: Excellence in Membrane Transport Research)

Profile

First structure of a monocarboxylate transporter from the Solute Carrier 16 family

Monocarboxylate transporters play important roles in certain cancers. We have reported structures of an L-lactate-transporting solute carrier family 16 homolog with bound substrate and inhibitor. The structures show the transporter in the pharmacologically relevant outward-open conformation. Structure–function analysis provides insights into the molecular working mechanisms of ligand binding and L-lactate transport.

Boshart P0 et al., Nature Commun 2019 : 10, 2649.

Structure of the L-lactate transporter SMCT: sideview, transmembrane helices and two L-lactate molecules are in rainbow colours, and black and red.

Different modes of Nav1.5 reorganization at the cardiomyocyte lateral membrane and T-tubules

Mutations in the gene encoding the voltage-gated sodium channel Nav1.5 cause many different cardiac arrhythmias, but the causes of this variety remain unknown. We show in mice that different mutations (Nav1.5 truncation and dystrophin deficiency (mdx)) cause different modes of Nav1.5 reorganization in specific cardiomyocyte domains (T-tubules and lateral membrane). Nav1.5 mutations therefore seem to site-specifically affect Nav1.5 localization, distribution, and possibly function.

Vermij et al., bioRxiv 674275.

Cardiomyocyte model showing the effects of dystrophin deficiency (mdx) and Nav1.5 truncation (DSIV) on Nav1.5 distribution (green).

Interaction of antiemetics with serotonin-gated ion channels

We have synthesised several analogues of the clinically used antiemetic drug granisetron. These granisetron probes feature photo-crosslinking groups and our modifications did not affect their high affinity for the target, the serotonin-gated type-3 ion channel (5-HT3R). Furthermore, these granisetron derivates showed specific photo-crosslinking with the human receptor. Detailed analysis after photo-crosslinking by protein-MS/MS identified the exact residue in the receptor binding site that was covalently modified.


Placental versus hepatic secretion of apolipoprotein A1 and E

High levels of atherogenic lipids in pregnancy are associated with health complications for the mother, the fetus and the newborn. By ex vivo human placenta perfusion and primary cell culture experiments we demonstrated that the human placenta releases high amounts of anti-atherogenic apolipoproteins, particularly apoA1 and apoE. These data indicate that the placenta plays an important role in maternal and fetal cholesterol homeostasis during human pregnancy.


Placental secretion of apolipoprotein A1 and E

TRPM4 in colorectal cancer (CRC)

In cooperation with the Translational Research Unit of the Institute of Pathology (Unibe) we showed that TRPM4 is highly expressed in human CRC tumor buds and associated with an infiltrative growth pattern. TRPM4 contributes to cell viability, proliferation, migration, invasion and cell cycle. TRPM4 plays a versatile role in cancer cell pathophysiology.

Kappel et al., Mol Oncology 2019 : 13, 2393 - 2405.

Photo-crosslinking of 5-HT3 receptor

Placental versus hepatic secretion of apolipoprotein A1 and E
In 2019 we finalized the process of digitizing and storing the whole archive of the Swiss Academy of Medical Sciences at our Institute (2,000 folders). Based on these sources, Dr. Magaly Tornay developed the SNSF-funded research project “Governing by Values: On the History of Medical Ethics and Bioethics in Switzerland”, which started in October 2019.

Medical ethics in Switzerland

In 2019 we finalized the process of digitizing and storing the whole archive of the Swiss Academy of Medical Sciences at our Institute (2,000 folders). Based on these sources, Dr. Magaly Tornay developed the SNSF-funded research project “Governing by Values: On the History of Medical Ethics and Bioethics in Switzerland”, which started in October 2019.

A new European Journal

The Institute hosted a conference with colleagues from 19 European countries in order to discuss the state of medico-historical research and the establishment of a new “European Journal for the History of Medicine and Health” in 2021. There is a need for a high quality journal serving as a platform for various European perspectives on medical history. Hubert Steinke will be Co-Editor-in-Chief of the journal.

Psychiatric hospitalisation

After five years of research, the Independent Expert Commission (IEC) on Administrative Detention published its Final Report, written by IMG staff member Urs Germann and Lorraine Odier. Among others, it shows the growing significance of psychiatric hospitalisation and calls for further research beyond the “watershed” of 1981.

Institute for the History of Medicine (IMG)

Bühlstrasse 26, 3012 Bern
www.img.unibe.ch

- Teaching students of medicine at the Universities of Bern, Basel and Fribourg as well as history students and master students at the biomedical engineering program
- 1 professor, 1 assistant and further scientists in SNSF-funded projects
- Rich library (100,000 vols.), archive documenting local and Swiss medicine (paper and digital), large collection of medical objects from University and University Hospital
- Research on medical theory and practice 18th to 20th century, Albrecht von Haller and the 18th century Republic of Letters, history of medical ethics, history of the quality of life concept
- External partners: Institute of Biomedical Ethics and History of Medicine, University of Zurich; Institute for the History of Medicine and Health, University of Lausanne; School of Liberal Arts, Indiana University; Center for African Studies, University of Florida; Department of History, University of Montreal; Institute for History and Ethics of Medicine, University of Cologne

Grants

- Swiss National Science Foundation (grants No. 100011_159614; 100011_184880; 10615-157963; 174862; 186439)
- Canton of Aargau, Departement of Health
- Swiss Academy of Medical Sciences

Profile

Haller online

In this SNSF-funded longterm project we are editing thousands of bookreviews and letters from Albrecht von Haller’s correspondence. These sources will allow us to get a better understanding of 18th century scientific networks. The website has been launched in May 2019 and presents an ever increasing number of transcriptions.
www.hallernet.org

Facsimile and text of a letter
### Institute of Pharmacology (PKI)

Imeslpital, INO-F, 3010 Bern
[www.pk.unibe.ch](http://www.pk.unibe.ch)

#### Profile

- Teaching students of medicine, dental medicine, pharmacy, biomedicine and biology as well as graduate students at the Graduate School for Cellular and Biomedical Sciences (GCB)
- 7 research groups
- Investigation of pathophysiological processes that contribute to inflammation and tumor diseases
- Aim: Discovering new drug targets
- External Partners: Institutes of Biochemistry, Experimental Immunology and Molecular Cancer Research, University of Zürich, Zürich, Switzerland; Department of Pharmacology, Toxicology and Clinical Pharmacy, Institute of Pharmacy, University of Frankfurt/Main, Germany; Department of Medicine, University of Tübingen, Tübingen, Germany; Department of Surgery, University of Toronto, Toronto, Canada; Department of Clinical Immunology and Allergology, Sechenov University Moscow, Russia; Swiss Edi: Research Group, Olten, Switzerland

#### Grants

- Swiss National Science Foundation (grant No. 310030-166473; 310030_184816; 310030-175561; 31003A_173215; 31003A_149387; 310030-146215; 310030E-132762; 310030A-138201; 31003A_170134; 310030_184757)
- Novartis Foundation for Biological-Medical Research
- HORIZON 2020 (Marie Sklodowska-Curie Action): MEL-PLEX
- Swiss Cancer League (KFS-3703-08-2015; KFS-3941-08-2016)

#### Highlights

**Characterizing ATG12 as a novel drug target in cancer**

ATG12 is known as an essential autophagy-regulating protein. We observed that cancer cells with reduced ATG12 expression undergo an oncotic cell death, owing to reduced mitochondrial biogenesis and function. Inducing oncosis by imposing an ATG12 deficiency in solid tumors might represent a novel anti-cancer therapy.


**Glycan-specific antibodies in immunodeficiency**

The surface of microbial pathogens and tumor cells is coated with glycans, which can be recognized by glycan-specific antibodies. An atlas of the glycan-specific IgG repertoire in primary antibody deficiency was established. We observed that individuals with immunodeficiency lack a significant proportion of antibodies not only to bacterial, but also to specific tumor antigens. Furthermore, our data suggest that broad antibody repertoire analysis may have advantages over current diagnostic approaches.


**BCL-2 family protein BOK is a positive regulator of uridine metabolism**

We report that the Bcl-2 family protein Bok activates a key enzyme involved in uridine metabolism (UMPS), and thereby nucleotide biosynthesis and activity of the chemotherapeutic drug 5-fluorouracil (5-FU). We propose Bok as potential biomarker for 5-FU resistance and to develop Bok mimetic compounds to re-sensitize 5-FU-resistant cancers.

Srivastava R et al., PNAS 2019 : 116, 15469 - 15474.

**Tumor glycosylation as an immune checkpoint**

We demonstrated that one of the most common tumor glycosylation patterns (hypersialylation) provides tumor cells the capacity to escape the immune attack by cytotoxic CD8+ T cells. The attenuation of T cell anti-tumor responses involves the interaction of tumor glycans with the inhibitory Siglec-9 receptor. The recognition of sugar-structures as immune checkpoints may support immunotherapeutic approaches against cancer.

Right panel: Journal cover page with artistic representation of our results (May 2019).


**The AMP kinase is a key mediator of the endothelial barrier-protective effect of S1P**

Breakdown of the endothelial barrier is a critical step in the development of organ failure in severe inflammatory conditions such as sepsis. By measuring transendothelial resistance, we show a key role of the S1P-AMPK signaling axis in endothelial barrier stabilization in human microvascular endothelial cells (HMEC-1). This is not seen in glomerular endothelial cells (GENCs), thus proving a tissue-specific regulation of endothelial barrier integrity by S1P.

Institute of Physiology

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Prof. Thomas Nevan
Managing Director

Prof. Walter Senn
Codirector

Prof. Stephan Rohr
Codirector

Prof. Ernst Neppi
Codirector

Prof. Marcel Eggert

Prof. Stéphane Ciocchi
(SNF)

Prof. Jan Kucera

Prof. Jürg Streit

PD Dr. Sonja Kleinlogel

Prof. Jean-Pascal Pfitzar
(SNF)

Dr. Shankar Sachidanandam

Profile

- Performs basic and translational research in cardiac- and neurophysiology.
- Five groups in cardiac physiology investigate the conditions that lead to heart failure and arrhythmias. They use electrophysiological and imaging techniques as well as computer simulations to elucidate the mechanisms of action potential propagation and excitation-contraction coupling on the subcellular, cellular and organ levels.
- Six groups conduct research in neurophysiology. The common theme is to understand neuronal network dynamics in the brain in health and disease. Strategies to cure retina degeneration are devised as well as developing new treatment strategies for chronic pain or spinal cord regeneration. The formation of emotional memories, anxiety, depression and perceptual vision are studied from molecules, over single cells to neuronal networks with electrophysiological, two-photon imaging, optogenetic and behavioral approaches.
- Two computational neuroscience groups develop probabilistic models of brain function at different levels, from single synapses to cognition.
- Predominantly engaged in the education of students of human and dental medicine, responsible for the entire field of physiology, covering all lectures and practical courses. Lecturers of our institute also participate in the training of students in veterinary medicine, pharmacy and the “Master in Biomedical Engineering”. Our institute coordinates the master program in Biomedical Sciences (BMSc), contributes to the BeNeFri program in Neuroscience, as well as other interfaculty lecture courses.
- External research partners: Humboldt University, Neurocure Cluster of Excellence, Berlin, Germany; Department of Pharmacy and Drug Science, University of Bari Aldo Moro, Bari, Italy; Department of Medicine, Wisconsin Institutes for Medical Research, University of Wisconsin, USA; Kirchhoff Institute of Physics, University of Heidelberg, Heidelberg, Germany; Medizinische Hochschule Hannover, Germany.

Grants

- ERC Consolidator Grant (Nevian)
- ERC Starting Grant (Ciocchi)
- SNF Professorship Grant (Pfitzar)
- Bertarelli Foundation Catalyst (Kleinlogel)
- Human Brain Project (Senn)
- IFK Decoding Sleep (Nevian, Senn)

Highlights

Ultrafast stretch events and cardiac electrophysiology

A hefty blow to the chest can have entirely opposite outcomes. While, for instance, baseball players have died after being hit in the chest by a baseball, patients undergoing fatal cardiac tachyarrhythmias have been saved by an appropriately timed thump to the chest. In the past, investigations of electrophysiological mechanisms underlying these phenomena have been hampered by the lack of experimental systems that allow very fast stretching of cardiac tissues while simultaneously recording its electrophysiological properties. In collaboration with the EPFL Neuchatel, we developed such a device that is based on a stretchable, extremely thin silicone membrane containing integrated gold and carbon electrodes. Applying high voltage pulses to the carbon electrodes results in quasi-linear stretching of the membrane by up to 12% and at rates up to 100x those encountered by cardiomyocytes during normal heart function. Impulse propagation during strain application is recorded by extracellular gold electrodes positioned along the linear cardiomyocyte preparations. Much to our surprise, impulse conduction remained virtually unchanged even at the highest strain amplitudes and rates which suggests that the strain sensitive element of the heart may actually not be the contracting cardiomyocyte itself but adjacent electrotonically coupled connective tissue cells. The developed device is ideally suited to investigate this hypothesis and lends itself to systematic studies of the effect of specific drugs or gene therapies having beneficial effects on myocardio-electrical coupling in the heart.


Stochastic computation by spikes

When thinking of a visual scene, we imagine the different objects of the scene while the focus of attention jumps stochastically from object to another. We modeled this imagination process with a generative network of spiking neurons that serially “dream” visual objects seen before. A reduced version of the network is implemented in the Heidelberg neuromorphic hardware.


Wafer from the BrainScaleS neuromorphic hardware (left) with example of single neuron voltage time course (blue) and multi-neuron spike raster plot (top right).

Mechano-active multielectrode array (MaMEA). Prepare (a) with strength-duration curve (b) and example of an electrogram (c).
Institute of Social and Preventive Medicine (ISPM)

Mittelstrasse 43, 3012 Bern
www.ispm.unibe.ch

Muka
Dr. Taulant Franco
Prof. Oscar Pozzilli (IS), Italy.
Fellow, Honorary Consultant in Public Health Medicine; Department of Epidemiology and Prevention, IRCCS Neuromed, Epigenomics Research Group, King’s College London; Institute of Applied Health Research, UKRI Innovation Clinical Community Medicine, University of Greifswald, Greifswald, Germany; Helmholtz Zentrum München, Munich, Germany; College London; Department of Epidemiology, Erasmus Medical Center, Rotterdam, the Netherlands; Institute of Zurich, Zurich, Switzerland; University of Lausanne; HELIUS, University of Amsterdam; School of Public Health, Imperial Research Center of the Ministry of Health of the Russian Federation; Department of Nuclear Medicine, University Hospital la Salud, Colombia; Universidad de la Magdalena, Colombia; Federal State Budgetary Organization National Cardiology Schulich School of Medicine&Dentistry, University of Western Ontario, Canada; Fundacion Universitaria de Ciencias de la Salud, Colombia; Universidad de la Magdalena, Colombia; Federal State Budgetary Organization National Cardiology Research Center of the Ministry of Health of the Russian Federation; Department of Nuclear Medicine, University Hospital Zurich, Zurich, Switzerland; University of Lusanne; HELLIS, University of Amsterdam; School of Public Health, Imperial College London; Department of Epidemiology, Erasmus Medical Center, Rotterdam, the Netherlands; Institute of Community Medicine, University of Greifswald, Greifswald, Germany; Helmholtz Zentrum München, Munich, Germany; Epigenomics Research Group, King’s College London; Institute of Applied Health Research, UKRI Innovation Clinical Fellow, Honorary Consultant in Public Health Medicine; Department of Epidemiology and Prevention, IRCCS Neuromed, Pozzilli (IS), Italy.

Grants

• SNF Ambizione (P2lodp3_18592371)
• Swiss National Science Foundation (31B30_185396)
• Horizon 2020 (EU_825162)

Highlights

New research group on Climate Change and Health
The group, led by Ana M. Vicedo-Cabrera, will investigate the impact of climate change on health, with particular focus on the evaluation of adaptation strategies, mitigation policies and their health co-benefits.

SwissRDL Medical Registries and Data Linkage
SwissRDL maintains and analyses the national hip and knee implant registry on behalf of the SRS foundation. The annual report 2019 shows 2-years revision rates for primary hip and knee implant operations per hospital (panel shows hips only). With this report Switzerland presents a high-quality report comparable with others from long-standing registry world wide. Additionally, SwissRDL has negotiated and implemented new registries in 2019 and will be running the Swiss Heart Surgery Registry, the percutaneous coronary intervention registry and the idiopathic lung disease registry starting in 2020.

Cardiometabolic Health Group
This newly established research group focuses on the aetiology of cardiometabolic disease including type 2 diabetes and cardiovascular disease and their associated factors such as dyslipidaemia, hypertension and obesity. The aim is to understand the role of lifestyle factors (e.g., diet, physical activity and sleep) on maintaining optimal cardiometabolic health and preventing cardiometabolic disease, as well as in identifying new biomarkers and pathways for prediction, diagnosis, prognosis and treatment of cardiometabolic disease.

Global disparities in cervical cancer rates among women with HIV
A study by an international group of researchers led by a team at the Institute of Social and Preventive Medicine, University of Bern, found that the incidence rates of cervical cancer are particularly high among women living with HIV in South Africa or Latin America. For this study recently published in the International Journal of Cancer, researchers used data from a collaboration between global HIV cohort research networks to compare cervical cancer rates in 45 countries across Europe, South Africa, Latin, and North America among women living with HIV who initiated antiretroviral therapy between 1996 and 2014. Read the abstract on PubMed.

Cardiometabolic Health Group

24-Step guide on how to design, conduct and successfully publish a systematic review and meta-analysis in medical research
Together with researchers from the Netherlands and the UK we have developed a clear step-by-step guide on how to perform a systematic review and meta-analysis of observational studies and clinical trials. This 24-step guide simplifies the process of conducting a systematic review, provides healthcare professionals and researchers with the tools to conduct methodologically sound systematic reviews and meta-analyses, and enhances the quality of synthesis efforts already underway. Read the abstract on PubMed.
Theodor Kocher Institute (TKI)
Freiestrasse 1, 3012 Bern
www.tki.unibe.ch

Profile

- Teaching bachelor and master students of the Medical, Science and Vetsuisse faculties in lectures and practical classes in immunology, vascular and cell biology. Educating graduate students of the Graduate School for Cellular and biomedical Sciences (GCB). Coordinating national PhD programs: "Cell Migration" and "Cutting Edge Microscopy".
- The TKI hosts 5 research groups studying central nervous system immunity in health and disease with a focus on multiple sclerosis, stroke, Alzheimer's disease and brain metastasis of tumors by employing advanced in vitro and in vivo imaging approaches.
- External partners: Prof. Thorsten Buch, University Zurich, Switzerland; Prof. Christer Betsholtz, Uppsala, Sweden; Dr. Yann Decker, University of the Saarland, Germany; Prof. Michael Detmar, University Heidelberg, Germany; Prof. Fabien Gossot, University of Lille, France; Prof. Mikio Furuse, Kyoto University Faculty.

Grants

- Swiss National Science Foundation (grants 310030_189080, 31003A_170131, 310030_189226), EU Horizon 2020 MSCA-ITN 2015-675619 BtRAIN and MSCS-ITN-2018-813294 ENTRAIN; Fidelity Bermuda Foundation; ARSEP, Swiss MS approaches.

Highlights

- Oxidative pathology in mitochondria of oligodendrocytes
We have established a novel transgenic mouse model which allows imaging of oxidative pathology in oligodendrocyte mitochondria by 2-photon microscopy. By expressing a reporter protein sensitive to variations in oxidative agents (Op1-roGFP2), specifically in mitochondria of myelin-producing oligodendrocytes, we can visualize - at a single organelle level - dynamic redox changes in the spinal cord of anesthetized animals. After establishing the surgical approach and data analysis, we now move toward dissecting the mechanisms of mitochondrial pathology during the development of anti-CNS inflammation.

- Successful finalization of the international PhD program BtRAIN
The EU Horizon 2020 funded ITN BtRAIN coordinated by Britta Engelhardt was successfully finished. BtRAIN has created novel knowledge on the vertebrate brain barrier signature genes and their specific role in regulating brain barriers function in development, health, ageing and disease. BtRAIN has educated 12 young brain barrier researchers in an interdisciplinary manner allowing them to obtain unique skills and thus making them highly desired researchers in the academic and non-academic sector in the brain barriers field.

- Clearance of cerebrospinal fluid from the sacral spine through lymphatic vessels
We have recently shown with dynamic in vivo imaging that routes of outflow of CSF in mice occur along cranial nerves to extracranial lymphatic vessels rather than directly to venous blood as commonly assumed. In this project, we used near-infrared and magnetic resonance imaging to demonstrate a cranial-to-caudal flow of CSF traces within the spinal column and revealed that the major spinal pathways for outflow occurred to lymphatic vessels at the sacral region. Outflow of CSF from the spine to lymphatic vessels may have significance for many conditions, including multiple sclerosis and spinal cord injury.

- Molecular composition and function of blood-brain barrier tight junctions
Complex tight junctions (TJs) between brain microvascular endothelial cells forming the blood-brain barrier (BBB) block uncontrolled paracellular diffusion of molecules across the BBB. Our studies advanced understanding of the molecular composition of BBB tight junctions and identified a novel role for BBB cell-to-cell contacts in regulating T-cell trafficking across these cellular junctions of the BBB ensuring central nervous system immune surveillance. Castro Dias et al., 2019, Sci Rep. 9(1):2033; Castro Dias et al., 2019, FBCNS, 16(1):30; Castro Dias et al., Int J Mol Sci. 20(21), pii: E5372; Wimmer et al., 2019 Front Immunol, 5:10:711.

- Brain metastasis formation: Melanoma cells breach the blood-brain barrier
Prior to colonization of the brain, metastatic melanoma cells breach the BBB. We have developed a high-throughput live cell imaging setup combined with an automated evaluation pipeline to identify inhibitors of this process. Proof-of-concept experiments confirmed applicability: Inflammatory conditions of the BBB accompanied by impaired barrier properties significantly increased melanoma cell intercalation with the BBB when compared to control conditions of the BBB.

- Overview of CSF efflux sites in mice

- Inflamed condition of the in vitro BBB (pMBMECs, Il1b) led to significant stronger melanoma cell (B78%NOVA) intercalation compared to control (Ctl).

Schematic representation of the molecular composition of BBB tight junctions
Institute for Infectious Diseases (IFIK)

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www.ifik.unibe.ch

Profile
- Covers the entire spectrum of microbiology integrated in research, education and diagnostic services under one roof, including virology, bacteriology, mycology, parasitology, molecular diagnostic and infection serology.

Grants
- Swiss National Science Foundation: project grants 162583; 162808; 169791; 170063; 179260; 189136; NRP 72 grants 177452; 177378; 177386; Sinergia grant 180317; Bilateral programme South Africa grant 170844
- European Union: EUREKA Eurostars grant “Heartt”; Marie Sklodowska-Curie Action training network HONOURS
- University of Bern: Interfaculty Research Cooperations “One-Health” and “Decoding Sleep”
- The Gut Microbiology lab of Prof. Hapfelmeier was awarded participation in the new SNF NCCR “Microbiomes” (nccr.ch).

Highlights
- First accredited nanopore sequencing laboratory worldwide
- As of 21 January 2019, the IFIK is the first laboratory worldwide to have received the ISO/IEC 17025:2018 accreditation for sequence-based identification of bacterial pathogens using Oxford Nanopore Technologies devices in a medical diagnostic setting. The new sequencing technology enables to taxonomically identify pathogens more rapidly and cost-effectively. The IFIK sequencing laboratory is also the first certified Nanopore Service Provider in Switzerland.
- The Oxford nanopore sequencing service at IFIK

New research group: Experimental Virology

As of July, the Institute for Infectious Diseases (IFIK) has established the “Experimental Virology” research group headed by Prof. Ronald Dijkman who currently holds a dual appointment as Assistant Professor at the Vetsuisse Faculty in Bern and IFIK. His research has an emphasis on characterizing critical determinants of species barriers for emerging respiratory pathogens, such as influenza viruses, for both epizootic and zoonotic risk assessment and the development of effective countermeasures against these diseases.

INFECT by anresis App

ANRESIS collects antibiotic resistance data from 30 representative laboratories all over Switzerland. But collecting is not enough, data should be used and help in decision making. Therefore, ANRESIS developed an interactive, user-friendly app showing real time local resistance rates and treatment guidelines from the Swiss Society of Infectious Diseases in a single interface.

New zebrafish breeding and research laboratory to develop experimental models of infectious diseases

The IFIK inaugurated a zebrafish laboratory including a small breeding facility with a microinjection system and fluorescent imaging capability. This biosafety level 2 laboratory will allow the development of new experimental models of infectious diseases, including (but not restricted to) central nervous system and systemic infections.

Biosafety Center Bern

The biosafety center started its activities in August 2019. It is responsible for biosafety at sitem-insel and the ifik and advises both university institutes as well as partner institutions on questions related to biosafety. It offices are situated in the sitem-insel building. It will also conduct active biosafety research. Kümin D., Gsell Albert M., K. Summermatter: Applied Biosafety 2019: Volume: 24 issue: 4, page(s): 200-206.

Hindbrain infection of a zebrafish embryo by GFP-expressing Streptococcus pneumoniae

Biosafety Center, “Open House” day sitem-insel, 30 August 2019

3D image representation of human airway epithelial cells

The INFECT by anresis App in action
A new pathway to "reprogram" killer cells

The so-called natural killer (NK) cells are cells of the innate immune system that recognize and eliminate infected cells or cancer cells. During a virus infection, NK cells also keep the body's own immune cells such as the T cells at bay to avoid excessive killing of intact body cells. Cardoso Alves et al. found a novel function of the death molecule TRAIL. NK cells without TRAIL sense various signals from their environment differently: they react more strongly to a signal that causes them to release the cell toxin granzyme B (GZMB). Consequently, mice without TRAIL have more protective CD8+ T cells and are therefore better able to remove virus-infected cells. (www.embopress.org/doi/10.15252/embr.201948789).

TRAIL programs NK cells by blocking the production of inflammatory messengers (IFNs) but promoting the formation of cell toxins (GZMB). NK cells lacking TRAIL produce more IFNγ but less GZMB, which results in greater antivirus CD8+ T cell response in infected mice.

Development of ngTMA pipelines

We investigate the role of machine and deep learning in addressing clinically relevant research questions, predominantly in gynecological and colorectal cancers. Pipelines for handling data from next-generation Tissue Microarray (ngTMA) construction to assessment of immunostaining and integration of scores with clinical, histopathological and other research data for statistical analysis are performed by the group. We create and implement tools for quality control and optimization of digital workflows and apply these tools as a basis for our clinically oriented biomarker studies.

Example of digital p53 analysis in an endometrial cancer within an ngTMA experiment

Changes in local immune cells during onset vs. remission of colitis

Understanding the mechanisms that drive remission induction and maintenance in the intestine is critical for a rational treatment of patients with inflammatory bowel diseases. We have recently established a reversible, relapsing-remitting mouse model of colitis with reproducible onset of colitis and induced remission (Brasseit et al., Mucosal Immunol 2016). In this model we monitor the composition of the intestinal microbiota during relapsing – remitting colitis and define its consequences on the metabolic profile in the feces and the host. Furthermore, we investigate how these changes influence the host immune response and vice versa at a single cell level. An ultimate goal is to identify strategies to specifically extend the remission period, or even prevent further relapses of the disease.

In the healthy intestine, luminal bacteria are separated by a mucus layer (green) from the intestinal epithelium (blue), containing mucus-secreting goblet cells (green) and the lamina propria, containing numerous immune cells.

Understanding the role of autophagy in retinoic acid therapy of breast cancer

Epithelial-to-mesenchymal transition (EMT) plays a key role in therapy-resistance and metastasis formation. In the present study, we therefore aim at reversing the EMT phenotype of breast cancer cells using differentiation-based therapy based on all-trans retinoic acid (ATRA). Cellular differentiation is often associated with upregulation of autophagy. Autophagy is a lysosomal degradation and recycling system and may support cellular differentiation by removing superfluous organelles, keeping energy levels or by regulating signalling by selective removal of proteins. Therefore, we study autophagy functions during therapy-induced MET and how modulation of autophagy can support differentiation-based therapy. Furthermore, we investigate how cancer associated fibroblasts influence cancer autophagy and therapy efficiency.

Role of DAXX and ATRX in PanNET

Almost, half of Pancreatic Neuro-endocrine Tumors (PanNEs) show loss expression of DAXX or ATRX. We could show that DAXX/ATRX loss correlates with an increased risk of metastasis. DAXX and ATRX negative tumors show chromosomal instability and ALT (Alternative Lengthening of Telomeres) activation, a mechanism for telomeres length maintenance. We focus on unravelling the mechanism underlying this new cancer-associated pathway using in vivo mouse model, in vitro cell culture and ex vivo human patient tissues. We specifically focus on the role of epigenetic changes occurring in DAXX/ATRX negative PanNETs.

Heterogeneous telomeric size as hallmark of ALT positive cells. Circos plot depicting different DNA methylation and chromosomal instability in DAXX/ATRX negative and positive PanNETs.
Institute of Forensic Medicine (IRM)
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www.im.unibe.ch

Profile
- Seven departments conducting research projects within the context of forensic sciences
- External funding for the department of Forensic Medicine and Imaging, the department of Forensic Toxicology and Chemistry, the department for Physical Anthropology and the Forensic Psychiatry Service
- World leading in forensic physics and ballistics, forensic imaging and alcohol consumption markers
- Successful performance of forensic genetic projects, traffic medicine projects and forensic psychiatry projects
- Training and continuing education of students of medicine and jurisprudence, medical personnel and the institutions of the administration of justice and the police
- External Partners: Institute of Legal Medicine, University of Bonn; Center for Medical Image Science and Visualization, CMU, Linköping University, Sweden; Institute of Forensic Medicine, University Medical Center Freiburg; Department of Psychiatry, Psychotherapy and Psychosomatics, Psychiatric Hospital, University of Zürich; Netherlands Forensic Institute, Den Haag; Department of Forensic Anthropology, Center of Forensic Medicine, Medical University of Vienna, Austria; Max Planck Institute for the Science of Human History, Jena

Grants
- Swiss National Science Foundation (grant No. CR313L_157024; 320030_179466/1)
- Opioid maintenance therapy (OMT) in Belarus and Switzerland – a multistep project to enhance methodological knowl edge and strengthen competencies in the further education of young scientists and physicians in the area of OMT, Swiss State Secretariat for Education, Research and Innovation (SERI) and Swiss Science and Technology Cooperation (SSCT)
- Attitudes and beliefs of Swiss stakeholders towards patients with alcohol use disorder in Switzerland: comparison between primary care, specialists, insurance, and legal experts, Swiss Foundation for Alcohol Research Grant Program (SSA)

Highlights

Forensic Science International: Mind and Law
A new journal within the Forensic Science International family was launched with the aim of promoting the interdisciplinary exchange of ideas between psychiatric, psychological, legal, and other related fields within the forensic spectrum and beyond. «Forensic Science International: Mind and Law» will be edited by Michael Liebrenz, department of forensic psychiatry, and focus on mental disorders and how the sufferings of those affected are dealt with in the legal context in different regions of the world.

A new comprehensive Swiss dataset for forensic genetics
In the past, forensic genetics in Switzerland suffered from a lack of comprehensive data. To fill this gap, we established a biobank, systematically collected throughout the whole country including a population sample from Ticino. We then established autosomal allele frequencies for the 23 most commonly used STR loci. Statistical evaluation of the data revealed only small genetic differences among regional subpopulations and among language subgroups. The autosomal allele frequencies can therefore be used for forensics and paternity/tampering testing as a valid nationwide dataset.


Multidisciplinary research on human remains from international excavations
Egypt, Turkey, Russia, and Madagascar - the team of the Department of Physical Anthropology joins international excavations every year providing their expertise on human remains to different research projects. The remains come from single or multiple graves, as skeletons or mummies, from burial mounds or rock-cut chambers and cover a time span from pre-roman to early modern period. Apart from a thorough morphological investigation, samples for radiocarbon dating, stable isotope and ancient DNA analysis are taken in the field enabling interdisciplinary studies on diet, provenance, population genetics and kinship of the deceased.


Monitoring of direct alcohol markers
Direct and indirect biomarkers are widely applied for the determination of alcohol consumption. They help to assess past or present alcohol consumption. In this study, different sampling strategies for the intermediary long-term marker phosphatidylethanol (PEth) were evaluated. Volumetric capillary DBS sampling for PEth is a simple and useful tool for compliance monitoring.


Deforming bullet (32 auto) causing a pear-like shaped temporary cavity

Fundamental research in wound ballistics
The formation of the temporary cavity (TC) in 12 cm long 10% gelatine cubes was recorded using high-speed-video. This was compared to cross sectional analysis of the cracks left by the TC in gelatine. However, the profiles of the TC in high-speed video did not correspond to the destruction found in gelatine. Moreover, neither the damage in gelatine nor the profile of the TC did reflect the deceleration of the bullet. But TC and damage in gelatine correlated with the energy deposited. The elastic behaviour of the target model might be the cause of this observation.


External fundings for the department of Forensic Medicine and Imaging, the department of Forensic Toxicology and

MTR laser and 3D camera
Institute of Dental Medicine (ZMK)

Freiburgstrasse 7, 3010 Bern

Director and Head of the Clinic of Oral Surgery and Stomatology

Prof. Antion Sculean
Executive Director and Director of the Clinic of Periodontology

• Undergraduate dental curriculum (for 3rd, 4th and 5th year students) with theoretical and practical courses
• Postgraduate curricula (3-4 years) leading to specialty and / or MAS degrees
• Continuing Dental Education Master Courses
• Fundamental, translational and clinical research covering most aspects in dentistry
• WHQ Collaborating Centre for Epidemiology and Community Dentistry, Milan, Italy; Department of Cariology, Institute of Odontologie, Sallgrena Academy, University of Gothenburg, Sweden; Department of Operative Dentistry, Periodontology and Preventive Dentistry RWTH Aachen, Germany; Department of Odontologie, School of Dentistry, University of Copenhagen, Denmark; University of Texas Health, USA; University of Oslo, Norway; University of São Paulo – FDUSP / FIOH, Brazil; Indiana University, USA; Federal University of Minas Gerais, Brazil
• The Department of Reconstructive Dentistry provides the following range of treatment: Treatment planning and prosthetic restoration; Interdisciplinary collaboration with other departments for complex cases; Computer-assisted all-ceramic crowns, bridges or long-term temporary restorations (CAD/CAM); Advice and prosthetic treatment for implant patients; Optimization of dental esthetics
• The Division of Gerodontology offers synoptic dental treatment for the elderly and frail taking into account their individual needs and wishes. It provides also service to the community in three outreach clinics

Grants
• Endowed professorship for preventive dentistry and oral epidemiology of the Lutz Zürrer Foundation
• Swiss Government Excellence Scholarships for Foreign Scholars and Artists (ESKAS Scholarship; 2018.0515)
• Research Internships Abroad – São Paulo Research Foundation (BEPE–FAPESP; 2019/08437-5)

Highlights

Resin infiltration to mask caries lesion

In this clinical study we found resin infiltration technique a very useful method to mask caries lesions having developed during treatment with fixed orthodontic appliances. Colour changes while re-wetting the lesions with ethanol seem to be a valuable indicator for the number of required etching procedures.


Research on Tissue Integration of Biomaterials

A preclinical study of a novel collagen-based biomaterial used for soft tissue augmentation has shown a short inflammatory phase and rapid ingrowth of blood vessels and proliferating mesenchymal cells leading to fast tissue integration. These data form the biologic rational for the clinical application of this biomaterial for soft tissue augmentation around teeth and dental implants.

Laboratory for Oral Molecular Biology

Van der Woude syndrome (WWS; OMIM # 119300) is an autosomal dominant condition and represents the most common syndromic form of orofacial clefts with a prevalence of 1/3500 to 1/100’000 newborns. We are interested in a WWS patient, who harbors a novel undescribed disease-causing IRF6 mutation. Currently, we are establishing genotype-phenotype correlations using WWS patient-derived keratinocytes and over-expression systems of the pathogenic WWS-associated variant. Ultimately, our research will result in a better understanding of the pathogenesis of WWS and of IRF6 gene function.

(A) WWS patient with typical characteristics: cleft lip/palate and lip pits in the lower lip. (B) Genetic analysis revealed a novel undescribed pathogenic IRF6 (IRF6; WWS) mutation in the conserved protein binding domain (SMIR4/AD) resulting in a truncated IRF6 protein. (C) Forced expression of IRF6, WWS shows weaker expression of the mutated protein compared to the IRF6 wild type (wt).

Laboratory of Oral Microbiology

Grants: Innosuisse 26940.1 PFLS-LS
The laboratory evaluates in vitro new treatment options for periodontal and peri-implant therapy. Both standard microbiological methods as well as different biofilm models are used. A collaborative study of the three Swiss university laboratories of oral microbiology (Basel, Bern, Zurich) was finalized which determined antibiotic resistance incl. the mechanisms behind in two periodontopathogens.


Further, in vitro the effect of Porphyromonas gingivalis and its cysteine proteases (gingipains) on epithelial cell adhesion to titanium-zirconium alloy surfaces (a dental-implant surface) was investigated in collaboration with universities in Krakow (Poland) and Malmö (Sweden).

The lower the better? Data on cognitive decline in multinimod elderly patients and systolic blood pressure

Hypertension trials often exclude patients with multimorbidity and lack generalizability. We aimed in a Dutch cohort of >1,200 elderly patients to determine if systolic blood pressure (SBP) in patients undergoing antihypertensive treatment is associated with 1-year changes in cognitively functioning or quality of life (QoL) in persons aged 75 years with or without complex health problems. We found an association between low SBP (<130mmHg) and more cognitive decline in those who were treated with antihypertensive medication, and stronger among those with complex health problems. Our results suggest that SBP thresholds for treatment should be redefined, especially for frail older persons.


More refusals of colorectal cancer screening when only colonoscopy offered among patients in primary health care

Guidelines recommend primary care physicians (PCPs) offer patients a choice of colorectal cancer (CRC) screening methods. However, PCP screen mostly patients with colonoscopy and few screen with fecal occult blood testing (FOBT). We conducted a cross-sectional data collection among PCPs. 91 PCPs collected data on 3,637 50-75-year-old patients. PCPs who only offered colonoscopy had lower screening rates (47% vs. 71%) and higher refusal rates (44% vs. 20%) than PCPs who offered both methods. Encouraging PCPs to offer both methods could reduce the number of refusals and raise CRC screening rates.

Martin Y et al., J Gen Intern Med. 2019 : 34(8), 1409-1411.

The feasibility of Network Meta-analyses of Pharmacological and Non-Pharmacological Interventions in Psychiatry is challenging

The way how network meta-analyses (NMA) of pharmacological and non-pharmacological interventions in psychiatry are methodologically conducted is challenging. This could compromise their validity. In our meta review of 12 published NMAs, we found that the major issues are a) the definition and classification of the control node in the geometry of the network, b) differences in baseline participants’ characteristics (i.e. previous medication exposure), study risk of bias and the level of masking between studies with pharmacological and non-pharmacological interventions.

Del Giovane C et al., JAMA Psychiatry 2019.
Institute of Complementary and Integrative Medicine (IKIM)

Inselspital, Freiburgstrasse 46, 3010 Bern; von Roll Campus, Fabrikstrasse 8, 3012 Bern
www.ikim.unibe.ch

Prof. Ursula Wolf
Director

PD Dr. Stephan Baumgartner
Deputy Director

PD Dr. Johannes Fleckenstein

Profile

- 3 research groups
- Teaching students of medicine and dental medicine
- Patient centered translational and clinical research in complementary and integrative medicine; investigating pharmacological and non-pharmacological therapies, developing of new methodology
- Investigation of efficacy, effectiveness and mode of action of pharmaceutical preparations as used in Complementary and Integrative Medicine
- Investigation of underlying physiological mechanism; implementation of Chinese Medicine / Acupuncture technique into clinical routine
- External partners: Paracelsus Spital Richterswil, Switzerland; Hospital of Fribourg (HFR), Switzerland; Swiss Group for Clinical Cancer Research SAAK, Breast Cancer Group, Switzerland; EMPA, St. Gallen, Switzerland; NTB Buchs, Switzerland; Neuchatel Platform of Analytical Chemistry, University of Neuchatel, Switzerland; Society for Cancer Research, Arlesheim, Switzerland; Children’s Hospital of Eastern Switzerland, St. Gallen, Switzerland; Institute of Integrative Medicine, University of Witten/Herdecke, Germany; Department of Sports Medicine, Goethe-University Frankfurt, Germany; Department of Clinical Pharmacology, University of Regensburg, Germany; Department of Anaesthesiology, University of Greifswald, Germany; ISS Inc.y Champaign, IL, USA; Multidisciplinary Laboratory of Pharmaceutical Sciences, Faculty of Pharmacy, Federal University of Rio de Janeiro, Brazil

Grants

- Swiss National Science Foundation: SNF Bridge Discovery 20B2-1_180983 / 1 and 32473B_162717
- Swiss Cancer League, Cancer Research Switzerland: 4259-08-2017
- Swiss National Science Foundation: SNF Bridge Discovery 20B2-1_180983 / 1 and 32473B_162717
- University of Bern, UniBE Initiator Grant

Highlights

The Pulse-Respiration Quotient: A Powerful but Untapped Parameter for Modern Studies About Human Physiology and Pathophysiology

We describe why the PRQ is a powerful parameter that captures complex regulatory states of the cardiorespiratory system. We show that the PRQ (i) changes during human development, (ii) is time-dependent (ultradian, circadian, and infradian rhythms), (iii) shows specific patterns during sleep, (iv) changes with physical activity and body posture, (v) is linked with psychophysical and cognitive activity, (vi) is sex-dependent, and (vii) is determined by the individual physiological constitution. We explain why and how the PRQ could be used for basic research on human physiology and for applications in medicine.


Swiss paediatrician survey on complementary medicine

Paediatricians attitude towards complementary medicine (CM) is positive, emphasised by great interest in CM training, a willingness to contribute to CM research and a high rate of paediatricians who use CM for themselves and their families. The percentage of paediatricians offering CM is currently rather low despite strong demand for CM for children. This study provides key pointers for the future development of complementary and integrative medicine for children in Switzerland.


Effects of integrative medicine pharmaceuticals in a fingerprint metabolomic analysis

We observed compelling evidence for specific effects of pharmaceuticals produced according to European Pharmacopoeia monograph 1038 (Praeparationes homoeopathicae) by a fingerprint metabolomic analysis in a highly controlled bioassay. We see this as a challenge to develop theoretical models to explain effects of such pharmaceuticals.


Circular radial region of interest (CRROI) of fingerprint metabolomic analysis

Impact of succussion on pharmaceutical preparations

We applied the Droplet Evaporation Method to investigate the effects of mechanical energy on phytopharmaceutical preparations. For all investigated preparations, significant differences were found between succussed samples and controls, calling for mechanistic studies to further characterize the effects induced.


DEM image of succussed (right) and of unsuccussed (left) Echinacea extract

Complementary medicine and its role in sports medicine

This review highlights the use and evidence for hands-on conservative pain treatments in athletes. Among complementary approaches it is neural therapy (injection techniques) providing best evidence. Acupuncture can be considered a Grade B recommendation.

Panel below: Journal article cover page. The abstract is bilingual addressing English and French readers.

Fleckenstein and Banzer. Sci Spo 2019:34,e77-e100.
Institute for Medical Education (IML)

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Dr. Sandra Trachsel
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MME

Dr. Kai Schnabel
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Prof. Sissel Guttmann
Director RVL

Dr. Philippe Zimmermann
Department Head
ASCR

Prof. Sören Huwendiek
Department Head
AAE

Research on effective health provider communication

Despite scientific breakthroughs, cancer remains the top two causes of death in Switzerland. So, "communication about approaching death" remains an important task for oncology professionals. We aim at supporting those professionals in performing such communication with confidence and positive impact for all involved. A new interactive e-learning module supports a blended learning approach and practical, safe training. Learning effects are investigated with RCT design. Partners: Palliative Care Unit, Inselspital; Swiss Cancer League.
https://tinyurl.com/project-swiss-cancer-league

The IML participates in EU-project on clinical reasoning

Insufficient clinical reasoning (CR) skills are a major cause of cognitive errors in patient care. Yet, there is a lack of explicit teaching of CR in the training of healthcare professionals. To close this gap, an EU-wide project has been launched in which the University of Bern will work together with European partner universities and institutions to develop a CR-curriculum and a train-the-trainer course for lecturers.
https://tinyurl.com/Clinical-decision-making

The IML is committed to interprofessionalism

Healthcare professionals face increasingly complex situations. One way to overcome this challenge is to improve interprofessional collaboration. Together with the Bern Nursing Education Center and the Bern University of Applied Health Sciences, IML developed several interprofessional educational activities. Three of these were awarded by the Swiss Academy of Medical Sciences:
https://tinyurl.com/Story-Interprofessionalism.

Supporting Clinical Skills Training with new videos

To manage the increasing number of students, Clinical Skills Training (CST) in the 3rd year was substantially redesigned. Students now exercise Clinical Skills CST in examining their peers before they practice on real patients. To support that the students are performing the tasks correctly, videos demonstrating the CST-Tasks were produced by the AUM. The video-production was supported by students and experts of the different clinical disciplines. IML`s new video studio and team supports the high quality of the videos.

Profiles

- Competence centre for medical education, serving the medical faculty in Bern and other medical schools and institutions nationally and internationally
- Contributing to innovative and sustainable development in medical education, supporting development of competency in health professionals aiming at optimal patient care
- Four departments: assessment and evaluation, teaching and media, IT-development and MAS Medical Education
- Investigating effectiveness of assessment and learning aids, innovative methods and fields of applications
- Developing tools for assessment and learning, complying with needs from digital transformation

Grants

- EU-Project: «Developing, implementing, and disseminating an adaptive clinical reasoning curriculum for healthcare students and educators», Project coordinator: University of Augsburg. Project head: PD Dr. Inga Hege. Project partner: IML: Prof. Sören Huwendiek, MME, Dr. Felicitas Wagner
- Swiss Cancer League: «Communication with cancer patients and their families about approaching death: Scaffolding conceptual and practical learning for health professionals.» Project head: Prof. Sissel Guttmann. Project partner: Prof. Steffen Eychmüller, University Centre for Palliative Care (UCPC), Inselspital, University Hospital Bern
- ERS (European Respiratory Society)/PhD-Grant: «How to improve continuing professional development to foster physician’s competencies and patient treatment», PhD Advisor: Prof. S. Huwendiek, MME
- SGAIM: «Definition of Competencies for Attending Physicians in General Internal Medicine Departments in Swiss Hospitals: a multicenter qualitative study» SGAIM Foundation Award. Project partner: Dr. Christine Roten, MD, MME, Dr. Martin Perrig, MD, MME, Department of General Internal Medicine, Inselspital, University Hospital Bern, University of Bern, Dr. Christoph Berendonk, MD, MME, IML

Highlights

- Professionalization of the work with simulated participants
  Dr. med. B. Brem has been elected as Member of the Board of Directors of the Association of Standardized Patient Educators (ASPE). She will hold the position of Member Liaison for two years. ASPE engages in providing an international network for the new field of human simulation, develops “Standards of Best Practice” and basics for certification in working with simulated participants (SP).

- Practice cooperation in interprofessional teams
  © Picture BZ Pflege and IML 2019

- Communication is a key competence

- Strengthening teaching of CR

- Examination of lymph nodes (© Picture IML 2019)
Clinics at the University Hospital, Inselspital

- Department of Anaesthesiology and Pain Medicine
- Department of Angiology
- Department of Cardiology
- Department of Cardiovascular Surgery
- Department of Clinical Chemistry
- Department of Cranio-Maxillofacial Surgery
- Department of Dermatology (DERK)
- Department of Diabetes, Endocrinology, Nutritional Medicine and Metabolism (UDEM)
- Department of Diagnostic and Interventional Neuroradiology
- Department of Diagnostic, Interventional and Pediatric Radiology (DIPR)
- Division of Magnetic Resonance Spectroscopy and Methodology (AMSM)
- Department of Emergency Medicine
- Department of ENT, Head and Neck Surgery
- Department of General Internal Medicine
- Department of Geriatrics
- Department of Hematology and Central Hematology Laboratory
- Department of Infectious Diseases
- Department of Intensive Care Medicine
- Department of Medical Oncology
- Department of Nephrology and Hypertension
- Department of Neurology
- Department of Neurosurgery
- Department of Nuclear Medicine
- Department of Obstetrics and Gynecology
- Department of Ophthalmology
- Department of Orthopedic Surgery and Traumatology
- Department of Osteoporosis
- Department of Pediatrics
- Department of Pediatric Surgery
- Department for Plastic and Hand Surgery
- Department for Pulmonary Medicine
- Department of Radiation Oncology
- Department of Rheumatology, Immunology and Allergology (RIA)
- Department of Thoracic Surgery
- Department of Urology
- Department for Visceral Surgery and Medicine
Achieving gender parity in acute care medicine

In this review article we bring together articles from business and human resource sciences, recommendations for involving women in leadership and gender diversity programs, and the experience and knowledge of women in leadership positions, in an attempt to move toward Sheryl Sandberg’s vision of a world in which “there will be no female leaders – there will just be leaders.” This approach will enable lasting and meaningful progress toward a universal standard that will work to the benefit of everyone.


A nationwide survey

Ultrasound has significantly increased safety and effectiveness in regional anaesthesia. Our nationwide survey including all active members of the Swiss Society for Anaesthesiology and Resuscitation (SSAR/SGAR) revealed that dual guidance appears to be the preferred approach for safely localizing nerves for PNB in Switzerland.


Implementing Mini-CEX and DOPS for postgraduate medical trainees

The way in which educational tools such as the Mini-Clinical Evaluation Exercise (Mini-CEX) and Direct Observation of Procedural Skills (DOPS) are implemented is of key importance, but many factors influence their implementation. We aimed to develop a comprehensive analysis of such factors and found 13 relating to four categories: organizational culture (e.g. value of teaching and feedback), work structure (e.g. time for Mini-CEX and DOPS, faculty development), instruments (e.g. content of assessment), and users (e.g. relationship between trainees and supervisors), and their interaction.


ERK and p38 contribute to the regulation of nociceptin and the nociceptin receptor in human peripheral blood leukocytes

Mechanisms involved in the regulation of nociceptin and its receptor (nociceptin opioid peptide receptor, NOP) in response to inflammation and pain in humans are insufficiently studied. We investigated specific signaling pathways contributing to the regulation of nociceptin and NOP in human peripheral blood leukocytes and found that ERK and p38 are two major signaling pathways regulating nociceptin and its receptor in human peripheral blood leukocytes under inflammatory conditions.

Zhang L et al., Mol Pain 2019: 15.

FACS data of human leukocytes
Prof. Dr. med. Marc Schindewolf
• Förderungspreis Forschung 2019 der Union Schweizerischer Gesellschaften für Gefässerkrankungen (USGG): PD Dr. med. Bangerter Rhyner Grant (Dr. Luise Adam)

Canada
Research Initiative Institution, Cambridge, United Kingdom, McGill University, Department of Human Genetics, Montreal, Department of Pharmacological and Biomolecular Sciences, Milan, Italy, University of Cambridge, Cardiovascular Strategic
Zürich, Switzerland, University Hospital Munich, Institute of Epidemiology and Prophylaxis of Cardiovascular Diseases, Hospital Basel, Department of Angiology, Basel, Switzerland, University Hospital Zürich, Department of Angiology,

• External Partners: Vorarlberg Institute for Vascular Investigation and Treatment (VIVIT), Feldkirch, Austria, University

venous thromboembolism research of vascular malformations and artherosclerosis expression, drug therapy and endovascular management of risk factor analysis and risk factor modulating therapies in peripheral artery disease, integration of omics technologies in

• Projects comprise analysis, classification and computational hemodynamic modeling of congenital vascular malformations, risk factor analysis and risk factor modulating therapies in peripheral artery disease, integration of omics technologies in research of vascular malformations and atherosclerosis expression, drug therapy and endovascular management of venous thromboembolism.

• External Partners: Vorarlberg Institute for Vascular Investigation and Treatment (VIVIT), Feldkirch, Austria, University Hospital Basel, Department of Angiology, Basel, Switzerland, University Hospital Zürich, Department of Angiology, Zürich, Switzerland, University Hospital Munich, Institute of Epidemiology and Prophylaxis of Cardiovascular Diseases, Munich, Germany, University of Colorado Hospital, Heart and Vascular Center, Denver, Colorado, USA, University of Milan, Department of Pharmacological and Biomolecular Sciences, Milan, Italy, University of Cambridge, Cardiovascular Strategic Research Initiative Institution, Cambridge, United Kingdom, McGill University, Department of Human Genetics, Montreal, Canada

Grants
• Bangeter Rhynner Grant (Dr. Luse Adam)
• Förderungspreis Forschung 2019 der Union Schweizerischer Gesellschaften für Gefässerkrankungen (USGG): PD Dr. med. Marc Schindewolf

Highlights

Comparison of interwoven nitinol with drug-eluting stents for endovascular treatment of femoropopliteal artery disease

We analyzed efficiency of interwoven nitinol stents with drug-eluting stents for the femoropopliteal artery with regard to clinical-ly-driven target lesion reintervention at 12 months in patients with symptomatic PAD. Data was prospectively collected. The results do not provide conclusive evidence to favor one stent over the other. Haine A et al., Eur J Vasc Endovasc Surg. 2019 Oct 25. pii: S1078-5884(19)31465-0. doi: 10.1016/j.ejvs.2019.09.002.

Adherence to Statin Therapy Favours Survival of Patients with Symptomatic Peripheral Artery Disease (PAD)

Patients with PAD are at a very high cardiovascular risk. Guidelines recommend a strict lipid lowering therapy with statins. However, adherence to statins is generally low. In our single center study we observed 691 symptomatic PAD patients. Those never receiving statins had a significant higher mortality rate (31%) compared to patients con-tinuously on statins (13%) or new on statins (8%). Moreover, patients on intensified statin medication had a low mortality of 9%, whereas those who terminated statin medication or reduced statin dosage had a higher mortality (34% and 20%, respectively).

Our data suggest that adherence to statin therapy is associated with reduced mortality in symptomatic PAD patients. A strategy of intensive and sustained statin therapy is therefore recommended.

Dopheide JF et al., accepted for publication in European Heart Journal – Cardiovascular Pharmacotherapy in Dec. 2019.

(A) Achieved mean LDL-C reductions depending on treatment incentives. (B) Kaplan-Meier survival curves of patients on and off statins. (C) Kaplan-Meier survival curves of patients with a switch in statin dosage.

Broadening the categories of patients eligible for extended venous thromboembolism treatment.

VTE causes >500.000 death/year in Europe and the risk of recurrence after cessation of anticoagulation is up to 10%/year. Traditional treatment algorithms seem to be overly simplistic. More granular definitions of risk factor groupings, e.g. minor transient/persistent, environmental/non-environmental risk, in combination with new extended anticoagulation therapy options may provide refined management strategies for patients who commonly receive shorter-duration therapy.

Department of Cardiology

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Profile

- Teaching at multiple levels: clinical training, lectures for undergraduate and graduate students, supervision of MD, Master and PhD students
- Continuous medical education program
- Pre-clinical research (electrophysiology, arterial hypertension)
- Broad range of clinical research activities, encompassing investigations of devices for the minimal invasive treatment of coronary artery disease and valvular heart disease; pharmacological therapies in patients with acute coronary syndromes; studies for the treatment of electrophysiological disorders as well as clinical trials of medicinal products.
- External partners: University and large hospitals in Switzerland and abroad, ETH, CSEM, industry

Grants

- SNF Project “Efficacy of Permanent Internal Mammary Artery Occlusion in Stable Coronary Artery Disease: a Double-Blind, Randomized, Sham-Controlled Trial”, Prof. C. Seiler
- SNF Project “Assisted reproductive technologies-induced alterations of the cardiac phenotype in mice and humans, underlying mechanisms and long-term consequences”, PD Dr. E. Rexhaj
- SWISSHEART Failure Network (SHFN) - a Swiss Personalized Health Network (SPHN) Project (2018DRI14 PHRT122), Prof. Stephan Windecker
- SNF Project “Assisted reproductive technologies-induced alterations of the cardiac phenotype in mice and humans, underlying mechanisms and long-term consequences”, PD Dr. E. Rexhaj
- SNF-Spark “Value of transesophageal Phrenic Nerve Pacing” Dr. A. Häberlin

Highlights

- EVOPACS Trial: PCSK-9 inhibitors to reduce LDL-C in patients with Acute Coronary Syndrome (ACS)
  Decline of low-density lipoprotein cholesterol (LDL-C) levels effectively reduces the risk of adverse events in patients with established atherosclerotic cardiovascular disease. The clinical benefit of statins in improving clinical outcomes is proportional to the magnitude of LDL-C reduction and is more pronounced in patients with acute coronary syndromes (ACS) compared to stable coronary artery disease. A new randomized, multi-centric double-blind placebo controlled trial demonstrated the safety and the efficacy of a new PCSK-9 inhibitor on top of a therapy with statin to reduce the LDL level in patients with ACS. Koskinas KC et al., J Am Coll Cardiol. 2019 Nov 19;74(20):2452-2462.

- SCOPE trial: Comparing two transcatheter aortic valve replacement systems
  Transcatheter Aortic Valve Replacement (TAVR) represents an alternative treatment to surgery for patients with severe aortic valve stenosis. In this randomized, non-inferiority multi-centric international clinical trial, two commercially available TAVR devices were compared with regard to early safety and efficacy. The self-expanding device did not meet non-inferiority compared to the balloon expandable device. An early composite safety and efficacy endpoint was useful in discriminating the performance of the two devices. Lanz J et al., Lancet 2019 Nov 2; 394 (10209) :1619-1628.

- BISTEMI trial: Comparing two drug eluting stents

- Electrophysiology
  The DEEP SEDATION study was started, an investigator initiated randomized controlled trial comparing Desmethyleneadine versus Propofol in the catheter ablation of atrial fibrillation. For ventricular tachycardia ablation, the novel approach of using stereotactic beam radiation therapy was newly implemented as an interdisciplinary project between the Divisions of Radio-Oncology, Radiology and Cardiology. In cardiac pacing, novel pacing modalities such as His-Bundle Pacing as well as leadless pacing are studied in clinical projects at the Inselspital as well as in translational projects in the SITEM.

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Department of Cardiovascular Surgery
Inselhospital, Freiburgstr. 18, 3010 Bern
www.unibe.ch/ cardio

Profile

- Teaching at multiple levels: clinical training, lectures for undergraduate and graduate students, supervision of clinical and pre-clinical research work for MD, Master and PhD students
- Clinical and pre-clinical research: State-of-the-art research on multiple themes (eg. aortic pathologies, heart transplantation with donation after circulatory death, and cardiovascular fluid mechanics) is promoted through cooperation amongst clinicians, research scientists and biomedical engineers
- External partners: Integrated Actuators Laboratory - Zentrum für künstliche Muskeln, EPFL Lausanne, Microcity Neuchâtel; Cardiac Surgery, University of Alberta, Edmonton, Canada; Department of Transplantation, Royal Papworth Hospital, Papworth Everard, Cambridge, UK; Laboratoire de Signalisation et Physiopathologie Cardiovasculaire, INSERM, Université Paris Sud, Université Paris Saclay, Châtenay-Malabry, France; St. Vincent's Hospital, University of New South Wales, Victor Chang Cardiac Research Institute, Sydney, Australia

Grants

- A collaborative platform for artificial muscles (collaboration with EPFL Lausanne & Nanocity Neuchâtel), Siemens Stiftung
- Prevention of paraplegia in thoraco-abdominal aneurysms PAPA-ARTiS, EU Horizon 2
- A publicly available collection of virtual and physical 3D models of congenital heart defects for surgical planning and training, Swiss Heart Foundation
- Optimizing quality of cardiac grafts obtained with donation after circulatory death (DCD) using endothelial-targeted reperfusion strategies (Principal Applicant), Swiss Heart Foundation
- EXoCET: EXosomes based Combination Therapy to target multiple signaling within cardioprotective pathways (Co-Applicant) Swiss National Science Foundation (Project No. ZC020_182948)
- Role of mitochondrial damage-associated molecular patterns (mtDAMPs) in heart transplantation with donation after circulatory death (Principal Applicant), International Society for Heart and Lung Transplantation
- Cardioprotection with HOPE (Hypothermic Oxygenated PErfusion) in heart transplantation with Donation after circulatory death (Principal Applicant), International Society for Heart and Lung Transplantation

Highlights

Management of the aortic arch in patients with Loeys-Dietz syndrome

We present management strategies through a review of our clinical experience with the largest Loeys-Dietz patient population worldwide and a comparison with our experience in patients with Marfan syndrome.

Mitochondrial integrity during early reperfusion in an isolated heart model of donation after circulatory death

We report that mitochondrial dysfunction occurs with shorter periods of ischemia than those required to provoke contractile dysfunction and that mitochondrial function and circulating mitochondrial-related parameters can be used as specific indicators of cardiac ischemia-reperfusion injury.
Wyss R et al., J Heart Lung Transplant 2019,38:647-57.

Johanna Dürmüller-Bol DBMR Research Award and Jörg Vollmar-Prize from the German Society of Vascular Surgery and Vascular Medicine

Dr. Maria-Nieves Sanz was honoured for her research on mitochondrial-related inflammatory processes in heart transplantation. This is the second time that the Department of Cardiovascular Surgery was bestowed with this prestigious award in the last seven years.

The 2019 Jörg Vollmar-Prize from the German Society of Vascular Surgery and Vascular Medicine was awarded to Dr. Silvan Jungi. We report an estimated 4-year limb salvage of 84%, and excellent bypass patency rates despite poor runoff vessels. The considerable value and positive outcomes of open bypass surgery is emphasized. Jungi S et al., Eur J Vasc Endovasc Surg 2019,57:393-398.

Symptomatic or aneurysmal aberrant subclavian arteries: results of surgical and hybrid repair

Surgical and hybrid repair provides satisfying results; however, the optimal procedure must be defined on an individual patient basis.

Vertical Right Axillary Mini-Thoracotomy for Correction of Ventricular Septal Defects (VSD) and Complete Atrioventricular Septal Defects (CAVSD)

Vertical right axillary mini-thoracotomy is a safe and effective approach for the repair of VSD and CAVSD in selected patient groups, and the outcome data appear comparable to those with median sternotomy.
Galactose based culture reveals distinct metabolic signatures in pyruvate dehydrogenase and complex I deficient human skin fibroblasts

The bioenergetic profile of a cell influences the metabolism. Our results show that the switch from glucose to galactose based culture, leads to clearly distinctive bioenergetic profile of fibroblasts characterized by an increase in oxygen consumption (OCR) and decrease in extracellular acidification rate (ECAR). Further we could show that NMR metabolomic profiling revealed clear separation of mitochondrial defects from controls under galactose but not glucose using partial least squares discriminant analysis (PCA). Our results show the importance of selective culture methods and the influence of bioenergetics in discriminating normal from mitochondrial deficient cells using untargeted fingerprinting NMR profiles. Hertig et al., Metabolomics. 2019 Feb 28;15(3):32.

Discovery and validation of temporal patterns involved in TBI brain metabolism in human

Traumatic brain injury (TBI) is recognized as a metabolic disease with acute impaired glucose metabolism, thus the utilization of alternative energy substrates could be beneficial for the patients. We assessed and identified different states of brain metabolism during the acute post-TBI critical care phase using cerebral microdialysis. Using combination of metabolomics and metabolic modeling, ketometabolism was identified as one of the key components associated to the patients’ outcome, and the potential target for TBI neuro-repair. The identified metabolic signature could predict short- and long-term patient outcome.


The bleeding assessment tool of the ISTH is a useful diagnostic tool in patients with suspected platelet function disorders

Aiming to determine the diagnostic accuracy of the bleeding assessment tool (BAT) of the ISTH in clinical practice, we conducted a large prospective cohort study in 555 consecutive patients referred between 2012 and 2017 with a suspected bleeding disorder. Presence of a platelet function disorder was associated with substantially higher BAT scorings compared to patients without. Our data suggest that the ISTH-BAT provides a useful screening tool for patients with suspected platelet function disorders.

Bone Biology Lab

Focus of our research is the assessment of biphasic synthetic calcium phosphates and collagen-based scaffold for bone regeneration. The degradation kinetics and the formation of new bone can be affected by the level of biomimetic HA coating of α-TCP. Mineralized collagen, but not the collagen alone, enhanced healing of the critical-size bone defects. Preliminary data indicated that the mode of periosteal manipulation influence the nature and kinetics of ectopic bone formation.

Use of enamel matrix derivative liquid in combination with a natural bone mineral on new bone formation

We have developed a new liquid carrier system for enamel matrix derivate (EMD-liquid) for improved adsorption of EMD to biomaterial surfaces. Using the derivate of EMD-liquid, bone regenerative potentials have been investigated in combination with a natural bone mineral (NBM) in vivo. The Panel shows the images of toluidine blue and fuchsin staining on the middle sections in the bone defects at 8 weeks in NBM alone and NBM + EMD-liquid groups.


Effect of the degree of conversion of resin-based composites on cytotoxicity, cell attachment, and gene expression

Resin-based composite (RBC) restorations are routinely used in dental practice. We comprehensively investigated the influence of the degree of conversion (DC), resin-based composites RBC composition, and the effect of additional violet light from one light curing unit (LCU) on cell attachment/growth, eluate cytotoxicity, and gene expression on human gingival Fibroblasts (HGF). The Panel shows Giemsa staining of HGF cells attached on a total of 20 combinations of RBCs.

Fujioka-Kobayashi M et al., Dent Mater 2019: 35(8), 1173-1193.

Translational research on biodegradable magnesium plate/screw system for osteosynthesis

Our degradable magnesium osteosynthesis system consisting of plates and screws was successful for the midface fixation in a minipig fracture model. The degradation of a relatively large amount of magnesium did not lead to an accumulation of alloying elements in the organs. Hence, a significant step has been made for clinical application of magnesium-based osteosynthesis devices. Application of magnesium plates in the mandible is still a challenge because of strong mechanical loading. The magnesium plate/screw system has continuously been improved by the newly initiated mechanical test.

Profile

- Investigation how T cell metabolism and T cell function are related in inflammatory skin diseases
- Study of the autoimmune response and the molecular events leading to skin blistering in pemphigus
- Assessing the mechanisms of the interaction of plakin family members with various intermediate filaments in epithelia and stituated muscle cells
- Understanding how stem cells contribute to homeostasis of the skin
- Investigation of epithelial barrier dysfunction and type 2 inflammation
- Carrying out clinical trials with different targeted therapies for atopic dermatitis, psoriasis, hidradenitis suppurativa and melanoma
- External Partners: Genetic Skin Disease Group, King’s College London, London, UK; Institute for Research in Biomedicine, Bellinzona; Department of Dermatology, Huashan Hospital, Fudan University, Shanghai, China; Department of Immunology, University Hospital Zurich, Zurich, Switzerland; Department of Dermatology, University of Freiburg, Germany; The Regional Dermatology Training Centre, Tanzania; Department of Pathology, C.H. U. Henri Mondor, Paris, France; Department of Dermatology, University Hospital Düsseldorf, Düsseldorf, Germany; Hautklinik, Universitätsklinikum Marburg, Germany; Institute of Molecular Systems Biology, ETH, Zurich; Centro Studi GISED, Bergamo, Italy; Ludwig Institute for Experimental Dermatology, University of Bielefeld; Institute for Biomedical Techniques, Department of Information Technology and Electrical Engineering, ETH Zurich

Grants

- SNF Pernphrhus-from pathogenesis to therapeutics (Pegasus): Epitope spreading in pemphigus
- SNF Sinergia: A One health approach to unravel novel genes and molecular pathways in dermatology
- SNF Pemphigus-from pathogenesis to therapeutics (Pegasus): Epitope spreading in pemphigus
- Electrical Engineering, ETH Zürich
- Dermatology, University of Lübeck; Institute for Biomedical Techniques, Department of Information Technology and Institute of Molecular Systems Biology, ETH, Zurich; Centro Studi GISED, Bergamo, Italy; Lübeck Institute for Experimental Dermatology, University of Lübeck; Institute for Biomedical Techniques, Department of Information Technology and Electrical Engineering, ETH Zurich

Highlights

A next-generation Tissue Microarray (ngTMA) automated analysis to find new markers to assess prognosis of melanoma

Finding new markers to assess prognosis of melanoma without the necessity to perform a surgical interventions is an important goal in melanoma research. We have developed in a completely automatic and random procedure a ngTMA block containing tissues from melanocytic nevi, primary and metastatic melanoma to evaluate the potential markers. Then sections cut from ngTMA block undergo immunohistochemically stainings and expression of the proteins is investigated using fully automated analysis and compared among the study groups. Seyed Jafari, SM et al. PLoS One. 2018 Nov 8;13(11):e0207019.

Identification and characterization of pathogenic variants of desmoplakin

Desmoplakin (DSP) is a component of the desmosomes, cell-cell adhesion complexes. Genetic alterations of the DSP gene are associated with skin, hair and or heart pathologies. The genotype-phenotype relationship of DSP variants is still unclear. We have developed a biochemical method, which enables to identify deleterious versus silent variants in the tail domain of DSP. The tail of DSP tail anchors the intermediate filament network. Our results indicate that binding of DSP to epidermal keratin and the heart muscle-specific desmin relies on the same recognition sites. Deleterious DSP C-terminal variants, in a homoygous or compound heterozygous state, that are associated with a skin phenotype are most likely also pathogenic in the heart and lead to cardiomyopathy.


Association of vascular endothelial growth factor subtypes with melanoma patients’ prognosis

The expression of angiogenic factors, such as the Vascular Endothelial Growth Factor (VEGF) subtypes and their receptors is upregulated on vascular endothelial cells during tumor angiogenesis. By using semantic map analysis, we investigated the associations between VEGF subtypes and receptors by using next-generation Tissue Microarray, patients’ characteristics and prognosis in a cohort of 137 melanoma patients. Low and high values of VEGF subtypes were distinguished in two clusters in the map. In particular, high values of VEGF C, especially VEGF-C, were associated with thick MM, which was linked to disease-free survival and ulceration.

Cazzaniga S et al. Acta Derm Venereol. 2020 Jan 7;100(1).

Human "T₉⁺" cells are a subpopulation of PPARγ-T²₂ cells

While T₁, T₂, and T₁7 cells are well-defined T cell lineages in humans, it remains debated whether IL-9-producing T₁ cells represent a bona fide "T₉⁺" lineage. Our understanding of the characteristics of vascular endothelial cells during tumor angiogenesis is still unclear. We have developed a biochemical method, which enables to identify deleterious versus silent variants in the tail domain of DSP. The tail of DSP tail anchors the intermediate filament network. Our results indicate that binding of DSP to epidermal keratin and the heart muscle-specific desmin relies on the same recognition sites. Deleterious DSP C-terminal variants, in a homoygous or compound heterozygous state, that are associated with a skin phenotype are most likely also pathogenic in the heart and lead to cardiomyopathy.


Detection of PPARγ+ T helper cells in human allergic contact dermatitis. (A+B) Representative pictures of immunohistochemical staining for PPARγ and CD3 (A) or CD4 (B) in lesional skin of acute allergic contact dermatitis. Scale bars: 50µm. Open arrows: PPARγ-CD3+ cells (A) or PPARγ-CD4+ cells (B). Closed arrows: PPARγ-CD3 cells (A) or PPARγ-CD4 cells (B).
Closed-Loop and Artificial Pancreas

We are developing and evaluating a broad set of insulin-delivery algorithms in different target groups. Current studies address the safety, efficacy and utility of commercially available hybrid closed-loop systems (LINEAR: NCT03932630) for type 1 diabetes, the Cambridge algorithm for fully-automated closed-loop in type 2 diabetes (APRenaL: NCT04025775; Diabetes Obes Metab 2019; Lancet Diabetes Endocrinol 2019) and type 1 diabetes (APAKal: NCT07409554; Diabetes Care 2019). In parallel we are evaluating "Do-it-yourself" (DIY) systems using elaborate in-silico environments (Diabetes Obes Metab 2019).

Innovation in Nutritional Medicine and Research

Optimising nutrition support is a core interest of our clinical research. The EFFORT trial has shown that an adequate nutritional therapy in patients significantly improves clinical outcomes (NNT 25) and reduces mortality (NNT 37) (Lancet. 2019, JAMA Network Open 2019). Leveraging novel technology (computer vision, artificial intelligence) to further enhance efficacy, effectiveness and personalisation of nutrition care is amongst our top priorities (DEEP VISION).

Grants

- **SNF Sinergia Programm** (Project CR583_183569) Design and Evaluation of a Vehicle Hypoglycemia Warning System in Diabetes (HEADWIND Project) – PI Prof. C. Stättler
- SNF (320038_185019 / 1) Effects of SGLT-2 inhibitor dapagliflozin on hormonal glucose regulation and ketogenesis in patients with type 1 diabetes - PI Prof. M. Laimer
- Scheibath Foundation - PI Prof. M. Laimer
- SNF Marie Heim-Vögtlin - PI PD Dr. M. Balmer
- Dexcom ISS - PD Dr. Michel Laun
- Internal partners: ETHZurich, ETHLausanne, HSGallen, CSEM Neuchatel, UniFR, University of Cambridge, Manchester, Swansea (UK), Rotterdam, Padova, Cologne, Graz, Toronto, Duke, Yale

Profile

- Seven research groups covering the areas of: Diabetes Technology (artificial pancreas, closed-loop/hybrid-loop systems, smart sensing and devices, algorithms/artificial intelligence); Metabolic Science (exercise-related fuel metabolism, cardometabolic studies, immuno-metabolism, deep phenotyping, metabolic imaging, physiological/mechanistic drug research); Metabolic Modelling (in-vivo, in-silico); Computer Vision and Food Science (automated food recognition, biometrics, body shape/composition analysis); Peptide Analytics, Biomarkers and Metabolomics (LC/GC MS, dry blood spot analysis); Inborn Errors of Metabolism; Neuroendocrinology (central regulation, functional imaging), Neuroendocrine Tumors; Thyroid Disease (MALDI-TOF mass spectrometry imaging)
- Teaching students of medicine, dental medicine, pharmacy, biomedical sciences, graduates of the Graduate School for Cellular, and Biomedical Sciences
- Postgraduate education (CAS in sex and gender-specific medicine, Certificate Course Clinical Nutrition)
- External partners: ETHZurich, ETHLausanne, HSGallen, CSEM Neuchatel, UniFR, University of Cambridge, Manchester, Swansea (UK), Rotterdam, Padova, Cologne, Graz, Toronto, Duke, Yale

Highlights

**HEADWIND: Design and Evaluation of a Vehicle Hypoglycemia Warning System in Diabetes**

In collaboration with the ETHZ and the HSG (SNF sinergia program), we lead a program applying artificial intelligence to analyze driving behavior during hypoglycemia based on car-derived parameters as well as physiological parameters. The overall target is to detect and predict hypoglycemia while driving in real-time. To this aim, we induce hypoglycemia while human participants are driving first in a simulator and then in real cars. (SNF-sinergia CRSII5_183569; SGED 2019; ATTD 2020).

For further information, please visit our website: www.udem.insel.ch

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Highlights

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For further information, please visit our website: www.udem.insel.ch

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Profile

- Neurovascular research: The research group focuses on treatment strategies for neurovascular diseases (aneurysms, arterio-venous malformations (AVMs) etc.) and especially the evaluation and treatment of acute ischemic stroke. The researchers combine preclinical development of techniques in bench-top and animal models and evaluate their application in large-scale international clinical studies (Swift direct). Furthermore, the group evaluated robotic approaches in neurovascular treatment.
- Advanced Neuroimaging and Artificial Intelligence Technology in Neuroradiology: The research of the Support Center of Advanced Neuroimaging (SCAN) focuses on the translation of quantitative imaging methods, artificial intelligence technology and automated image analysis into clinical practice, sequence development and methodological counseling for neuroimaging studies. In this domain, the researchers of the SCAN have participated in international neuroimaging trials (e.g. ENINGMA Epilepsy and Parkinson’s disease), neuroimaging challenges and data analysis networks in cerebrovascular research, neuro-epidemiology, sleep-related disorders, epilepsy and neurodegenerative disorders.
- CSF Disorders: The multidisciplinary group of the Neuroradiology develops new imaging modalities and interventional/surgical treatment options for patients with CSF leakage syndromes and idiopathic intracranial hypertension
- Ultra High Field MRI: MR-physicists and clinical neurologists support the Translational Imaging Center (TIC) at the stem-inel in MR methodology and explore new indications for UHF MRI in CNS disorders (as e.g. structural epilepsies, neuroimmunological and vascular disorders)
- Teaching activities: Medical Faculty, Faculty of Human Sciences, Biomedical Engineering, University of Applied Science Bern, European Academy of Neurology (EAN), European Stroke Organisation (ESO), Medical Image Computing and Computer Assisted Intervention Society (MICCAI)
- External Partners: Department of Neurology, UCLA, USA, Excellence Center for Ultra-High Field MRI, Austria, Department of Neuroradiology and Informatics, Technical University Munich, Germany, Division of Neuroradiology, Western Ontario Hospital Toronto, Canada

Grants

- SINERGIA Project “Predict and Monitor Epilepsy After a First Seizure: The Swiss-First Study” (PI Prof. Wiest)
- SINERGIA Project “Hydroid” (PI Dr. Mosimann)
- SNF Project “Introduction of High Field Optimized Fast 3D MR Spectroscopic Imaging for IDH Typing of Gliomas and Assisted Surgical Neuro Navigation” (PI Dr. Slotboom)
- HORIZON 2020 INSPIRE MED (CI Dr. Slotboom)
- Novartis Innovation Grant (PI Dr. McKinley)
- SFIN Grant “Imagine” (USZ, UZH, Inselspital, USB, EOC, CHUV, HES-SO.Local PI Bern: R. Wiest)
- SNF SPARK “Using Deep Neural Networks to Bridge Clinical and Quantitative Analysis of Intracranial EEG in Epilepsy”
- SISF Grant UValése Translational developments of sequence and post-processing techniques for clinical applications of chemical exchange saturation transfer imaging (CEST) for ultra-high field MRI - a use case for a public-private partnership between TIC and Olea Medical®
- Heart Foundation (PI PD Dr. P. Mosimann)

Highlights

The Translational Imaging Center at the stem-inel has become operational by the 1st of July 2019. The Bernese system is one of the first clinically approved UHF systems in the world and enables researchers to investigate specific pathologies obscured by conventional MRI (as e.g. subtle cortical dysplasias in epilepsy or cortical MS plaques). The TIC is granted support and expertise in MR physics by Siemens Healthineers, funding of the SNF and SISF, whereby the first scientific projects could be started in 2019.

The Peter-Huber Preis of the Swiss society for neuroradiology has been granted to Dr. Thomas Dobrocky: “Assessing Spinal Cerebrospinal Fluid Leaks in Spontaneous Intracranial Hypotension With a Scoring System Based on Brain Magnetic Resonance Imaging Findings”.

Certificate of Advanced Studies in Neuroimaging (CAS)

The SCAN and the stem school at the Swiss Institute for Translational and Entrepreneurial Medicine have conceptualized a joint program to educate physicians in the emerging field of AI technology in imaging. The main objective of the CAS is to equip Medical Doctors with the necessary skills to take a leading role in the AI-driven transformation of medicine. The course program was developed for medical professionals and provides them with the necessary knowledge and skills to translate medical problems to data science problems. By working on an AI project of their own, the participants will know the fundamentals of Artificial Intelligence and can thus judge the feasibility and adequacy of proposed AI solutions.
A new predictive indicator in the diagnosis of endometriosis

The «Cloverleaf Sign» has been discovered on MRI and it has been shown that it accurately predicts significant longer OR times, increased intraoperative blood loss, as well as a higher rate of bowel resections. Therefore, it has a profound impact on OR planning and patient counselling.


Gadolinium-Based MRI Contrast Agents Induce Mitochondrial Toxicity and Cell Death

This was the first study conducted in human neurons to assess for toxic effects on cellular function from MRI contrast agents routinely used in clinical practice. The results demonstrated that agents with lower kinetic stability broadly induce higher rates of cell death and greater impairment of mitochondrial function. Bower et al., Invest Radiol. 2019 Aug;54(8):453-463.

Liver MRI susceptibility-weighted imaging (SWI) compared to T2* mapping in the presence of steatosis and fibrosis

We demonstrated that both the SWI and T2* relaxation times are strongly dependent on the liver steatosis grades, while SWI is more dependent than T2*. Both parameters are useful predictors for liver fibrosis when using a multiparametric approach. However, the stage of fibrosis may be overestimated in the presence of steatosis, which should be taken into account when using SWI and T2* in the assessment of fibrosis.


Improved identification of DNA double strand breaks: γH2AX-epitope visualization by confocal microscopy and 3D reconstructed images

Currently, in the context of radiology, irradiation-induced and other genotoxic effects are determined by visualizing DSB-induced DNA repair through γH2AX immunofluorescence and direct counting of the foci by epifluorescence microscopy. This procedure, however, neglects the 3D nature of the nucleus. Confocal microscopy and 3D reconstruction enables an improved identification of double strand break induced γH2AX foci, allowing for an unbiased, ameliorated quantification. Ruprecht N et al., Radiat Environ Biophys. 2019 May;58(2):295-302.

Monitoring the tumor burden in vivo by optical imaging in a xenograft SCID mouse model

We transfected the human melanoma cell line MeWo with pEGFP-C1 or pDsRed1-N1, and investigated solid xenografts and metastases. EGFP-/DsRed-xenograft tumors had identical growth kinetics. In vivo the emitted light by DsRed tumors/metastases was much brighter than by EGFP. DsRed metastases were earlier (3 vs. 5 weeks) and much more sensitive detectable than EGFP metastases.

Division of Magnetic Resonance Spectroscopy and Methodology (AMSM)

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Prof. Roland Kreis
Prof. Peter Vermathen
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Dr. Gabriele Bonanno
Dr. Arun Joseph
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Makie Hoefemann
Martyna Dziadosz
Kadir Simak
Damian Hertig

Profile

- Magnetic resonance imaging (MRI) and spectroscopy (MRS) methods in collaboration with clinical partners in prospective studies of different organs
- Development of novel methods to suit pertinent needs to study physiology and pathology
- MRI and MRS studies in brain, kidney, muscle, liver, and heart
- High resolution NMR studies on biopsies, cell cultures and body fluids
- Participating in University teaching programs for students of medicine, chemistry, biochemistry, and biomedical sciences
- External Partners: Max-Planck-Institute for Cognition- und Neurosciences, Leipzig, Germany; Max Planck Institute for Biological Cybernetics, Tuebingen, Germany; University Children's Hospital and Children’s Research Center, Zurich, Switzerland; Department of Physiology, University of Lausanne, Lausanne, Switzerland

Grants

- Swiss National Science Foundation (grant No. 320030_175984; 320030_170062)
- Marie-Curie Initial Training Network (H2020-MSCA-ITN-2018-813120)

Highlights

Overview

MRI and MR spectroscopy (MRS) are powerful and extremely versatile methods for non-invasive studies and diagnostic examinations in humans. We use MRI and MRS methods in collaboration with clinical partners primarily in prospective studies of different organs. We develop novel methods to suit pertinent needs to study physiology and pathology, together with the underlying mechanisms, in situ. Most current studies are performed in brain, kidney, muscle, liver, and heart (see figure). In addition, high resolution NMR studies are performed on biopsies, cell cultures, body fluids.

Effect of short term exercise on ectopic lipid reservoirs in liver and muscle

European Innovative Training network (ITN)

The newly funded Marie Curie ITN INSPIRE-MED (inspire-med.eu) has started this year. It has a focus on multi-parametric and multi-modal MR/MRS and PET techniques in a European multicenter research and training network. The specific aim of our subproject includes novel synchronized acquisition and processing methods aiming at standard modeling evaluations in contrast to the use of dictionaries and machine learning methods.

Renal Function

Renal Function in native and transplanted kidneys has been investigated by multi-modal MRI and MRS in preparation of a Sinergia Grant. In collaboration with the Nephrology Department we aim at a better perception of the physiologic basis behind functional MR-parameters and why they may be changed in renal disease. Reproducibility and comparability studies have been performed. In clinical studies fMRI measurements were performed comparing living-donor versus deceased-donor kidney transplantation. Funded by sitem-insel Support Funds and in collaboration with the Nephrology we are currently preparing a study for localized non-invasive in vivo determination of potassium by 39K MR Imaging and Spectroscopy on the 7T MR-Scanner.

High-Resolution Magic Angle Spinning NMR

High-resolution magic angle spinning (HR-MAS) NMR techniques were applied to correlate in vivo and in vitro NMR spectra of tissue and body fluids. HR-MAS studies have been performed on biopsies and on cell cultures and analyzed by “metabonomical” methods. In collaboration with the vendor Bruker we established a perfused bioreactor system and performed measurements of living 3D cell cultures inside the NMR with changing conditions.

Brain Physiology

An SNF grant aims at MR acquisition and postprocessing methods that are tailored to the observation of brain metabolism, yet are also transferable to other organs. Exchange processes between amide protons and water are studied. MRS is also optimized for reproducibility in longitudinal studies and for detection of low-concentrated metabolites like phenylalanine and NAD+. Diffusion properties of brain metabolites are investigated with dedicated methodology (see figure) in collaboration with the Clinic for Neurology to elucidate microstructural properties where we make use of the fact that metabolites, in contrast to water, are essentially confined to intracellular space. In collaboration with a Max Planck Institute in Leipzig, where we use a MRI system with the strongest gradient system available for human research, we try to identify macromolecular signals to aid clinical MR spectroscopy.

Group logistics

After two years of “exile” at Erlachstrasse, we moved back to Freiburgstrasse, where our previous home, the old nephrology pavilions, have made room to the new sitem-insel building, where we moved into new offices and where we are part of the novel translational imaging center (TIC). This is equipped with a high-end 3T MR scanner, but also one of the worldwide few clinical 7T whole body MR scanners. With the move, the group now also hosts two Siemens MR Research Scientist (Dr. Bonanno and Dr. Joseph) in charge of support of clinical researchers from Inselspital performing research at 7 Tesla.

Arrival of one of the new MR scanners for the Translational Imaging Center at sitem-insel

Effect of diffusion time on brain spectra

Effect of short term exercise on ectopic lipid reservoirs in liver and muscle

Bioreactor for online Metabolomics of living cells inside the NMR.

Oscillating Gradient Spin-Echo (TD= 8.3 ms)

Pulsed Gradient Spin-Echo (TD= 155 ms)
3rd Swiss symposium on refugee and migrant health

On 13 December 2019, we host the 3rd Swiss symposium on refugee and migrant health. The annual symposium is now a well-established event in the communities of research and practice in the field. This year, the focus will be on the social aspects and the integration of refugees and migrants. The health of young adults and the work of “doctors without borders” are also key topics. In addition the importance of the social circumstances of asylum seekers and migrants will be addressed. We aim to achieve a more holistic view on the health of refugees and migrants.

www.telenotfallmedizin.ch

Department of Emergency Medicine

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• Teaching medical students, residents and postgraduate nursing students in emergency medicine and point of care ultrasonography
• 7 research groups
• Investigation of research questions with direct relevance to emergency patients
• EURAC Bozen, CHUV Lausanne, HUG Genf, Max Planck Institute for Human Development Berlin, Institute of Educational Measurement Oslo, Sarah Lawrence College New York, Charité Berlin, Erasmus Medical Center Rotterdam

Grants

• SNF and industry funding
• Foundational professorship for tele-emergency medicine and e-/m health
• BAG funded projects in migrant health
• Foundational career funding for young clinical scientists

Highlights

Foundational Professorship in tele-emergency medicine

Thanks to the support of the Touring Club Switzerland (TCS), the University of Bern is able to establish an assistant professorship for tele-emergency medicine. The professorship is concerned with “eHealth” in the field of emergency medicine. It is based at the University Emergency Centre (UNZ) at the Inselspital, University Hospital Bern. The professorship is unique in its academic orientation in German-speaking Europe and one of the very few in the world.

With the support of a professorship for tele-emergency medicine, the increasing bottleneck in Swiss emergency care is to be counteracted, which is aggravated by the ageing population and the lack of doctors.

The endowed professorship will be advertised nationally and internationally; according to the UNZ and TCS, it is expected to be filled at the beginning of 2020.

Swiss symposium on tele-emergency medicine and digital health

The rapid development of digitalisation in acute medicine encompasses all areas of medicine and all medical professionals in their everyday working environment from the use of apps and electronic decision aids with artificial intelligence to the support of rural areas through telemedically supported treatment systems. Whatever one may think of this development, it affects everyone who works in emergency medicine. Opportunities, such as the potential mitigation of a shortage of physicians in emergency care, and risks, such as the frequent lack of digital education, privacy concerns, and a lack of academic justification in the application of Digital Health products, must be addressed and discussed.

The department thus hosts the first Swiss symposium on tele-emergency medicine and digital health on 13 February 2020. www.telenotfallmedizin.ch
Research meets patient organisation

The Hearing Research Laboratory also focuses on the development of improved tinnitus diagnostics. The research work was presented at the public information event “Tinnitus - Development and Strategies for Treatment”. The event was organized by pro audito bern and the ENT Clinic. The lecture hall was well occupied showing the large interest in this topic.

Cochlear Morphology Characterization

In collaboration with the INRIA research group of the Université Côte d’Azur, France, we developed a novel method to characterize the shape of the cochlea. The method can be used to improve surgical planning approaches and to refine the postoperative fitting for cochlear implant patients.


Politzer Society Award 2019

Dr. Stefan Weder was nominated for the Politzer Prize in the category “best paper in clinical science” with his research project “Towards a Better Understanding of Electrocochleography: Analysis of Real-time Recordings” and won the gold medal.

DGA: Young Scientist Award 2019

The Politzer Society was founded in 1978 and has held an international conference every second year. The scientific focus is on research and the latest developments in otology and neurotology. This year, the 32nd meeting took place together with the World Congress in Otology in Warsaw, Poland. Dr. Stefan Weder was nominated for the Politzer Prize in the category “best paper in clinical science” with his research project “Towards a Better Understanding of Electrocochleography: Analysis of Real-time Recordings” and won the gold medal.

Politzer Society Award 2019

At the annual conference of the German Society of Audiology (DGA) Tim Fischer received the Young Scientists Award 2019. The prize was awarded for his research work on the influence of microphone directivity on the spatial hearing ability of bilateral cochlear implant wearers.

Cochlear Morphology Characterization

In collaboration with the INRIA research group of the Université Côte d’Azur, France, we developed a novel method to characterize the shape of the cochlea. The method can be used to improve surgical planning approaches and to refine the postoperative fitting for cochlear implant patients.

Department of General Internal Medicine

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**Profile**

- **Research focus:** multimorbidity, venous thromboembolism, anticoagulation, variation in care, overtreatment, thyroid diseases, primary care, epidemiology, public health, chronic pain, opioid use, drug metabolism and safety, nicotine dependence, pharmacolearning, machine learning
- **8 research groups**
- **Overall aim:** To improve quality of care and to promote evidence-based, high-value, safe, patient-centered, data-informed, and sustainable health care
- **Teaching:** (lectures, clinical teaching, skills training, small group teaching) and assessment (practical and theoretical examinations) of students in medicine and dental medicine
- **High-quality post-graduate training of general internal medicine and specialty residents**
- **International academic partners:** Division of General Internal Medicine, University of Pittsburgh Medical Center, USA; Department of Medicine, Ottawa Hospital Research Institute, University of Ottawa, CA; Departments of Primary Care/Public Health and Internal Medicine, Leiden University Medical Center, NL; Division of Hospital Medicine and Clinical Pharmacology/Experimental Therapeutics, Department of Medicine, Department of Epidemiology and Biostatistics, and Center for Tobacco Control Research and Education, University of California, San Francisco, USA; The Thyroid Studies Collaboration on 5 continents; Department of Neurobiology, Care Sciences and Society, Institute of Environmental Medicine, Division of Physiotherapy, Karolinska Institutet, Stockholm, Sweden; Occupational and Industrial Orthopaedic Collaboration on 5 continents; Department of Neurobiology, Care Sciences and Society, Institute of Environmental Medicine, Division of Physiotherapy, Karolinska Institutet, Stockholm, Sweden; Occupational and Industrial Orthopaedic Center, New York University Langone Orthopaedic Hospital, New York, USA; Division of Research, Kaiser Permanente Northern California, Oakland, USA; The European Drug Emergencies Network Research Group; Global Health Institute Barcelona, Spain; Kenya Medical Research Institute, Nairobi, Kenya
- **Swiss academic partners:** Venous thromboembolism network including 22 Swiss university and non-university hospitals; Institute of Biostatistics, Epidemiology, Biostatistics and Prevention, University of Zürich; Department of Health Sciences, Helbling, Dübendorf; Horten Center for Patient Oriented Research and Knowledge Transfer, University of Zürich; Department of Internal Medicine, Centre Hospitalier Universitaire Vaudois, Lausanne; Swiss Tropical and Public Health Institute, Basel

**Grants**

- Swiss National Science Foundation (grant no. 33C30_185616, NRP74 407440_167339/1, 179346, 320030_172676, 32003B_189132, 31003A_160206, 32003B_179346)
- OPERAII- European Union’s HORIZON 2020 (grant agreement 634238)
- Novartis Foundation for Medical-Biological Research, Bangerter-Rhyner Foundation, Swiss Society of General Internal Medicine Foundation, Eurospine Task Force Research Grant, UNITAID: BOHEMIA (Broad One Health Endectocide-based Malaria Intervention in Africa)
- Intramural grants: CTU grant (Inselspital), UnibE ID Grant

**Highlights**

**Skeletal Effects of Levothyroxine for Subclinical Hypothyroidism in Older Adults: A Randomized Trial**

Both thyroid dysfunction and levothyroxine therapy (LT4) have been associated with bone loss, but studies on the effect of LT4 for subclinical hypothyroidism (SHypo) on bone yields conflicting results. In a nested sub study of the double-blind placebo-controlled TRUST trial, 196 participants with SHypo were randomized to LT4 with dose titration vs. pla.cebo. The unadjusted 1-year change in lumbar spine bone mineral density (BMD) was similar between LT4 (+0.8%) and placebo-treat- ed groups (+0.6%); between-groups difference +1.4%: 95% CI -0.1 to 2.9, p=0.059). Likewise, there were no between-group differences in 1-year change in Trabecular Bone Score (-1.3%: 95% CI -3.1 to 0.6, p=0.19) and total hip BMD (+0.2%: 95% CI -1.1 to 0.1, p=0.61). Over 1-year levothyroxin had no negative effect on bone health in older adults with SHypo. J Clin Endocrinol Metab. 2020 Jan 1;105(1). pii: dgz058. doi: 10.1210/clinem/dgz058.

**Aspirin compared with anticoagulation to prevent venous thromboembolism after knee or hip arthroplasty: a large retrospective cohort study**

Guidelines recently allowed the use of aspirin as an alternative to anticoagulants for venous thromboembolism (VTE) prophylaxis in patients undergoing knee or hip arthroplasty, but there is limited data on contemporary use and outcomes with aspirin in real-world practice. In a large retrospective cohort study using data from >100,000 patients undergoing knee or hip arthroplasty in the US MedAssets Health System, more than a fourth of all patients received aspirin as the sole anti thrombotic agent postoperatively. Thromboprophylaxis with aspirin-only was not associated with a higher risk of postoperative VTE compared with anticoagulants after knee or hip arthroplasty. J Gen Intern Med 2019;34:2038-2046.

**Effects of Nicotine Metabolic Rate (NMR) on withdrawal symptoms and response to cigarette smoking after abstinence**

The NMR is a phenotypic biomarker that is highly correlated with the rate of nicotine clearance, an important determinant of smoking behavior and nicotine dependence. In our study fast metabolizers by NMR had lower blood nicotine concentrations and greater craving/with- drawal scores compared with slow metabolizers, but not greater reward after smoking, thus supporting the idea that fast metabolizers are likely smoking more to relieve craving/withdrawal symptoms. Selection of medications and/or doses of medications guided by NMR may be useful in optimizing smoking cessation therapies. Clin Pharmacol Ther. 2019;105:641-51.
New standards for measuring muscle and bone function

For assessment of muscle function, standardized tests are needed to measure movement and strength of upper and lower extremities. However, many tests have been insufficiently standardized or are not feasible for use in clinical settings. Based on systematic reviews, we propose a new standardization of gait speed assessment in older people. In addition, imaging techniques give additional information on muscle and bone quantity and quality. We are working on standards for measuring sarcopenia with body impedance analysis. In a joint collaboration with the CTU unit, the ARTORG center, and the osteoporosis unit, we were able to develop a new standard for measuring bone strength (HR-pQCT-Based Distal Radius and Tibia Strength).


New technology for sensory assessment in hospital setting

Sensory function deficits are often not recognized at the time of hospital admission in older patients. However, reliable information on hearing and vision is key for optimal patient management, to ensure adequate patient-provider communication and to avoid in-hospital complications such as delirium and falls. In 2019, in a joint project with the audiology unit, we evaluated a prototype device using a calibrated voice recording as an alternative to the whispered voice test, and measured speech levels in patient-provider communication settings. The ultimate goal is to develop a feasible and clinically relevant brief hearing test for patients at hospital admission.

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Grants
- Swiss National Science Foundation (grant No. 310030-185233; 314730_173127; R’EQUP 316030_183501)
- SNF Professorship (PP0DP_183721)
- Landsteiner Foundation for Blood Transfusion Research (LSBR1719), Amsterdam, The Netherlands
- Product and Process Development Cellular Products Sanquin (PPOC) program Sanquin (PPOC-14-010, PPOC-15-010, PPOC-16-34, PPOC-17-44), Amsterdam, The Netherlands
- Dutch Thrombosis Foundation (TSN 201604), Amsterdam, The Netherlands
- Health Services Research grant, Swiss Cancer Research (HSR-4085-11_2016)
- 3rd Call for Proposals for Personalized Health and Related Technologies RHRT project #2019-717

Profile
- Teaching students of medicine, dental medicine, pharmacy, biomedical sciences as well as graduate students at the Graduate School of Graduate School for Cellular and Biomedical Sciences (GCB)
- 9 research groups
- Investigation of epidemiological and pathophysiological processes as well as diagnosis, prognosis and therapeutic approaches of blood-related disorders, pathophysiological processes that contribute to inflammation and tumor diseases
- External partners: INSERM & UMR-S 1176, Université Paris-Sud, University Paris-Saclay, Le Kremlin-Bicêtre, France; Department of Molecular Medicine, The Scripps Research Institute, La Jolla, CA, USA; Division of Developmental Therapeutics, Research Centre for Innovative Oncology, National Cancer Center Hospital East, Chiba, Japan; Department of Biochemistry, Cardiovascular Research Institute Maastricht, Maastricht University, Maastricht, The Netherlands; Institute of Research in Immunology and Cancer, University of Montreal, Montréal, Canada; Division of Hematology/Onco, Boston Children’s Hospital and Department of Pediatric Oncology, Dana-Farber Cancer Institute, Harvard Medical School, Boston, MA, USA; Steering committee and Advisory Board of Hereditary TTP Registry (www.ttpregistry.net); Department of Internal Medicine- Infectious Diseases and Center of Molecular Medicine, University Medical Center Amsterdam, The Netherlands; Department of Experimental Immunohematology, Sanquin, Amsterdam, The Netherlands; Departments of Biochemistry & Molecular Biology and Pathology, University of Oklahoma Health Sciences Center, Oklahoma, OK, US; EBMT SAA Working Party, Late effects working party of the ASTCT, DCEG National Institutes of Health National Cancer Institute, Rockville, Bethesda, USA

Highlights
The International Hereditary Thrombotic Thrombocytopenic Purpura Registry
Congenital thrombotic thrombocytopenic purpura is an autosomal recessive inherited disease with a clinically heterogeneous course and an incompletely understood genotype-phenotype correlation. In 2006, the Hereditary TTP Registry started recruitment for a study which aimed to improve the understanding of this ultra-rare disease. We found a larger proportion of compound heterozygous than homozygous carriers of ADAMTS13 c.4143_4144dupA with overt disease onset at <3 months of age, despite the fact that ADAMTS13 activity was <16% in 18 of 20 homozygous, but in only 8 of 14 compound heterozygous carriers. An evaluation of overt disease onset in all patients with an available sensitive ADAMTS13 activity assay shows that residual ADAMTS13 activity is not the only determinant of age at first disease manifestation. Registered at clinicaltrials.gov identifier NCT01257269.
Van Dorland HA et al., Haematologica 2019 : 104, 2101-2115.

Complement C3 inhibition by compstatin Cp40 prevents hemolysis
Our results show that Cp40 can inhibit complement-mediated extravascular hemolysis of autoimmune hemolytic anemia (AIHA) sera opsonized RBCs. In addition our data show that complement regulation by Cp40 can be even more beneficial since both intra- and extravascular hemolysis are prevented. Together, we conclude that Cp40 is a promising new treatment for complement-mediated AIHA.

Molecular minimal residual disease negativity and decreased stem cell mobilization potential predict
excellent outcome after autologous transplant in NPM1 mutant acute myeloid leukaemia (AML). Outcomes vary if intensively treated NPM1mut AML receive consolidation treatment with autologous SCT in first complete remission depending on the NPM1mut minimal residual (MRD) status and on the stem cell mobilization potential. These results may contribute to improve the selection of appropriate candidates for autologous SCT within the subgroup of NPM1mut AML patients and to identify those that possibly should rather undergo allogeneic SCT if possible.
De Santiago de Benito A et al. Haematologica 2019, in press.

Platelet protein S limits venous but not arterial thrombosis propensity by controlling coagulation in the thrombus
Platelet protein S (PSplt) controls platelet activation as well as coagulation in thrombosis in large veins but not in large arteries. PSplt limits thrombin generation in the thrombus and insures that highly activated platelets and fibrin remain localized at the injury site.
Calzavarini S et al., Blood 2019, in press.
Successful ESTHER project: Introduction of point-of-care HIV viral load measurement in Macenta, Forest Region, Guinea-Conakry, West Africa: «Go Viral»

Dr. Cornelia Staehelin and her coworkers in Guinea-Conakry have successfully implemented HIV viral load testing in Macenta within an ESTHER Project.

Machine learning methods for understanding outbreak dynamics

During a large outbreak of vancomycin-resistant enterococci (VRE) we gathered all relevant data generated during medical routine on more than 60,000 admissions for data science purposes. This permitted an accurate analysis of risk factors for being affected by VRE. Machine learning methods were then employed to tie a network of VRE transmissions among patients. After securing funding from the SNF, we will continue delineating the dynamics of this outbreak with novel analytical methods in order to be prepared for future outbreaks.

For further information contact the PI of this SNF grant, Andrew Atkinson, PhD, Andrew.atkinson@insel.ch.

HCC Incidence in HIV/HBV coinfection

Hepatocellular carcinoma (HCC) is a major cause of morbidity and mortality among HIV/HBV coinfected individuals. Among 3,625 study participants enrolled in a European multi-cohort study, 60 individuals (1.7%) developed an HCC. Age at initiation of HBV therapy and the presence of liver cirrhosis were predictors of HCC. The incidence of HCC remained below the commonly used screening threshold in patients without cirrhosis, who started on tenofovir disoproxil fumarate (TDF) when aged <46 years old. Wandeler, G et al, J Hepatol. 2019 Aug;71(2):274-280.
Department of Intensive Care Medicine

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Grants
- Study grant: The international sham-controlled randomized clinical trial PhINEST to treat dysphagia in ICU patients post mechanical ventilation (Phagenesis Ltd.)
- ESICM Basic Science Award: Assessment of inhalative Phage Therapy in Methicillin Resistant Staphylococcus Aureus (MRSA) Ventilator Associated Pneumonia (VAP) in Rats – AEROPHAGE
- Personalized Health and Related Technologies [PHRT-110] Grant: Personalized Swiss Sepsis Study (PSSS): Detection and modelling of sepsis using machine learning to analyse continuous ICU monitoring, laboratory, microbiology, and -omics data for personalized sepsis management

Profile
- Key objective: to advance the pathophysiological understanding of critical illness and associated (multiple) organ failure and to develop innovative strategies to fight the increasing threat of antibiotic resistance. Special emphasis is placed on alternatives to antibiotics such as phage therapy and anti-virulence approaches.
- Clinical trials (including international multicenter trials) with internal and external partners focusing organ failure, metabolism/immune responses and severe infections.
- New translational research group “Immunosuppression in Critical Illness” founded in 2019. “European Group on Immunology in Sepsis, EGIS” co-founded by KIM
- Experimental research projects include animal models particularly on the pathophysiology of circulatory, cerebral and gastrointestinal function.
- Experimental research (including animal models particularly on the pathophysiology of circulatory, cerebral and gastrointestinal function) is performed in cooperation with other groups, including ARTORG and Institute for Infectious Diseases, University of Bern.
- Full spectrum of training in Intensive Care Medicine, including training to qualify as an ICU specialist, training for medical students, advanced training for specialist nurses to qualify as certified experts in intensive nursing (INDS HF) and training for individuals to become professional healthcare assistants.
- Educational methods including simulator-based training, Echocardiography, Sonography and ECMO training
- External Partners: European Group on Immunology in Sepsis (EGIS): multiple research partners across Europe, Prof. M. Moller, Prof. A. Perner, Center for Research in Intensive Care (CRIC), Copenhagen, Denmark, Prof. K. Amrein, Medical University of Graz, Vienna, Austria, Prof. W. Doehrer and Prof. von Haehling, Cachexia Research Groups, Charité University Medicine Berlin, and Goettingen University, Germany, Prof C. Storm, Charité Center of Excellence for Cardiac Arrest, Berlin, Germany, Marzia De Lucia, PhD PD, Head of EEG imaging, Centre for Research in Neuroscience – Department of Clinical Neurosciences, CHUV – UNIL, Laurence G. Rahme, Prof. of Surgery, Microbiology and Immunobiology, Harvard Medical School, Director of the MGH Molecular Surgical Laboratory, Center for Surgery, Innovation and Bioengineering, Massachusetts General Hospital, Boston, MA, USA
Introduction of CAR-T treatment in lymphoma and ALL patients.

In January 2019, the first Swiss patient received CAR-T treatment at the University Hospital/Inselspital. Up to 20 patients with relapsed/refractory ALL or lymphoma are treated 2019 at the Inselspital. We developed an assay to monitor persistence of CAR-T cells in the follow-up of such patients.

First population-based data on incidence, mortality, and survival of soft tissue (STS) and bone sarcoma (BS) diagnosed in Switzerland.

We performed a retrospective study with data from the National Institute for Cancer Epidemiology and Registration (NICER) database in Switzerland. The age-standardized incidence and mortality rates in 2011-2015 were 4.43 and 1.42 per 100,000 person-years for STS, and 0.91 and 0.42 for BS. Incidence rates of STS and BS have been stable since 2001. The longer RS in STS can be attributed to advances in sarcoma patient management.
Department of Nephrology and Hypertension

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Profile

- Teaching programs for students in medicine, biomedical sciences (BMS) and membrane biochemistry as well as supervision of bachelor and master students for their elective modules, master thesis and PhD students for Cellular and Biomedical Science (CGB)
- 8 research groups
- Investigation of the mechanisms that contribute to renal function, loss upon hypoxyia and chronic allograft failure, formation of kidney stones, development of renal fibrosis and atherosclerosis, as well as factors influencing the steroid hormone metabolism and the role of the transportome in renal function
- Aim: improving diagnostic tools and implement of novel therapeutic approaches to enhance patient care

External partners:
- Department of Pharmacology and Toxicology, University of Lausanne, Switzerland
- Institute of Social Medicine, University of Cambridge, Cambridge, UK; Università Cattolica del sacro Cuore, Rome, Italy; University of Stellenbosch, South Africa; Baker Heart Research Institute, Melbourne, Australia; University of Texas Southwestern Medical Center, Division of Nephrology, Dallas, USA; Johns Hopkins University, Department of Physiology, Baltimore, USA

Grants

- Swiss National Foundation (310030_182482; 310030_188762/1; 314C0_1866785/1; 310030_182272; CR133_180326; CSRS-5_190714)
- NCCR Kidney.ch
- NCCR-Kidney.CH Junior Grant Award
- NCCR TransCure
- Swiss Transplant Cohort study
- Swiss Cancer League (KFS 3966-08)
- Foundation Johanna Dürmüller-Boll

Highlights

Role of membrane transport proteins in health and disease

Solute Carrier (SLC) transporters play important roles in the maintenance of biological barriers in different organs, as well as intracellular organelles. Using information from various publicly available databases and scientific literature, we have recently identified ~150 putative new SLCs from human, of which ~25 have known transport function and 40% are orphans. SLCs are still relatively unexploited as therapeutic targets, and their characterization can potentially open the way for novel personalized therapeutic applications. Gyimesi & Hediger, manuscript in preparation.

Role of the TWEAK/Fn14 pathway in experimental Calcineurin Inhibitor Toxicity (CIT)

Using clinical and gene-wide association data from the National Swiss Transplant Cohort Study, we demonstrated that a frequent single nucleotide polymorphism within TWEAK’s 3’UTR is predictive both for CIT lesions in the allograft and for premature allograft failure (A). We showed in vitro that the 3’UTR is critically involved in regulation of TWEAK expression potentially via post-transcriptional mRNA stabilization (B). We try currently to clarify the mechanisms of TWEAK’s post-transcriptional regulation and its consequences on inflammation and renal fibrosis. Silder et al. Manuscript in preparation.

Redox protein MEMO affects phosphotrophic bone hormone FGFR3 secretion and signaling

FGF receptor signaling in osteoblasts depends on the presence of the redox protein Mediator of Cell Motility 1 (MEMO). Ablation of MEMO in mice severely alters bone structure and energy metabolism. We investigate the interactions of MEMO with FGFR3 biology using redox proteomics, experimental kidney disease models, bone tissue-specific mouse models and primary culture techniques of bone and kidney cells to define fundamental biological mechanisms. This may ultimately help to define diagnostic or therapeutic targets for CKD-MBD, a frequent complication of chronic renal failure that carries high morbidity and mortality. Moor et al., JBMH Russ 2018;195-200.

Association of urinary sex steroids and glucocorticoids with muscle mass and strength in healthy adults

We examined the association of urinary excreted sex steroids and glucocorticoids with lean mass and grip strength in the SKPOGH cohort. Our results strongly support a direct link between sex steroid and glucocorticoid hormones in a physiological range with muscle mass and function. Steroid hormones seem to exert age-specific anabolic effects on lean mass and handgrip strength: Deficits in physical performance in the aged muscle may be attenuated by androgens, whereas glucocorticoids in a physiological range seem to increase skeletal muscle mass at all ages, and muscle strength in particular in younger adults. Both et al., J Clin Endocrinol Metab. 2019;104(6):2195-2215.

Association of 24-hour urine TH-cortisol excretion with handgrip strength after multivariable adjustments

USP of a patient with a relapse of an adrenocortical tumor producing corticosterones, glucocorticoids and sex steroids (blue dots) compared with age-matched reference intervals (Box plots). A surgical intervention can be planned after adrenal imaging.

Profile

- Teaching programs for students in medicine, biomedical sciences (BMS) and membrane biochemistry as well as supervision of bachelor and master students for their elective modules, master thesis and PhD students for Cellular and Biomedical Science (CGB)
- 8 research groups
- Investigation of the mechanisms that contribute to renal function, loss upon hypoxyia and chronic allograft failure, formation of kidney stones, development of renal fibrosis and atherosclerosis, as well as factors influencing the steroid hormone metabolism and the role of the transportome in renal function
- Aim: improving diagnostic tools and implement of novel therapeutic approaches to enhance patient care

External partners:
- Department of Pharmacology and Toxicology, University of Lausanne, Switzerland
- Institute of Social Medicine, University of Cambridge, Cambridge, UK; Università Cattolica del sacro Cuore, Rome, Italy; University of Stellenbosch, South Africa; Baker Heart Research Institute, Melbourne, Australia; University of Texas Southwestern Medical Center, Division of Nephrology, Dallas, USA; Johns Hopkins University, Department of Physiology, Baltimore, USA

Grants

- Swiss National Foundation (310030_182482; 310030_188762/1; 314C0_1866785/1; 310030_182272; CR133_180326; CSRS-5_190714)
- NCCR Kidney.ch
- NCCR-Kidney.CH Junior Grant Award
- NCCR TransCure
- Swiss Transplant Cohort study
- Swiss Cancer League (KFS 3966-08)
- Foundation Johanna Dürmüller-Boll

Highlights

Role of membrane transport proteins in health and disease

Solute Carrier (SLC) transporters play important roles in the maintenance of biological barriers in different organs, as well as intracellular organelles. Using information from various publicly available databases and scientific literature, we have recently identified ~150 putative new SLCs from human, of which ~25 have known transport function and 40% are orphans. SLCs are still relatively unexploited as therapeutic targets, and their characterization can potentially open the way for novel personalized therapeutic applications. Gyimesi & Hediger, manuscript in preparation.

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Association of 24-hour urine TH-cortisol excretion with handgrip strength after multivariable adjustments

USP of a patient with a relapse of an adrenocortical tumor producing corticosterones, glucocorticoids and sex steroids (blue dots) compared with age-matched reference intervals (Box plots). A surgical intervention can be planned after adrenal imaging.
Intra-arterial Urokinase as a promising treatment option after mechanical thrombectomy

Mechanical thrombectomy is the current standard of care in acute ischemic stroke patients presenting with a large vessel occlusion. In this work, it was shown that additional administration of intra-arterial Urokinase can be beneficial in patients with failed procedures, without increasing the risk of systemic or intracranial bleeding. Multicenter validation of the findings and planning of a randomized-controlled trial is currently ongoing. Kaesmacher et al., JAMA Neurology 2019, in press.

Highlights
**Deep Brain Stimulation (DBS)**

Our research has pioneered the development of segmented leads and is focused on the development of a DBS algorithm to predict the most effective stimulation area in the Subthalamic Nucleus (STN) based on a comprehensive probabilistic stimulation map (PSM) of the STN integrating structural and connectivity imaging data, as well as electrophysiological biomarkers. Using computer-based biophysical electric field models, it is possible to estimate the stimulated nerve tissue around the electrode which is referred to as the volume of tissue activated (VTA). Reducing and confining the VTA to specific parts of the STN obtained from the PSM could improve outcome or even reverse stimulation induced cognitive impairments in STN DBS patients.

Nguyen TAK et al., Brain Stimul. 2019.

**Paracrine factors: A powerful tool for brain repair**

In 2019 we investigated the potential of Endothelial progenitor cells-derived conditioned medium (EPC-CM) on the survival of cultured neuronal progenitor cells. Taken together, our findings identified EPC-CM as a powerful tool to promote survival of cultured neurons and further support the importance of paracrine factors in the actions of stem and progenitor cells for brain repair.

Di Santo S et al., Cell Transplantation 2019.

In 2019 we investigated the potential of Endothelial progenitor cell-derived conditioned medium (EPC-CM) on the survival of cultured neuronal progenitor cells. Taken together, our findings identified EPC-CM as a powerful tool to promote survival of cultured neurons and further support the importance of paracrine factors in the actions of stem and progenitor cells for brain repair.

**Trauma**

The New England Journal of Medicine published the findings of the randomized controlled, blinded TOSCAN trial. The TOSCAN trial evaluated the role of routine CT scanning after evacuation of chronic subdural hematoma, and showed that doing CT scans only in cases of neurological evaluation decreased the number of second surgeries, had a positive impact on morbidity and mortality, and lowered costs.


**Neurooncology**

Research of the neuro-oncological team is focused on intra- and perioperative mapping and monitoring of neurological function as well as intra-operative differentiation of tumor tissue, infiltration and peri-lesional healthy brain. The team has published a key paper, which showed how post-operative transcranial magnetic stimulation can predict neurological outcome after surgery. The HORAO project reached its first milestone as the winners of the crowdsourcing competition were elected during the final conference. Preliminary studies are evaluation an array of technologies for improved tumor boarder identification, paving the way for larger trials starting 2020.

Deep neural network automatically characterizes lesions on Ga-68-PMSA-PET/CT

We developed a deep neural network to characterize automatically the prostate cancer lesions on Ga-68-PSMA-PET/CT. The preliminary test within the pelvic area confirms the potential of deep learning methods.


Digital PET/CT detects more lesions compared to analogue PET/CT

Digital PET/CT detected more prostate cancer lesions compared to an analogue PET/CT. A significantly higher rate of pathological PET/CTs was found in the patient group with the lowest PSA levels. The differences could be plausibly explained by the measured imaging characteristics of the scanner.


Anle138b treatment ameliorates tau pathology in a transgenic mouse model

Late-stage oligomer modulation effectively ameliorated tau pathology in hTau mice and consequently rescued metabolic function. Molecular imaging by FDG-PET could serve for monitoring effects of such treatment.


Dopaminergic loss and its metabolic correlates in dementia with Lewy bodies

Relative glucose hypermetabolism and disturbed metabolic connectivity of limbic and basal ganglia circuits correlated with the extent of nigrostriatal degeneration in DLB patients. The identification of specific metabolic network alterations in patients with early nigrostriatal degeneration may serve as an additional supporting biomarker for early diagnosis of DLB.

M. Huber et al. Mov Dis (2019) accepted.

Immunohistochemistry of frontal cortex, treatment (A) vs. vehicle (B)

Cerebral FDG-PET baseline and after 3 mo treatment vs. placebo

Architecture of the proposed framework

Grants

- Swiss National Science Foundation (grant no. 200021_188914; IZKSW3_188350)
- Swiss Cancer League (grant no. KFS-4723-02-2019)
- FreeNovation Fund, Novartis Foundation
- UlmInn Forschungsförderung (grant no. 43/2018)
- Deutsche Forschungsgemeinschaft (grant no. RO5194/1-1)
- Berger-Janser Stiftung (grant no. 11/2019)
Department of Obstetrics and Gynecology
Inselspital, Friedbühlstrasse 19, 3010 Bern

Division of Obstetrics and Feto-Maternal Medicine

Areas of Scientific Investigation:
- Stem cells, exosomes, non-coding RNA and sPIF in peripartum Neuroregeneration
- Basic function of Astroglia in peripartum hypoxia and inflammation
- Transmembrane transporter and biomarker in preeclampsia and preterm birth
- Clinical research in preterm birth, fetal neurosonography, labor induction, postpartum hemorrhage
- Teaching: preclinical and clinical at master, MD and PhD level; postgraduate preclinical / clinical training
- External Research partners: Dept. Pediatrics and Neonatology, Maastricht University; Dept. Neurosurgery, Cellular and Molecular Physiology, Yale University; Dept. Obstetrics, Heching Union Medical College Hospital (PUMCH); Dept. Obstetrics, Basel University; Dept. Obstetrics, Schiller University Jena; Dept. Obstetrics, University of Insurbia, Varese; Risch Laboratory, Bern; Bionercept, LLC, Cherry Hill, New Jersey

Grants
- SNF Grant: “Salt Supplementation in pregnancy at high risk to develop preeclampsia”
- EU Grant COST: “International network for translating research on perinatal derivatives”
- Bangerter Stiftung: “Mesenchymal stem cells-derived exosomes as a neuroregenerative therapy”
- CTU Grant: “Preimplantation factor: Biomarker for preterm birth?”
- Grant Swiss Society of Obstetrics and Gynaecology / Bayer
- Investigator-initiated trial (Ferring): “Patient blood management in obstetrics”
- Investigator-initiated trial (Ferring): “Induction of labor: misoprostol vaginal insert vs tablets”
- Perinatal medicine intramural fund: Several grants for clinical studies

Highlights

Stem cells in peripartum brain damage: Translation

Researcher Group

Division of Gynecology and Gynecological Oncology

Head Research
Prof. Daniel Surbek
Director and Co-Head Research Laboratory Perinatal Medicine

Group Leaders
Prof. Luigi Riso Deputy Director Group Leader
PD Dr. Beatrice Mosimann Group Leader
PD Dr. Marc Baumann Group Leader
PD Dr. Andreina Schoenberkin Co-Head Research Laboratory Perinatal Medicine Group Leader
PD Dr. Martin Müller Group Leader

Profile
- Pathophysiology of endometriosis and associated pain
- Pathophysiology of gynaecologic cancers
- Quality of life before and after urogynaecologic interventions
- Physiologic and pathophysiologic changes of the pelvic floor during exercises
- Transgender studies
- External partners: Center for Gender Variance, Univ. of Basel; Berner Fachhochschule, Bern; Dell Medical School, Univ. of Texas, Austin; Croydon Univ. Hospital, London; Dept. of Obstetrics & Gynaecology Inst. for Molecular Biosciences, Queensland; Clinical Pathology & Cytology, Karolinska Univ. Hospital, Stockholm; Dept. of Urogynaecology, Princess-N-Hospital, Southampton; Dept. of Obstetrics & Gynecology, Ospedale Santa Chiara di, Trento

Grants
- Swiss National Science Foundation
- Contura AG, Copenhagen, Denmark
- Bern Center for Precision Medicine (BCPM)
- Innosuisse – Swiss Innovation Agency
- Stiftung für Klinisch-experimentelle Tumorforshung

Highlights

Progestosterone resistance in endometriosis
Dienogest is one of the promising options for the long-term hormonal management of endometriosis. By analyzing the prevalence of somatic mutations, cellular composition (progestin receptor level in stromal cells, Fig. A), or transcriptomic profile between endometriotic lesions from non-responders (GO, Fig B), we intend to unveil the mechanism of Dienogest resistance which occurs in several patients.

Endometrial cancer (EmCa) with sentinel lymph-node (SLN) dissection
SLN removal offers a convincing balance between oncological safety and perioperative morbidity in early-stage EmCa. However, lymphovascular space invasion may represent a risk factors for recurrence and therefore lymph-node evaluation is crucial. Immoden S. et al. Eur J Surg Oncol. 2019.
Retinal artery occlusion is associated with compositional and functional shifts in the gut microbiome and altered trimethylamine-N-oxide levels

Retinal artery occlusion (RAO) is a sight-threatening complication of cardiovascular disease and commonly occurs due to underlying atherosclerosis. As cardiovascular disease and atherosclerosis in particular has been associated with compositional alterations in the gut microbiome, we investigated this association in patients with clinically confirmed non-arteritic RAO compared to age- and sex-matched controls. On the phylum level, the relative abundance of Bacteroides was decreased in patients with RAO compared to controls, whereas the opposite was observed for the phylum of Proteobacteria. Several genera and species such as Actinobacteria, Bifidobacterium spp., Bacteroides stoolcoproducers, Faecalibacterium prausnitzii were relatively enriched in patients with RAO, whereas others such as Odontobacter, Parasutterella or Lachnospiraceae were significantly lower. Patients' gut microbiomes were enriched in genes of the cholesterol metabolism pathway.

The gut derived, pro-atherogenic metabolite trimethylamine-N-oxide (TMAO) was significantly higher in patients with RAO compared to controls (p = 0.023) and a negative correlation between relative abundances of genera Parasutterella and Lachnospiraceae and TMAO levels was found in study subjects. Our findings propose that RAO is associated with alterations in the gut microbiome and with elevated TMAO levels, suggesting that RAO could be targeted by microbiome-altering interventions. Gut enterotypes in the cohort. Three enterotypes exist in the cohort based on the abundance of microbial genera. Patients (n = 29) and controls (n = 30) are denoted by squares (©) and triangles (Δ), respectively. Boxplots represent the abundance of Bacteroides, Eubacterium and Prevotella, the proposed drivers of the three enterotypes (b). Blue is enterotype 1, purple is enterotype 2 and orange is enterotype 3. Scientific reports (2019) 9:15303.
Department of Orthopedic Surgery and Traumatology

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Prof. Klaus A. Siebenrock
Chairman

Prof. Loth M. Bemmerker
Head Spine Surgery

Dr. Michael Schär
Head Shoulder, Elbow and Sports

PD Dr. Frank M. Kienle
Head Knee Surgery and Tumor Surgery

PD Dr. Fabian Krause
Head Foot and Ankle Surgery

PD Dr. Johannes D. Bastian
Head Orthoeriatrics

PD Dr. Sven Höpple
Head Translational Medicine and Spine Research

Dr. Monic Danti
Pediatric Spine Surgery

Prof. Matthias Zumstein
Head Shoulder, Elbow and Sports

Prof. Markus Keel
Consultant Hip and Spine Surgery

Profile

- Hip research focus on the development of osteoarthritis and pre-artritic deformities like femoroacetabular impingement
- Spine research is focused on disc degeneration models and tissue-engineered disc regeneration and the treatment of osteoporotic fractures
- The shoulder & elbow team is working on statistical shape modeling of shoulder morphology, rotator cuff regeneration incl. stem cells and the investigation of surgical techniques and implants
- The main focus of knee research is the regenerative treatment of the anterior cruciate ligament; prosthetic infections and anti-infectious implants are also research topics
- Arthritis of the ankle joint, innovative treatment of ankle fractures and AMIC plastic in osteochondral lesions is the main focus in foot & ankle research
- Orthogeriatrics investigates the impact of orthogeriatric pathways and rehabilitations protocols on the clinical outcome in geriatric patients
- Translational medicine is the latest research field which will come to focus in the next years
- External Partners: Musculoskeletal Research Unit, Vetsuisse Faculty, University of Zürich, Zürich, Switzerland; AO Research Institute, Davos; RMS Foundation, Bettlach; SUVA

Grants

- Zumstein M, Dommer L: Timepoint and direction of secondary dislocation in early conservative treatment in distal radius fractures using 3D imaging. Hans Neuenchander Fonds, 2019
- Gantenbein B, Benneker L: «Working Package 1 of the H2020 Project »IPS spine« Project Agreement #825925 - Induced pluripotent stem cell-based therapy for spinal regeneration»
- Siebenrock KA, Beckmann NA, Fellowship Hip Surgery, Hans Neuenchander Fonds & Depuy Synthes, 2019

Second Swiss Orthogeriatrics Day, 31.10.2019

The number of elderly patients in orthopaedics and traumatology is steadily increasing and represents a new challenge in interdisciplinary and interprofessional treatment. Accordingly, in 2018 the «1st Swiss Orthogeriatrics Day Bern» was initiated by PD Dr. med. J. D. Bastian and hosted together with Prof. Dr. med. A. E. Stuck to provide a platform for all occupational groups involved in the treatment of the elderly for a rich and informative interdisciplinary exchange. On the 31 October in 2019, we were able to build on our previous success with the «2nd Swiss Orthogeriatrics Day» and hosted that symposium with PD Dr. med. J. D. Bastian together with Prof. Dr. med. A. Schönberger. The program included the following sessions:

Session 1:
- «Key Note Session» Gerontotraumatology from an emergency medicine point of view
- «Parallel Workshop-Sessions of different disciplines» with «Hands-on» about «The hands in old age - The key to independence» (Ergotherapy), «Malnutrition in orthogeriatrics» (Nutrition Counselling), «Geriatric assessment» (Geriatrics), «Co-management in the Rhine Valley» (Geriatrics), «Guided Tour around the Emergency Department, Inselspital» (Emergency Medicine), «Sonography in Gerontotraumatology» (Emergency Medicine), «Challenges in an acute-care hospital» (Nursing Staff), «The orthogeriatric patient» (Physiotherapy), «Patient discharges in succession scenarios» (Social Services), «Osteoporotic vertebral fractures» (Orthopaedics), «ADAPT system for Gamma3 nail» (Orthopaedics), «Tricky Trauma Cases – think outside the box!» (Orthopaedics), «TFN-A on the sawbone» (Orthopaedics)

Session 2:
- Unstable joint fractures - how does post-treatment work? As a debate session with specialist contributions with «Pros» and «Cons» for one or another treatment.

About 170 participants from 16 different cantons in Switzerland joined this event and made it a great success. For 2020, the 3rd Orthogeriatrics Day is already under preparation.

Sarcoma Center Bern

Sarcomas are a group of rare cancers originating from bone and the soft tissues. The Sarcoma Center Bern is one out of five university centers specialized on the care of sarcoma patients in Switzerland. Currently, the Sarcoma Center Bern is the only Swiss center fully certified according to ISO 9001:2015 and the German Cancer Society (DKG). The center, led by the Department of Orthopaedic Surgery, belongs to the University Cancer Center Inselspital (UCI). At the Sarcoma Center Bern 17 medical disciplines make use of state of the art diagnostic and treatment modalities and provide personalized care in order to ensure the best treatment for sarcoma patients.

A recent multicenter population-based retrospective registry study performed together with the National Institute for Cancer Epidemiology and Registration (NICER) was the first study analyzing population-based data on the temporal trends of incidence, mortality and survival of soft tissue and bone sarcomas diagnosed in Switzerland. The study showed a stable, low incidence rate of approximately five soft tissue sarcomas (STS) and one bona sarcoma (BS) per 100,000 person years, respectively. This related to an overall annual incidence of 366-448 STS and 63-77 BS in Switzerland between 1996 and 2015.
Department of Osteoporosis

Inselspital, Imhofs-Pavillon, 3010 Bern
www.osteoporose.insel.ch

Profile

- Epidemiological Research: focused on the epidemiology of fractures and the socioeconomic burden of osteoporosis and fractures; continued development and update of the country-specific fracture risk prediction tool (FRAX)
- Translational research: development of bone strength prediction and clinical fracture risk assessment by finite element analysis using high resolution peripheral quantitative CT (HR-pQCT) at the forearm and the tibia
- Clinical research (Phase II-IV): development of new investigational drugs for the treatment of osteoporosis and the reduction of fracture risk
- Experimental first in man clinical research: investigation of local bone remodeling and mechanoregulation of bone fracture healing in healthy, aged, and osteoporotic humans; targeted stimulation of bone anabolism with in situ bone active agents; osseointegration of bone implants
- Osteoporosis lectures for medical students in the context of geriatrics (year 4); practical courses on pharmaceutical medicine for students of human medicine (year 2); module “bone measurement technologies” of the “Biomechanics Labs” for students of biomedical engineering
- External research partners: Centre for Metabolic Bone Diseases, University of Sheffield, UK; ETH Zurich, Institute for Biomechanics, Zurich; Free University of Amsterdam (ACTA), Amsterdam, the Netherlands; Swiss Paraplegic Center, Rehabilitation and Geriatrics, Geneva; Zhejiang Chinese Medical University, Hangzhou, China

Grants

- MICROMESO SNF project grant no 165510 until 2020 (Co-PI)
- PARATY Fondation pour l’Ostéoporose until 2020 (PI)
- Osteoporosis lectures for medical students in the context of geriatrics (year 4), practical courses on pharmaceutical medicine for students of human medicine (year 2); module “bone measurement technologies” of the “Biomechanics Labs” for students of biomedical engineering
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Highlights

BMP-7 for Implant Osseointegration in Osteoporosis

Osseointegration of dental and orthopaedic implants is seriously impaired in an osteoporotic environment. By using bone morpho-
genetic protein-7 (BMP-7) we are able to significantly improve implant healing when used at a low dose in a delayed drug-release system, and were also able to prevent drug-related side-effects (ectopic bone formation, inflammation, cyst formation etc.). Hunziker et al.: The Slow Release of BMP-7 at a Low Dose Accelerates Implant Healing in an Osteoporotic Bony Environment. Ready for submission, 2021

SNSF DACH Fracture study (DACH-Fx)

In collaboration with ETH Zurich, University Hospital Innsbruck, Ulm University, and Insel Orthopaedics, this study follows distal radius fracture patients over one year. Detailed clinical assessment and HR-pQCT imaging are performed at six time points. This study will characterize fracture healing at the level of single trabeculae as well as the outcome contribution of age, osteoporosis, and bone biomarkers. In addition, computational models are built to study local mechanical regulation and to simulate the healing process.

HR-pQCT-Based Estimation of Bone Strength

High-resolution peripheral computed tomography (HR-pQCT) allows to reconstruct the in vivo geometry of distal fracture sites, including at the radius and the tibia, with a resolution of 61 microns. A novel measurement protocol with multiple bone sections and a homogenised finite element methodology was developed for evaluation of bone strength at the distal radius and tibia. An extensive in vivo reproducibility study was conducted to determine the precision of this methodology. Finally, reference values and clinical predictors of bone strength for HR-pQCT-based distal radius and tibia strength were established in young Swiss women and men.

Periodontitis and Periimplantitis in Osteoporosis

Periodontitis (POIS) and periimplantitis (PIIS) are very frequent diseases in dentistry and patients with osteoporosis. We investigated the differences between the two diseases in generally used experimental in vivo-models. Bone resorption was found to be significantly higher in the PIIS model. Huang et al.: What are the differences in the inflammatory process between experimental periimplantitis and periodontitis? Submitted for publication, 2019.

Top panel: in situ photograph of a ligature-induced paradontitis (tooth) and an implant with periimplantitis. Bottom panel: same set-up as above, imaged by CBCT.

Osteointegrated experimental dental implant in the osteoporotic bone

Failure zone of the distal radius calculated by homoge-
nised finite element analysis

2D radiograph of a right wrist showing a fracture of the radius (left). Cut away of the radius with region of fracture healing after 1 and 3 weeks, respectively. Image courtesy of Penny Atkins.
Department of Pediatrics

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Prof. Steffen Berger
Director a.i.

Prof. Christoph Aebi
Extraordinary and Head Paediatric Infectiology

Prof. Christa Plück
Extraordinary and Head Paediatric Endocrinology, Diabetology and Metabolism

Prof. Philipp Latzin
Extraordinary and Head Paediatric Pneumology and Allergology

Prof. Johann Schaller
Head a.i. Haematology and Oncology

PD Dr. André Rössler
Extraordinary and Head Paediatric Neurology and Neurodevelopmental Unit

Prof. Maja Steinlin
Extraordinary and Head Paediatric Diabetology and Endocrinology, and Allergology

Profile

• Teaching at every level of the School of Medicine and other courses at the University of Bern
• Providing training in Paediatrics and all its subspecialties as well as in Medical Genetics (FMH, FAMH)
• Offering PhD training in programs of the Graduate Schools of the University of Bern
• Very broad research spectrum ranging from basic to translational to clinical projects
• Genetics: Projects to identify novel disease-associated genes in rare diseases. Partners: University Hospitals Switzerland Haematology/Oncology: Strategies to manage fever in neutropenia; trying to understand the situation of childhood cancer survivors (in collaboration). Studies on mechanisms of genomic instability hematological malignancies (in cooperation with the DKFZ); Preclinical research focusing on new drugs and delivery for pediatric sarcoma
• Infectiology: Multidimensional research of pediatric sepsis as host organization of the Swiss Pediatric Sepsis Study; Partners: Children’s Hospital, University of Queensland, Australia; University Children’s Hospital’s Zurich, Department of Pediatrics, University of Lausanne, CHUV, Lausanne
• Neuropaediatrics: SNPSR and CP registry studies in collaborations with other Swiss Children’s Hospitals; Brainfit-Study in collaboration with the University Children’s Hospital’s Zurich and Neuroimaging Center Salzburg, Austria; HERO-Study (reorganization after pediatric stroke, collaborations); Onset-Study (influence of age at acute brain problem collaboration)
• Pneumology: Research topics: lung development in health and disease, novel lung function tests and state-of-the-art (reorganisation after paediatric stroke, collaboration); Onset-Study (influence of age at acute brain problem collaboration)
• Neurology and Neurodevelopmental Unit: Functional resting state MR is increasingly important for research of brain development and effects of brain lesions as stroke, brain tumors trauma during childhood. We analyzed different networks in healthy children and also in children after stroke or brain tumors. We could show, that especially cognitive networks do develop after the early childhood with increasing connections not only in cortical networks but especially in thalamocortical networks. In children after stroke or brain tumors network alterations correlate with cognitive and motor outcome. Steiner et al., submitted.
• Genetics: Projects to identify novel disease-associated genes in rare diseases. Partners: University Hospitals Switzerland, University of Bern
• Very broad research spectrum ranging from basic to translational to clinical projects
• Offering PhD training in programs of the Graduate Schools of the University of Bern
• Providing training in Paediatrics and all its subspecialities as well as in Medical Genetics (FMH, FAMH)
• Teaching at every level of the School of Medicine and other courses at the University of Bern

Grants

• HORIZON 2020 - PERFORM, Partner (2017-2020), - DIAMONDS, Partner (2020-2024), Infectiology
• SNF Fr. 182719 (Latzin), 179905 (Yammine), 168173 (Ramsey), 182871 (Frey), Pneumology
• Uniscentia, Novartis and IFCAH-ESFE grants; Endocrinology
• Krebsforschung Schweiz-4709-02-2019 (Leibundgut), Oncology

Highlights

Curcuminoids as potential drugs against breast and prostate cancer

Curcumin, the major component of commonly used spice turmeric, can inhibit activities of steroid metabolizing enzymes CYP17A1, a target of drugs for prostate cancer, and CYP19A1, a target for drugs against breast cancer. Our results propose that curcuminoids may be modified to generate better lead compounds for potential drugs against prostate and breast cancer. Rodríguez-Castaño P et al., Int J Mol Sci. 2019 20(18).

Curcumin docked into active site of steroid metabolizing enzyme CYP17A1.

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Curcumin docked into active site of steroid metabolizing enzyme CYP17A1.
Department of Pediatric Surgery

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www.kinderkliniken.insel.ch/de/kinderkliniken/kiki-chirurgie

Grants

- Batzebär grants PD Dr. Fauser-Kan, PD Dr. Liebs
- AO Foundation grants PD Dr. Ziebarth, Dr. Slongo

Profile

- Participates in University teaching programs for students of medicine (PBL-teaching, bedside teaching, practical year, clinical skills training, surgical rotation for students)
- Oncobiology research (laboratory research in childhood malignancies); study of the expression of Tim-3 and Gal-9 molecules and their roles in leukemia and in pediatric and adult solid tumors. PD Dr. E. Fasler-Kan
- Research of Necrotizing enterocolitis of the newborn (Hosting of the Swiss national NEC registry): Prof. S. Berger, PD Dr. U. Kessler
- Research of Pathophysiology of necrotizing enterocolitis and appendicitis: PD Dr. U. Kessler
- Health related quality of life research after pediatric extremity fractures (clinical research and follow up studies): PD Dr. T. Liebs, Dr. N. Kaiser
- Development of new operative strategies in pediatric hip surgery and clinical outcome research after surgical treatment of pediatric hip and knee diseases, including slipped capital femoral epiphysis and ruptured anterior cruciate ligaments. PD Dr. K. Ziebarth

Highlights

The Tim-3-Galectin-9 Pathway and Its Regulatory Mechanisms in Human Breast Cancer

Human cancer cells operate a variety of effective molecular and signaling mechanisms which allow them to escape host immune surveillance and thus progress the disease. We have recently reported that the immune receptor Tim-3 and its natural ligand galectin-9 are involved in the immune escape of human acute myeloid leukemia (AML) cells. We now found that studied breast tumors expressed significantly higher levels of both galectin-9 and Tim-3 compared to healthy breast tissues of the same patients and that these proteins were co-localized. Surface-based galectin-9 was able to protect breast carcinoma cells against cytokine T cell-induced death. Furthermore, we found that cell lines from brain, colorectal, kidney, blood/mast cell, liver, prostate, lung, and skin cancers expressed detectable amounts of both Tim-3 and galectin-9 proteins. We conclude that the Tim-3-galectin-9 pathway is operated by a wide range of human cancer cells and is possibly involved in prevention of anti-tumor immunity. Yasinska et al., Front. Immunol. 2019; 10:1594.

Definition of pediatric hypotension: Gaps between two percentile-based definitions and Pediatric Advanced Life Support/Advanced Trauma Life Support guidelines

Data are lacking to provide cutoffs for hypotension defining shock in children based on outcome studies. Pediatric Advanced Life Support (PALS), and Advanced Trauma Life Support (ATLS) definitions are based on normal populations. We compared different normal population based cutoffs including fifth percentile of systolic blood pressure (PS-SP) in children and adolescents from the German Health Examination Survey for Children and Adolescents (KIGGS, 12,199 children aged 3-17 years), US population data (Fourth Report), and cutoffs from PALS and ATLS guidelines. The KIGGS PS-SP was on average 7 mm Hg higher than Fourth Report PS-SP (5-10 mm Hg depending on age-sex group). For children aged 3 to 9 years, KIGGS PS-SP at median height follows the formula 82 mm Hg + age; for age 10 to 17 years, the increase was not linear. Current PALS/ATLS guidelines largely underestimate the rate of hypotension in children with trauma. Sarganas et al., J Trauma Acute Care Surg. 2019;86: 448-453.

Long term results of modified Dunn procedure in SCFE

AIMS: The modified Dunn procedure has the potential to restore the anatomy in hips with severe slipped capital femoral epiphyses (SCFE). However, there is a risk of developing avascular necrosis of the femoral head (AVN). In a retrospective analysis involving 46 hips in 46 patients treated with a modified Dunn procedure for severe SCFE (slip angle > 60°) between 1999 and 2016. At nine-year-follow-up, 40 hips were available for clinical and radiological examination. Mean preoperative age was 13 years, and 14 hips (30%) presented with unstable slips. Mean preoperative slip angle was 64°. The modified Dunn procedure for severe SCFE resulted in a low rate of AVN, low risk of progression to OA, and high functional scores at long-term follow-up. The slip deformities were mainly corrected but secondary impingement deformities can develop in some hips and may require further surgical treatment. Lerch et al., Bone Joint J 2019;101-B:403-414.

Breast cancer cell based pathobiological pathways showing LPHN-induced activation of PKCα, which triggers the translocation of Tim3 and galectin-9 onto the cell surface which is required for immune escape.

Fifth PS-SP percentiles curves for boys (upper panel) and girls (lower panel) aged 3 to 17 years according to the KIGGS population (red), the Fourth Report references (blue), thresholds of the PALS and the ATLS (green).
We investigated whether intradermally administered indocyanine green (ICG) and near infrared fluorescence imaging (NIRFI), sentinel lymph nodes (SLNs) in malignant melanoma and Merkel cell carcinoma can be identified transcutaneously as well as by traditional lymphoscintigraphy (LS).


In this study, non-vascularized partial joint transfer provides a mobile and stable PIP joint up to 4 years after reconstruction. The surgical technique presented herein is complex depending on additional soft tissue, tendon and nerve injuries but results in great patient satisfaction.


Non-vascularized partial joint transfer for Finger Proximal Interphalangeal joint reconstruction: a series of 9 patients

In this study, non-vascularized partial joint transfer provides a mobile and stable PIP joint up to 4 years after reconstruction. The surgical technique presented herein is complex depending on additional soft tissue, tendon and nerve injuries but results in great patient satisfaction.


Osteosynthesis of fifth metacarpal neck fractures with a photodynamic polymer bone stabilization system

Figure 1: (a) Polyethylene terephthalate balloon filled with monomer ex situ. (b) After implantation the resin is cured in situ by application of visible light.

Figure 2: Fifth metacarpal neck fracture in posteroanterior and 45° oblique radiographs (a) preoperatively and (b) after fracture reduction and osteosynthesis with an intramedullary photodynamic polymer. (c) Radiological follow-up 3 months after operation. Journal of hand surgery. European volume, p. 1753193419880775. Sage 10.1177/1753193419880775.

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Multi-walled carbon nanotubes (MWCNTs) as a hazard in chronic obstructive pulmonary disease (COPD)

We have shown in a mouse model of COPD that MWCNTs instilled intratracheal in a realistic occupational dosage induce mucus hypersecretion, and phenotypic shifts and activation in macrophage and dendritic cell populations in the lung. These findings demonstrate enhanced susceptibility for adverse health effects induced by MWCNT in individuals suffering from COPD (PD Dr. Fabian Blank).

**Highlights**

**Honors**

Prof. Thomas Geiser was appointed as individual member of the Swiss Academy of Medical Sciences (SAMW).

**Awards**

Kristina Krempaska (Supervision: PD Dr. Manuela Funke-Chambour) received a Poster Award by the Swiss Society of Pneumology for the project “Anti-fibrotic and apoptotic effects of Azithromycin in IFP and control fibroblasts”. Moreover, she was awarded with the best preclinical project at the DBMR research day: “Azithromycin has enhanced anti-fibrotic properties on IFP fibroblasts” (Supervision: PD Dr. Manuela Funke-Chambour). Dr. Sabina Guler was awarded by the Swiss Society of Pneumology by the Poster Discussion award.

**Impaired Wound Healing of Alveolar Lung Epithelial Cells in a Breathing Lung-On-A-Chip**

In collaboration with ARTORG Lung, we developed a microfluidic in vitro system to study lung epithelial wound repair on a lung-on-chip system that includes breathing movements. The data support the development of novel therapeutic strategies in chronic lung diseases such as lung fibrosis or emphysema (Prof. Th. Geiser/Prof. O. Guenat).

**Grants**

- Swiss National Science Foundation - SINERGIA SNF grant nr. CRSII3_160704/1 (Prof. Thomas Geiser)
- Lungenliga Schweiz - Patient specific pluripotent stem cell derived organoids - a step to precision medicine, 2019 (Prof. Thomas Geiser)
- OPO-Stiftung Zürich and Lungenliga Bern: Efficient Pulmonary Mucosal Allergen Immunotherapy in allergic Airways Disease Employing Virosomes as Nanocarriers for Antigen and Adjuvant (PD F. Blank)
- Lungenliga Schweiz/Lindenhoft-Stiftung: miRNA in pulmonary hypertension (PD J. Richter Hefit)
- CTU-Forschungs-Grant: Clinical impact of dehydroepiandrosterone sulfate on disease severity & physical performance in fibrotic interstitial lung disease (Dr. Sabina Guler)
- Lungenliga Schweiz 2015 (PD Dr. Manuela Funke-Chambour)
- Lungenliga Bern 2019 (PD Dr. Manuela Funke-Chambour)
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Dr. Evelyn Hermann
Dr. Olguin Bolín
Dr. Mohamed Shalan
Prof. Stefan Eychmüller
PD. Dr. Peter Manser

Prof. Michael K. Fix
PD. Dr. Michaela Medova

Profile

- Teaching: Undergraduate lectures are provided for students in medicine, physics, biomedical engineering, molecular and biomedical sciences as well as in dental medicine at the University of Bern; undergraduate teaching does also include practical training for medical students; postgraduate lectures in medical physics are given at the Department of Physics at the Swiss Federal Institute of Technology in Zurich; a CAS program for palliative care is run in cooperation with various faculties of the University of Bern and the Bern University of Applied Sciences; various PhD and MD-PhD positions are provided in radiation biology and medical physics
- Clinical Research: (a) Prostate cancer: salvage radiotherapy in biochemical failure, metabolic signatures; (b) head neck cancer: role of upfront neck dissection in the context of primary radiotherapy; single vocal cord irradiation, outcome of immune checkpoint inhibitor and chemotherapy followed by dose and volume adapted chemo-radiotherapy; (c) brain tumor: Automatic segmentation of brain tumors, postoperative radiosurgery; (d) palliative care: Advanced care planning/anticipation; best care for the dying; community palliative care
- Medical physics research: (a) Dynamic Trajectory Radiotherapy (DTRT) and Modulated Electron Beam Radiotherapy (MERT); (b) Independent Dose Calculation and Radiation Protection in the Context of Robotic Stereotactic Radiotherapy; (c) Efficient Quality Assurance for External Beam Radiotherapy and Accurate Dose Calculation for Brachytherapy; (d) Medical Imaging Related Research Topics
- Radiation Biology: (a) Omics-related (transcriptomics, (phospho-)proteomics and single-cell proteomics, metabolomics) studies on receptor tyrosine kinase (MET) signaling, radiation therapy response and resistance and head and neck squamous cell carcinoma (HNSSC); (b) Roles of novel posttranslational modifications (phosphorylations, ubiquitinations, acetylations) in cellular responses to DNA-damaging agents and targeted therapies; (c) Translational research to explore new prognostic and predictive biomarkers
- Collaborations: Multidisciplinary and multinational national and international collaboration within the SAKK & EORTC networks; Institute for Biomedical Engineering, Swiss Federal Institute of Technology (ETH), Zurich, Switzerland; Princess Margaret Cancer Center, Toronto, Canada; Laboratory of Artificial Intelligence in Medicine and Biomedical Physics, Stanford Medicine, USA; ACRF Image X Institute, University of Sydney, Australia

Grants

- Swiss National Science Foundation
- Innosuisse
- Rising Tide Foundation
- Research Framework Agreement, Varian Medical Systems Inc.
- Swiss Bridge Foundation
- sitem-insel Support Funds
- Werner und Heddy Berger-Janser Stiftung
- Krebsforschung Schweiz

Highlights

Bypassing the imputation of missing values in differential expression analysis of proteomic data

Imputation of missing values in proteomic datasets significantly affects results of differential expression analyses. We developed ProtRank as a new method that identifies differentially expressed proteins by ranking their observed changes with respect to changes of other proteins and thus overcomes the need to impute missing values.


The impact of imputation on differential expression analysis

Disease Control With Delayed Salvage Radiotherapy for Macroscopic Local Recurrence Following Radical Prostatectomy

For patients with isolated local recurrence after radical prostatectomy, dose-escalated salvage radiotherapy to the prostate bed provides inferior tumor control compared to early intervention. Despite dose escalation, no increased incidence of high grade toxicities was observed.


Adaptive step size algorithm to increase efficiency of proton macro Monte Carlo dose calculation

Accurate and efficient dose calculation is essential for treatment planning purposes in radiation therapy and Monte Carlo (MC) simulations are commonly known as the most accurate method to perform this task. Our group works on this topic and developed an adaptive macro Monte Carlo method for efficient dose calculations of proton beams.

Kueng R et al., Radiation Oncology, 14:165, 2019.

Metabolomics reveals tepotinib-related mitochondrial dysfunction in MET-activating mutations-driven models

Inhibition of the MET oncogene by small molecule inhibitor tepotinib alters metabolic signature of cancer cells and affects their mitochondrial function. Metabolic state of cancer cells upon MET inhibition can serve as an early indicator of efficacy of anti-MET therapies.


Changes of mitochondria-related metabolites upon MET inhibition.
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Profile

- Participating in University Teaching programs for students of medicine, biomedicine and biology; master students of biology are also coming from foreign Universities
- 3 Basic research groups and one clinically oriented research group
- Aim: Understanding of immune regulation and the development of potential new therapies for inflammatory and other diseases
- Rheumatology, Clinical Immunology and Allergology offered optional teaching courses for last-year medical students, in addition to regular courses and lectures; these interactive workshops were again very popular
- Clinical research focuses large-vessel vasculitides, immune regulation in pregnancy, pathogenesis of arthritides and osteo-immunological interaction; in Allergology drug-hypersensitivity and immune-deficiencies are in the research focus
- Collaboration with a variety of partners within the medical faculty of Bern, with partners in Germany, France, the USA and Italy are ongoing
- External Partners Immunology: The Jenner Institute, University of Oxford, Oxford; Department of Dermatology, Zürich and Italy are ongoing
- Grants
  - Swiss National Science Foundation (SNF), Bern. Project: “Unraveling the role of CD23 in the context of allergy”. Project period: 1.04.2018-31.03.2022
  - Swiss National Science Foundation (SNF), Bern. Project: “Understanding the role of antibody affinities in the regulation of allergic responses”. Project period: 1.04.2019-31.03.2023
  - National Institute of Health R01 Subaward with Prof. Theodore Jardetzky from Stanford University, Project: “Virus-like particle based vaccine to treat Fel d 1-induced allergy in human subjects by immunizing cats against their own major allergen, Fel d 1. The vaccine successfully induces Fel d 1 binding antibodies in cat and these antibodies neutralize the allergens thereby reducing symptoms of allergy in owners. Thoms F et al., JACI 2019; 144, 193-203.

Highlights

A new vaccine for cat allergies

Cat allergy in human subjects is usually caused by the major cat allergen Fel d 1 and is found in approximately 10% of the Western population. Currently, there is no efficient and safe therapy for cat allergy available. We developed a new virus-like particle based vaccine to treat Fel d 1-induced allergy in human subjects by immunizing cats against their own major allergen, Fel d 1. The vaccine successfully induces Fel d 1 binding antibodies in cat and these antibodies neutralize the allergens thereby reducing symptoms of allergy in owners. Thoms F et al., JACI 2019; 144, 193-203.

A new vaccine for peanut allergy

Peanut allergy is a severe and increasingly frequent disease. We generated vaccine candidates against peanut allergy based on the plant-virus derived VLP coupled to the whole extract of roasted peanut or to the single major allergens Ara h 1 and Ara h 2 in a mouse model of peanut allergy. The vaccines demonstrated strong immunogenicity and were able to protect allergic reactions to allergens extracts. Storni F et al., J Allergy Clin Immunol. 2019 Dec 19. pii: S0091-6749(19)31709-9.

A novel treatment protocol for giant cell arteritis

Large vessel vasculitides such as giant cell arteritis and Takayasu arteritis are still treated with long-term, high-dose glucocorticoids, leading to a broad range of severe side effects. After our first RCT (published in Lancet 2016) which showed a glucocorticoid-sparing effect of >50%, we sought to prove that blockade of the IL-6 pathway can induce disease remission and replace glucocorticoids. The interim analysis shows a very successful outcome, and, in addition, it proves that high-resolution ultrasound qualifies to monitor inflammatory changes in the arterial vessel wall.

Disruptive anti-IgE inhibitors prevent mast-cell dependent early airway response in viable atopic lung tissue

IgE plays a central role in the pathophysiology of allergic asthma. In patients suffering from allergic asthma, airway mast cells are loaded with allergen-specific IgE. Inhibition of allergens leads to rapid activation and degranulation of these cells and the induction of allergic symptoms. In this study we demonstrate that disruptive anti-IgE inhibitors prevent mast cell activation in atopic precision-cut lung slices. Jiménez Delgado S et al., JACI 2019; 145, 719-722.e1
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- Bern Center for Precision Medicine (190402)
- Lungenliga Bern (NCT03924206)
- Bernese Cancer League (160901; 170907; 180401)
- China Scholarship Council (201606230238; 201708530258; 201606740067; 201808170004; 201906240084)
- External Partners: Institute General Physiology, University of Ulm, Germany

- Investigate how the mitochondria-related nucleotide metabolism and the DNA damage response machinery is associated with the tumor initiating capacity and chemotherapy response of NSCLC cancer stem cells
- Identify the molecular mechanisms conferring resistance to clinical standard therapies, with the goal to unravel druggable vulnerabilities - the ‘Achilles’ heel’ - of therapy-resistant cancer cells for development of innovative strategies to treat lung cancer
- Determine the immunosuppressive potential of the tumor-derived mesenchymal subset, which may serve as a novel therapeutic target in lung cancer; further, we are interested in identifying the molecular mechanisms underlying T cell dysfunction in the setting of lung cancer
- Ddentify the molecular mechanisms conferring resistance to classical standard therapies, with the goal to unravel druggable vulnerabilities - the ‘Achilles’ heel’ - of therapy-resistant cancer cells for development of innovative strategies to treat lung cancer and MPM
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- Teaching students of medicine as well as graduate students at the Graduate School for Cellular and Biomedical Sciences (GCB)
- Evaluation and therapeutic improvements by unportal minimally-invasive thoracic surgery techniques
- Investigation of different clinical and functional parameters with the aim of achieving better outcome and improvement of patient’s comfort
- Determine the immunosuppressive potential of the tumor-derived mesenchymal subset, which may serve as a novel therapeutic target in lung cancer; further, we are interested in identifying the molecular mechanisms underlying T cell dysfunction in the setting of lung cancer
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- Swiss Cancer League (KFS-4265-08-2017; KFS-3772-08-2015)
- China Scholarship Council (201606230238; 201708530258; 201606740067; 201808170004; 201906240084)
- Bernese Cancer League (160901; 170907; 180401)
- Lungenliga Bern (NCT03924206)
- Bern Center for Precision Medicine (190402)

- WEE1 is a drug target that promotes standard chemotherapy in MPM
- Malignant pleural mesothelioma (MPM) is a rare but very deadly cancer, notoriously known for dearth of therapeutic options. In a kinase CRISPR/Cas9 knockout screen, we identified the G2-M checkpoint kinase WEE1, whose inhibition synergistically enhances the efficacy of standard chemotherapy, leading to enhanced MPM cell death in vitro and in vivo.
- Xu et al. Mol Cancer Ther, 2019 Nov 6 [Online].

- Surgical smoke project
- Surgical fumes that arise when electrosurgery is used during surgery contain a variety of carcinogenic compounds. Our team has recently analyzed the exposure to these fumes in our operating rooms (ORs) during surgery, using real-time mass spectrometry analysis. The results of our study will be used to further improve already existing smoke evacuation systems as well as to develop additional preventive and protective measures against hazardous volatile substances in the OR.

- Adoptive T cell therapy (ACT) project
- Antigen-specific tumor infiltrating lymphocytes (TILs), represents a transformative living immunotherapy for the treatment of cancer. In lung cancer patients, we previously demonstrated that TILs show hallmarks of exhaustion coupled with downregulation of Wnt/β-catenin signaling and reduced expression of TCF7, an end effector of Wnt signaling. Therefore, we are testing a novel intervention aimed at targeting Wnt/β-catenin to generate ex vivo expanded TILs with renewed and sustained functional ability to eliminate malignant lung cancer cells.

- Thoracoscopic approach does not compromise survival
- The full thoracoscopic approach to major pulmonary resections is considered challenging and controversial as it might compromise oncological outcomes. The aim of this work was to analyse the results of a full thoracoscopic technique in terms of nodal upstaging and survival in patients with non-small-cell lung carcinoma (NSCLC).
- We found that long-term survival is not compromised by a full thoracoscopic approach. Our results compared favorably with those of video-assisted techniques. Lutz JA et al. Eur J Cardiothorac Surg. 2019; 1(5);2:263-270.

- Influence of an early assessment and management of dysphagia after lung resection
- Aspiration resulting from oropharyngeal dysphagia has been recognized as a serious complication after lung resection. In a single-center, randomized controlled trial, we investigated the influence of a clinical dysphagia assessment before the initiation of oral intake after surgery. The risk of pneumonia and the median length of hospital stay were compared between the intervention and the control group. We demonstrated that early detection of postoperative dysphagia can significantly decrease the risk of postoperative pneumonia. Schmidt Leuenberger et al., Ann Thorac Surg, 2019, 108 (4), 1059-1064.
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Prof. George N. Thalmann
Prof. Fionia Burkard
PD Dr. Marianne Krathof-de Julio
Prof. Katia Monastyrskaya
PD Dr. Roland Seiler

Profile
- Teaching students of medicine, biomedicine and biology as well as graduate students at the Graduate School for Cellular and Biomedical Sciences (GCB) at the University of Bern, CH and Leiden, NL
- 4 Groups
- Investigation of pathophysiological processes that contribute tumor diseases and bladder dysfunction
- Aim: Understand the molecular processes of urological cancer progression and bladder dysfunction
- External Partners: Alvaro Aytes Institut d’Investigació Biomèdica de Bellvitge, Program Against Cancer Therapeutic Resistance / Prostate Cancer Lab, Barcelona, Spain; Theodore Alexandrov EMBL-Heidelberg, Structural and Computational Biology / Spatial Metabolomics Lab, Heidelberg, Germany; Cristina Müller, Center for Radiopharmaceutical Sciences ETH-FS-USZ, Paul Scherrer Institut, Zürich, Switzerland; Sylvia LeDevedec, Leiden Academic Centre for Drug Research, Netherlands; Rosalyn Adam, Urological Diseases Research Center, Boston Children’s Hospital, Boston, USA; Duke University, Durham USA; Swiss Institute for Bioinformatics, NCCR-ICO, ETH-PSI-USZ, Paul Scherrer Institut, Zürich, Switzerland

Grants
- Swiss National Science Foundation (#179265, #184933, #189149, #189369, #175773/1, #182966/1)
- Department of Defense W81XWH-19-1-0432
- Cancer League 4KS 4718-02-2019
- HORIZON 2020 Marie Sklodowska-Curie Action (STOp 748836)
- Bern Center for Precision Medicine
- Wings for life Spinal Cord Research Foundation
- Novartis Foundation for medical-biological Research
- Innosuisse Grant Nr 41236.1 IP-LS

Highlights

Preclinical Models for the Translational Study of Urological Cancers
We have generated patient derived organoids (PDOs) and patient derived xenografts (PDXs) of primary and metastatic PCa and other malignancies on tumor types (Karkampouna et al. 2018). We have also developed and implemented a clinically relevant culture system for studying tumor tissue ex vivo. This technique allows cultivation of tumor slices and needle biopsies in a tissue culture setting without loss of normal architecture, viability, proliferative properties, or expression of specific markers. We have shown that the effects of drug treatment in this system are consistent with those observed on organoids (in vitro) and PDXs (in vivo).

Discovery of diagnostic and prognostic biomarkers of the “point of no return” for bladder function
Lower urinary tract dysfunction (LUTD) has different underlying mechanisms and multiple confounding factors. LUT symptoms are often shared between the diseases of different origin, delaying and complicating therapy. We are in the possession of a unique collection of human biopsy samples from patients with well-characterized bladder functional phenotypes. Our approach relies on generation and analysis of big gene expression data to reveal the triggers of LUTD. To achieve our goal of unbiased classification of LUTD and identification of molecular drivers of pathologic bladder remodelling, we apply machine learning algorithms to the transcriptome data.

Microvasculature on CHIP
Metastasis spread is a multistep process that depends on the capacity of specific cancer cells to leave the primary tissue, extravasate in the lymphatic and/or vascular network, and survive in the flow, extravasate and colonize the distant tissue. Organ on chip technology has provide us novel tools that overcome some of the limitations of traditional 2D and 3D in vitro models, reduce the phylogenetic problem due to differences between human and animals and if considered as a pre-screening tool significantly reduces the number of animals for further in vivo validation.

Urodynamic studies in awake mice with obstruction and SCI.
Along with recruiting the human spinal cord injury (SCI) patients for a longitudinal study of gene expression changes during neurogenic LUTD, we carry out a translational project to perform urodynamic studies in awake mice with obstruction and SCI. We investigate the impact of obstructive and neurogenic LUTD on bladder remodeling and functionally validate the findings in human patients. Our earlier studies in bladder outlet obstruction have detected signs of macrophage infiltration by analysing the total RNA in the BPO patients’ biopsies for macrophage-specific gene expression markers. We are now using mouse models of SCI and PBOO to investigate the role of infiltrating macrophages in bladder dysfunction. Our aim is to develop tools to mitigate the impact of cellular inflammatory pathways, preserving bladder contractility.

Myeloid cells (green) and infiltrating macrophages (red) in control mouse bladder

Hierarchical clustering of significantly regulated mRNAs and miRNAs in benign prostatic obstruction patients

Prostate Cancer Organoids

Urodynamic studies in awake mice with obstruction and SCI.
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First Swiss national Kidney Paired Exchange Donation and Transplantation

On 20 November 2019 a kidney from a living donor from Bern was implanted in Zurich and vice versa a kidney from Zurich was transplanted in Bern. This was the result of a joint effort of all transplant centres in Switzerland and Swisstransplant to offer living donor transplantation for couples with immunological contraindication to transplant in their respective combinations, either due to HLA or bloodgroup incompatibility. Using a novel software, two compatible donor and recipient pairs were identified among 12 candidates and cross over transplantation performed. Both procedures in Zurich and Bern went well and were finished early afternoon on the same day. Both recipients were discharged without complications and with good kidney function.

Terlipressin in surgical patients

A randomized placebo controlled trial was concluded and revealed that the splanchnic vasopressor terlipres- sin can reduce the incidence of renal but not hepatic complications after major liver surgery.

CAS in Sex- and Gender-Specific Medicine established

In cooperation with the Medical Faculty of the University of Zurich, the University Clinic for Visceral Surgery and Medicine established the first continuing education course in gender medicine in Switzerland. National and inter- national experts will reveal the importance of gender in the various medical disciplines.

New curriculum in Hepatology

The University of Bern has new a curriculum in Hepatology. In 2019 Dr. Abdoul Sow and Dr. Daye Ka, both from Senegal, were the first to receive a Certificate of Advanced Studies in Hepatology and Dr. Yuly Mendoza from Colombia a Diploma of Advanced Studies in Hepatology. This unique curriculum allows physicians interested in liver diseases to follow a struc- tured teaching covering all aspects of hepatology in a leading university clinic in this field.

Outreach into the undeveloped world

In 2019 we have started collaborative work with colleagues at the University Hospitals of Harare and Nairobi. Our objective is to address why there is intestinal failure in young children caus- ing stunting with poor physical and cognitve development as a result of abnormal colonization of the intestine by microbes that do not normally cause any problems. The work is funded by the generosity of the Botnar foundation in collaboration with colleagues at the ETH and University of Basel. As a result of this outreach, we have already hosted colleagues from Zimbabwe in Bern, and Benjamin Misselwitz and Andrew Macpherson have been working in Harare and Nairobi.
Clinics at the University Hospital, Universitäre Psychiatrische Dienste (UPD)

University Hospital of Old Age Psychiatry and Psychotherapy (APP)
University Hospital of Child and Adolescent Psychiatry and Psychotherapy (KJP)
University Hospital of Psychiatry and Psychotherapy (PP)
Electrical stimulation reduces negative emotions and thereby improves memory

We applied transcranial direct current stimulation (tDCS) to the prefrontal cortex in 94 healthy individuals (n = 43 young, n = 51 elderly) while they performed an episodic memory task. In young but not in older participants, moderation models indicated that the reduction of negative emotions via tDCS led to better memory performance. Our study provides a further link between emotion and memory: as increased activity in the prefrontal cortex is crucial for successfully coping with affective interference, anodal tDCS seems to help preventing negative and probably distracting thoughts.

Boosting slow wave sleep by closed loop acoustic stimulation to enhance memory performance in older adults

Reduced slow wave sleep is associated with cognitive decline and recent work suggests that it might even causally contribute to its development. As part of ongoing collaborations, we successfully adapted a closed loop acoustic stimulation approach that is able to detect slow waves in a sleeping person’s electroencephalogram and presents short and soft tones temporally aligned with the peaks of the slow waves. This procedure has shown to improve slow wave sleep as well as specific forms of short-term memory functions. In our project we are particularly interested in a range of cognitive tasks as well as the long-term effects of our intervention in an at risk group.

Alternating direct current stimulation improves memory performance

Associative memory is among the first cognitive functions negatively affected by aging. In a randomized, sham-controlled, and double-blind study, healthy older adults (n=28; mean age 71) received anodal transcranial direct current stimulation (tDCS), alternating current in the theta range (tACS), and sham stimulation over the left ventrolateral prefrontal cortex each once during encoding of an associative memory task. Our results suggest that tACS might be more effective to improve associative memory performance than tDCS - at least in higher aged samples.
University Hospital of Child and Adolescent Psychiatry and Psychotherapy (KJP)

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Profile
- 4 research groups, 22 staff members
- Investigating mental disorders with an onset during childhood and adolescence up to emerging adulthood
- Focus on: neurobiological mechanisms, early precursors, innovative treatment development, psychotherapy research, e-mental health
- Teaching students of Medicine and Psychology at all educational levels
- External Partners: Department of Child and Adolescent Psychiatry, Psychiatric Clinics of the University of Basel, Basel, Switzerland; Institute of Pharmacology and Toxicology, University of Zurich, Zurich, Switzerland; Department of Economics, University of Zurich, Zurich, Switzerland; Developmental Clinical Psychology Research Unit, Faculty of Psychology and Educational Sciences, University of Geneva, Geneva, Switzerland; Department of Child and Adolescent Psychiatry, Heidelberg University, Heidelberg, Germany; University Hospital of Child and Adolescent Psychiatry, Medical University Vienna, Vienna, Austria; University of Trondheim, Trondheim, Norway; The National Centre of Excellence in Youth Mental Health, University of Melbourne, Australia; Division of Sleep Medicine, Harvard University, USA; Department of Psychology, University of Zurich, Switzerland; Psychiatry and Human Behavior, University of California, Irvine, USA; Department of Psychiatry and Human Behavior, Brown University, USA; Child Psychiatry Branch, National Institute of Mental Health, USA; Upstate Medical University, Syracuse, New York, USA; School of Psychiatry, University of New South Wales, Sydney, Australia & Neuroscience Research, Sydney, Australia; Center for Psychiatric Neuroscience, Department of Psychiatry, Lausanne University Hospital, Lausanne, Switzerland

Grants
- Interfaculty Research Cooperation Grant - Decoding Sleep: Sleep subtypes in adolescent depression: Sleep physiology and treatment (PI: Dr. Tarokh)
- SNF Scientific Exchange Grant (182891): The Neurobiological Mechanisms of Auditory Verbal Hallucinations in Youth with Borderline Personality Disorder (PI: Dr. Michel)
- SNF Project Grant (182839): Neurobiological Mechanisms of Pain Dependent Stress-Regulation in Adolescent Non-Suicidal Self-Injury (PIs: Prof. Dr. Kaess & PD Dr. Koenig)
- SNF Project Grant (184943): Sleep Neurophysiology: A Window onto Adolescent Mental Health (PI: Dr. Tarokh)

Highlights

Effective Brief Psychotherapeutic Intervention for Adolescent Non-Suicidal Self-Injury

We developed a manualized brief cognitive behavioral intervention for the treatment of adolescent non-suicidal self-injury (NSSI). Within a randomized controlled trial, the brief psychotherapeutic intervention conducted by trainees in psychotherapy led to an equal and quicker reduction of NSSI when comparing it to more intensive outpatient psychotherapy conducted by professional psychotherapists.

Kaess et al. European Child and Adolescent Psychiatry, in press.

Depressed Adolescents Show Altered Sleep Neurophysiology

Depression is often accompanied by subjective sleep complaints and alterations in sleep neurophysiology. In this study, we show that more depressive symptoms are associated with less sleep spindle activity. Sleep spindles, a unique feature of the sleep EEG, have been hypothesized to protect sleep against environmental disturbances. Thus, diminished spindle activity may be a subtle sign of disrupted sleep and its association with depressive symptoms in early adolescence may signal vulnerability for depression.


First Brain Imaging of Abnormal Involuntary Movements in Early Psychosis

We measured motor symptoms, regional cerebral blood flow (rCBF) and cortical grey matter volume (GMV) in patients with first episode psychosis (FEP), clinical high risk (CHR) and controls. Our video-based analysis demonstrated that abnormal involuntary movements were more frequent in CHR and FEP as compared to clinical controls and correlated with increased rCBF but decreased GMV in the premotor cortex. Thus, our study shows for the first time an association between motor symptoms and biological markers such as rCBF and GMV in early psychosis.


Sibling Study Shows Altered Cortisol Response to Memories of Early Adversity in Adolescent Non-Suicidal Self-Injury

We compared adolescents engaging in non-suicidal self-injury (NSSI) and their siblings with regard to hair cortisol concentrations as well as salivary cortisol concentrations before and after an interview assessing experiences of childhood adversity within the family environment. Our results showed that adolescents engaging in NSSI reported more severe experiences of childhood adversity, exhibited higher hair cortisol levels and had a significantly higher decrease in salivary cortisol levels during the trauma interview. The results point to increased basal hypothalamic-pituitary-adrenal axis functioning but decreased reactivity to stress such as traumatic memories in adolescents with NSSI.


Promising Findings for Transcutaneous Vagus Nerve Stimulation in Adolescent Depression

In a first experimental trial, we investigated the effects of transcutaneous vagus nerve stimulation (tVNS) on emotion recognition in adolescents with major depressive disorder (MDD). Compared to sham stimulation, tVNS decreased the recognition of briefly presented stimuli of negative valence in patients with MDD. These findings suggest a potential anti-depressant effect of tVNS by reducing the negative bias in adolescents with MDD. Further research is needed, addressing the potential of tVNS as treatment option for MDD in children and adolescents.

Koenig et al. Psychological Medicine, in press.

Frequency of NSSI at baseline, after 4 months, and after 10 months

Electrode placement for tVNS (A) versus sham (B) stimulation

Depressive Symptoms and Sigma Power

Functional correlates (rCBF) of abnormal involuntary movements in psychosis risk and first episode psychosis

Salivary cortisol levels in adolescents engaging in NSSI and their siblings before and after a trauma interview
University Hospital of Psychiatry and Psychotherapy (PP)

**Profile**

- Teaching students of medicine, biomedical science, and psychology within the curriculum, supervision of masters and doctoral theses projects, as well as teaching at various graduate schools at the University of Bern
- 15 research groups
- Clinical translational research topics include psychosis pathobiology and treatment, sleep, addiction, stress, and depression
- Aims: understand the etiology of psychiatric disorders and develop novel non-pharmacological treatment options
- External partners: Dept. of Psychiatry, Northwestern University, Chicago, USA; Dept. of Psychiatry, Vanderbilt University, Nashville, USA; Dept. of Psychiatry, Philipps-University Marburg, Germany; Dept. of Psychiatry, Karolinska Institute, Stockholm, Sweden; Dept. of Psychiatry, University of Freiburg, Germany; Dept. of Psychiatry, University of Würzburg, Germany; University of Amsterdam, The Netherlands; University of Milano, Italy; Depts. of Psychiatry, University of Basel and University of Zurich; Neurocenter of the University of Lugano

**Grants**

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- Prof. Daniela Hubl
- Prof. Andrea Fedenigel
- Prof. Thomas König
- Prof. Wolfgang Tscharner

**Highlights**

**Sleep orchestrates indices of neural plasticity**

Animals and humans spend on average one third of their lives in sleep, but its functions remain to be specified. We used electrophysiological (TMS, EEG), behavioral (memory, vigilance) and molecular (BDNF, cortisol) indices and provide first integrative evidence in humans that sleep strengthens relevant connections between neurons while keeping overall neural network function stable.


**Hand gestures improved with single session TMS**

Schizophrenia patients have severe gesture deficits, that are linked to alterations in the cerebral praxis network. This double-blind, randomized controlled trial tested the effects of single sessions of transcranial magnetic stimulation and detected one superior protocol. This protocol is now being tested in a parallel-arm RCT.


**Electrophysiological Brain Connectivity**

Leading EEG experts provided a consensus review on the current state of the theory and the available algorithms for the estimation of brain connectivity based on electrophysiological data. They developed a common theoretical framework, and explained the common pitfalls.


**Coping styles improved with specialized interventions after suicide attempts**

Coping strategies are critical for secondary suicide prevention. We demonstrated 11% less dysfunctional coping in the ASSIP group and 6% more problem-focused coping compared to the control group after 24-months. Thus, specialized psychotherapy was superior over treatment as usual.

Would you like to receive further copies of the annual report from the Medical Faculty?

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Cover Picture: Graphic illustrating the influence of individual genes and proteins in a medical treatment. Khrystyna Okhrim and the Bern Center for Precision Medicine, Prof. Dr. Mark Rubin.

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