



Universität
Basel

Faculty of Medicine



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RC2NB

ANNUAL REPORT

2021

About RC2NB

The Research Center for Clinical Neuroimmunology and Neuroscience Basel (RC2NB) is based on a non-profit foundation. It was founded in 2019 by the University Hospital Basel with participation of the University of Basel to continue and enhance the long-standing commitment and internationally renowned, clinically oriented research for patients with multiple sclerosis and other neuroimmunological diseases. RC2NB coordinates and supports several competitively funded research groups, dedicated to improving the clinical, imaging, biochemical, molecular, and cellular characterization of the disease process and understanding the benefits and side effects of newly developed therapies. Switzerland's largest MS center, the estab-

lished high-quality patient cohorts coordinated from here, the local, national, and international networks as well as academic partner institutions and collaborating industry provide optimal conditions for RC2NB's mission. With its interdisciplinary team and its alignment of basic research, clinical research, and patient care, RC2NB aims at rapid translation of research results into advances of patient treatment and diagnosis. Main activities of RC2NB include the development of innovative digital biomarkers, the establishment of structures and expertise for managing and processing large volumes of highly complex data, and the application of cutting-edge analytic approaches, including artificial intelligence.

Vision

RC2NB's mission is to strengthen internationally recognized expertise and innovative research projects and complement them with the development and validation of digital biomarkers and innovative methods of information processing and artificial intelligence.

Mission

Improving the life of people with MS and neuroimmunological diseases through the development of innovative tools that comprehensively characterize the disease process, facilitate the development and implementation of better treatments and enable personalized disease management.

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1 | Introduction

The year 2021 was marked by the implementation of the new structures within the centre and the coordination with internal and external partner institutions and groups. In parallel, important milestones were achieved in all three workstreams. A few may be mentioned here as representative:

In **Workstream 1**, the completion and evaluation of the feasibility study for dreaMS, which formed an important basis for the recognition of the dreaMS App as "software as medical device" by Swissmedic and the EU. All necessary preparations for the first large two-year validation study for dreaMS with participants of the Swiss MS cohort study are completed with approval by the competent authorities in November 2021. The completion of the technical fine-tuning for dreaMS and the digital platform and the inclusion of the first patients is foreseen in the course of the first quarter of 2022.

In **Workstream 2**, important prerequisites for the application of Neurofilament Light (NfL) Chain as a blood-based biomarker for MS in daily practice were created with the validation of the modelling of z-values

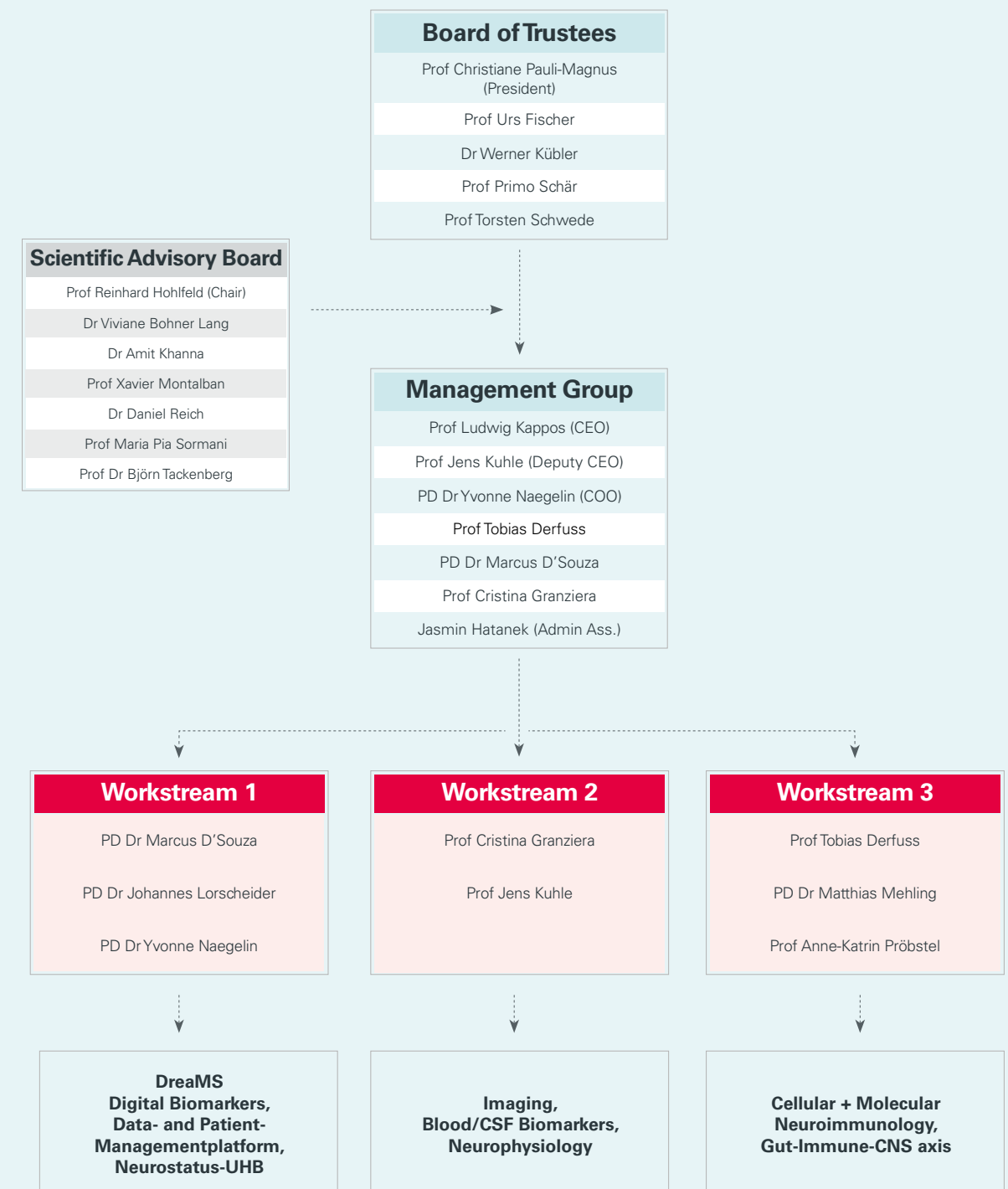
for NfL in two large, independent MS cohorts. Important publications underline the progress made in developing innovative methods for better tissue characterisation in neuroimaging.

In **Workstream 3**, in addition to contributions to a better understanding of the effects and side effects of immunotherapeutics, we can report important highlights with the award of an Excellence Professorship to Anne-Katrin Pröbstel and the establishment of her research group in the Department of Biomedicine. Number and quality of publications, various scientific prizes and mandates in national and international organisations provide further evidence of the productivity and recognition of the researchers joining forces in and with the RC2NB. The continued support from the University Hospital, the University, national and international research institutions and from our sponsors and cooperation partners in industry constitutes an important prerequisite and motivation of further progress!

Ludwig Kappos
CEO

2 | Governing Bodies

2.1 | Organizational Chart



2 | Governing Bodies

2.2 | Board of Trustees

Members

- Prof Christiane Pauli-Magnus (President of the Board, Head of Department of Clinical Research, University Hospital Basel)
- Prof Urs Fischer (Chairman Neurology, University Hospital Basel)
- Dr Werner Kübler (CEO University Hospital Basel)
- Prof Primo Schär (Dean Medical Faculty, University of Basel)
- Prof Torsten Schwede (Vice-president Research, University of Basel)

The Board of Trustees held two meetings, on 20 January 2021 and 26 August 2021.

2.3 | Scientific Advisory Board

Members

- Prof Reinhard Hohlfeld (Chair), Munich, Germany
- Dr Viviane Bohner Lang (Patient Representative), Allschwil, Switzerland
- Dr Amit Khanna, Basel, Switzerland
- Prof Xavier Montalban, Barcelona, Spain
- Dr Daniel Reich, Bethesda, United States of America
- Prof Maria Pia Sormani, Genova, Italy
- Prof Björn Tackenberg, Basel, Switzerland

The international RC2NB Scientific Advisory Board (SAB) meets annually and independently reviews the work and provides advice to the RC2NB. The first and constituting meeting was held (virtually) on 16 June 2021.

Quote from the SAB report issued after this meeting:

„Together with support from industry, the sustained strong commitment of both the University Hospital and the University of Basel provides excellent conditions for ensuring continuation of the superb translational neuroimmunology research for which the MS center in Basel is known. The scope and quality of the presented projects is outstanding.

In particular, the SAB appreciates that most projects are already supported by competitively acquired external funding, testifying to the excellence of the projects and their PI’s. The SAB recommends that RC2NB uses its own resources to foster collaboration and synergies between the different groups of investigators.“

2.4 | Management Group

Members

- Prof Ludwig Kappos, CEO, Workstream 1 and 2
- Prof Tobias Derfuss, Workstream 3
- PD Dr Marcus D’Souza, Workstream 1
- Prof Cristina Granziera, Workstream 2
- Prof Jens Kuhle, Deputy CEO, Workstream 2
- PD Dr Yvonne Naegelin, COO, Workstream 1
- Jasmin Hatanek, Management Assistant, Administration

Members of the management group represent the three workstreams of RC2NB and meet monthly to facilitate continuous exchange on and coordination of ongoing and planned research projects.

3 | Scientific Achievements

Three workstreams - One vision

Three closely linked workstreams pursue the common goal of the RC2NB. Interdisciplinary teams collaborate within and across the workstreams to develop innovative tools for

monitoring the health of patients with MS, better understand the disease process, enable personalized disease management, and find better treatments.

DreaMS Study Team – (from left to right): Corne de Jong (Healios), Nancy Wochnik, PD Dr Marcus d'Souza, Dr Tim Wölfle, Silvan Pless, Vera Müller, PD DrYvonne Naegelin, Prof Ludwig Kappos, Guilhem Dupont (Healios), Dr Andrea Wiencierz, PD Dr Johannes Lorscheider



3.1 | Workstream 1: Digital future

Research Group Leaders

PD Dr Marcus D’Souza (Neurostatus-UHB)

PD Dr Johannes Lorscheider (dreaMS)

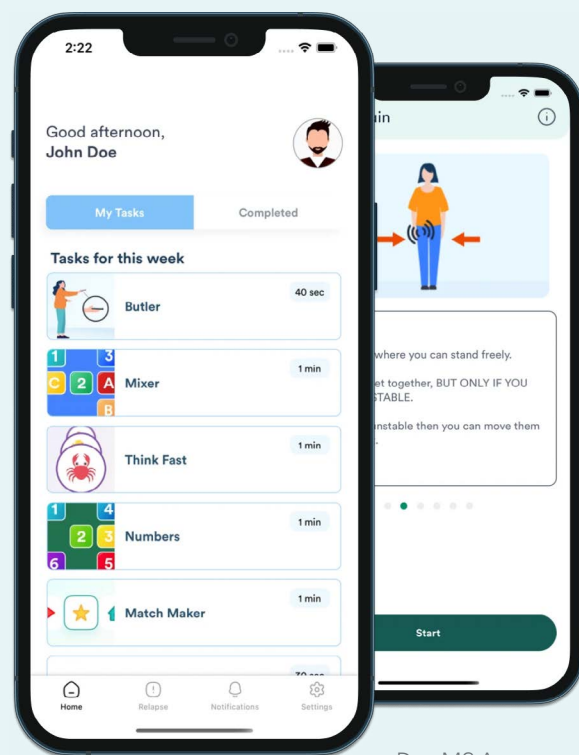
PD DrYvonne Naegelin (dreaMS)

The goal of the project "dreaMS" (Yvonne Naegelin, MD and Johannes Lorscheider, MD) is to develop and validate digital biomarkers for people with Multiple Sclerosis (MS). The built-in sensors in today's smartphones allow differentiated analyses of movement patterns. At the same time, smartphones are broadly used on a daily basis and can therefore provide a more realistic picture of the activities of their owners than visits to the doctor's office, which are only semi-annual or annual. DreaMS uses a smartphone app for this purpose that provides measures of movement and fine motor skills but also visual function and cognition through specifically developed tasks ("challenges") and also assesses mood and quality of life. This app is developed in cooperation with the Basel-based company Healios Ltd, which specializes in software solutions in the field of "digital health". This cooperation for the development of the dreaMS App as a set of digital biomarkers for MS and other neuroimmunological diseases was initially supported by the Swiss Agency for Innovation (Innosuisse). The project also aims at creating a digital solution that allows the customized use and

secure processing of information obtained with this application by people with MS, health care professionals and researchers in daily management and clinical research. This digital solution also allows the integration and common analysis of additional diagnostic and digital health tools as well as data from laboratory tests or neuroimaging for a comprehensive assessment of disease evolution and treatment effects. A first feasibility study (NCT04413032) was completed in 2021: in this study, dreaMS was tested for technical reliability, acceptance and meaningfulness in a group of MS patients and healthy volunteers. The pre-defined reliability criteria were met by 72 of 133 features extracted from the active tasks ("challenges") included in dreaMS. In ratings by the participants all challenges reached high scores for acceptance and meaningfulness for Multiple Sclerosis. Based on these results in May 2021 the dreaMS app obtained the qualification for software as class I medical device (SAMD) for Switzerland and the EU.

As next step in the development a large 2-year validation study (NCT05009160) with approx. 400 participants from the

Swiss MS cohort and a group of healthy volunteers was designed and approved by the institutional review board of northwestern Switzerland (EKNZ) and Swissmedic in November 2021. First patients will be included within the first quarter 2022. First discussions for the development of validated digital biomarkers to be accepted as endpoints in clinical trials were held with both the European Medicines Agency (EMA) in October and with the U.S. Food and Drug Administration (FDA) in November 2021. Both agencies appreciated and welcomed the concept of developing apps for assessing and monitoring disease symptoms in subjects with MS and acknowledged the interaction with RC2NB at a very early timepoint of development.



DreaMS App

Healios Ltd refined their in house procedures during the course of the project and as defined within our agreement establishing several ISO standards for ensuring to meet all standards for further regulatory approvals.

The research activities of Neurostatus-UHB are also part of workstream 1. In November 2021 Neurostatus-UHB was established as a public limited company (stock corporation under Swiss law) owned by the University Hospital. Activities of this spin-off include licensing the implementation and use of Neurostatus-EDSS and its digital version, the Neurostatus-eEDSS, in international corporate sponsored and academic clinical trials for MS and other neuroimmunological disorders. Neurostatus does also provide certification and training for users of the Neurostatus-(e)EDSS and engages in the development, testing and implementation of new digital clinical assessment and data capture tools for use by health care professionals. In 2021, Neurostatus-EDSS was licensed to 92 phase II/III MS trials, of which 30 are using the digital version. The digital Neurostatus-EDSS was developed and is provided in collaboration with three eCOA (electronical clinical outcome assessment) companies under non-exclusive Neurostatus-UHB licenses.

For clinical trials Neurostatus-eEDSS is provided in combination with a review and quality assessment service by an EDSS-Expert team. Income generated through these activities and licensing is used to support research activities coordinated by RC2NB.

3.2 | Workstream 2: Innovative imaging and analysis of body fluids

Research Group Leaders

Prof Cristina Granziera (advanced neuroimaging research –ThINk Basel)

Prof Jens Kuhle (Swiss MS Cohort Study and Body Fluid Biomarker Laboratory)

The year 2021 has been a very fruitful period for both Clinical Neuroimmunology/SMSC and the advanced neuroimaging group (ThINk Basel).

ThINk Basel has grown to a leadership role in the Basel area, in Switzerland and in the international neuroimaging and neurology community. Reflecting scientific achievements are 20 original research publications, co-authored or authored by ThINk members in highly ranked neurology and neuroimaging journals. In all those works, ThINk investigators exploited the sensitivity and specificity of advanced magnetic resonance imaging to identify new mechanisms of disease progression and brain repair in multiple sclerosis patients. In 2021, Cristina Granziera was awarded a 2-years prolongation of the Swiss National Fund Professorship with the project entitled “INSIDER-RAP: ImagiNg the Interplay between Axonal DamagE and Repair in Multiple Sclerosis – Remyelinating lesions And Pathways”. In collaboration with PD Dr M. Bach Cuadra (University of Lausanne), Prof A. Depeursinge and Prof J. Henning (University of Applied Sciences Western Switzer-

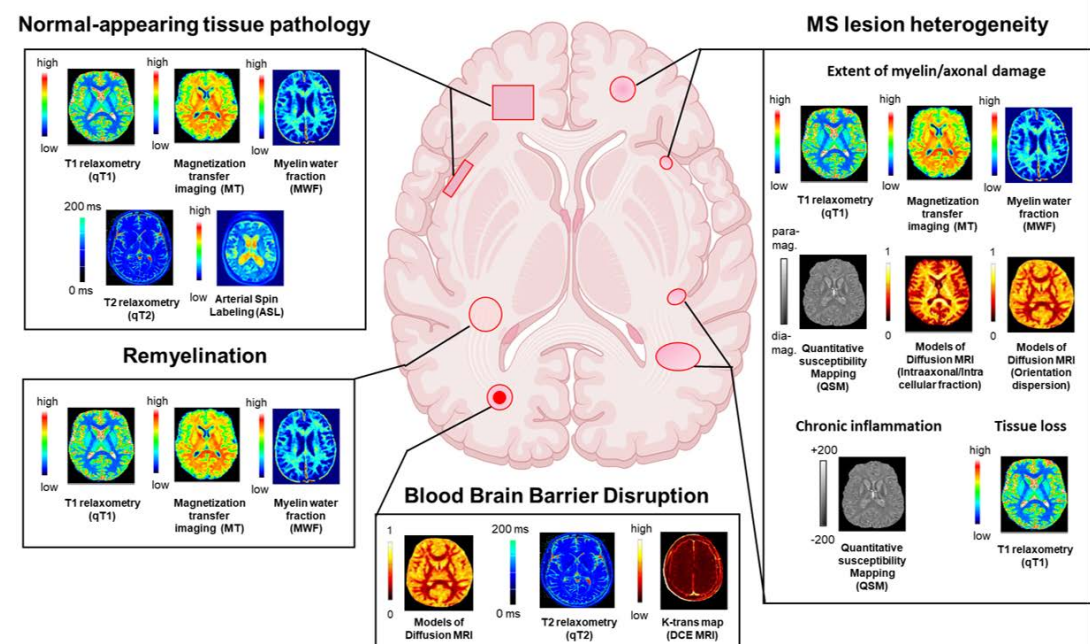
land) C. Granziera is a co-PI of a project funded by the “Hasler Foundation” entitled “MSxplain- Explaining AI decisions in personalized healthcare: towards integration of deep learning into diagnosis and treatment planning for Multiple Sclerosis”. In 2021, C. Granziera was elected secretary of the prestigious White Matter group at the International Society of Magnetic Resonance Imaging, and also co-president of the Medico-Scientific Advisory Board of the Swiss MS Society and member of the Executive Board of the Department of Biomedical Engineering at the University of Basel.

A central resource for our MS research is the Swiss Multiple Sclerosis cohort (SMSC; PI Jens Kuhle) which was founded in 2012 and comprises eight leading Swiss MS centers. It is one of the internationally largest MS research databases covering more than 1500 well characterised patients. Patients are followed every 6 or 12 months and high quality clinical, radiological as well as biological data are collected. Examinations are performed by Neurostatus certified raters and MRIs obtained according

to a predefined, standardized protocol in all participating centers. Central MRI analysis is done by ThINK Basel. More than 250'000 bodyfluid samples from 10'202 timepoints have been stored meanwhile. In 2021 the Clinical Neuroimmunology/SMSC group has relevantly contributed to international research on biomarkers in MS. We demonstrated that an intrathecal IgM synthesis (present in about 25 % of persons with MS (pwMS)) predicts a more active and severe disease course and an increased need of a high-efficacy disease modifying treatment (Ref. 79). We published work on generating a reference database for serum neurofilament light chain (sNfL) in more than 10000 serum samples from more than 5 000 control persons. The value of this normalised NfL

metric (NfL Z score or percentile) for use in clinical practice was illustrated in two independent patient cohorts comprising >5'000 pwMS, the SMSC and the Swedish MS registry (Müller, Benkert, ...Kuhle et al., Lancet Neurology 2022,). Members of the Clinical Neuroimmunology/SMSC team authored >60 peer reviewed publications in 2021 and - in addition to ongoing funding by the Swiss National Research Fund - received research grants from the Progressive MS Alliance and the Swiss MS Society. In 2021, J. Kuhle was elected as member of the Scientific Leadership group of the international registry MS Base and as president of the Scientific Research Grant Committee of the Swiss MS Society.

Brain imaging discloses different aspects of pathology in MS lesions and normal-appearing MS tissue



3.3 | Workstream 3: Recording and understanding the dysregulated immune system

Research Group Leaders

Prof Tobias Derfuss (Cellular and Molecular Neuroimmunology)

PD Dr Matthias Mehling (Immunosenescence, Protective Immunity under DMT)

Prof Anne-Katrin Pröbstel (Experimental Neuroimmunology)

The research group of Prof Tobias Derfuss at the Department of Biomedicine consists of two Postdocs, three PhD students and several master students. It receives funding from the Swiss National Science Foundation, public foundations as well as industry. The research is focused on three main topics: (I) Immunomonitoring of immune therapies in MS; (II) Relevance of B-cell receptor mediated antigen capture by B cells; (III) Discovery of new autoantibodies and functional testing of autoantibodies in neurological autoimmune diseases. Achievements in 2021 include:

1. Further elucidation of immune alterations during dimethylfumarate therapy (Diebold et al., in press Ann Neurol 2022),
2. Identification of a MS-associated B-cell population by CSF mass cytometry (Ref. 52);
3. Understanding the relevance of anti-SARS-CoV2 specific IgM and IgA antibodies on virus neutralization was investigated using recombinant human antibodies (Callegari et al. EMBO reports in revision);
4. B cell antigen capture in CNS was shown to be a trigger of autoimmunity in animal models using B-/T-cell receptor transgenic animals (Kim at al. unpublished);
5. Discovery of a differential IgM reactivity in the CSF of MS patients led to the identification of a novel autoantigen (Callegari et al.

unpublished). 6. Using myasthenia as an exemplary autoantibody mediated disease cooperation of different auto-reactive antibody clones was shown to be pathogenic by inducing clusters of acetylcholin receptors in the membrane (Rose et al., in preparation).

The new treatment options developed for the treatment of MS allow effective control of inflammatory disease activity. Although these drugs have rather specific immunomodulatory than broad immunosuppressive effects, functional immunological alterations and premature immunosenescence occur in susceptible patients. Against this background, the research group of PD Dr Matthias Mehling at the Department of Biomedicine has initiated in 2021 two studies to assess the impact of DMTs on protective immunity in MS. The first study, organized together with Jens Kuhle, aims to determine the rate of confirmed SARS-CoV-2 infection and severity of COVID-19 in a cohort of MS patients and to quantify SARS-CoV-2-specific antibody response. The study is nested within the Swiss MS Cohort (SMSC). PCR-confirmed SARS-CoV-2 infections, severity of COVID-19 according to the WHO clinical progression scale and immunizations with SARS-CoV-2 vaccines were captured by questionnaires.

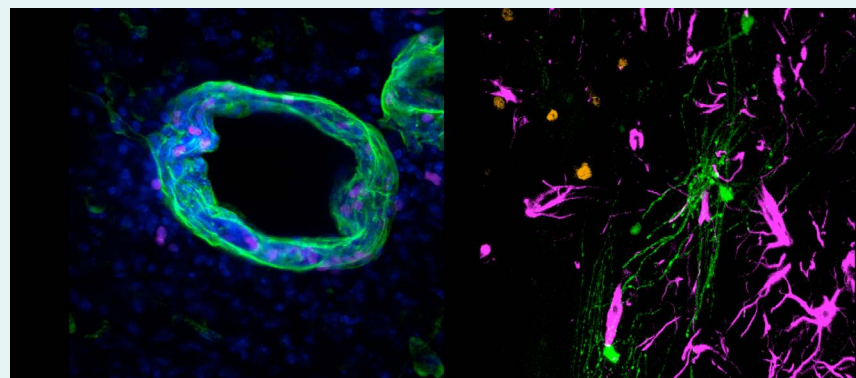
res used for interviews every 6 or 12 months. Anti-SARS-CoV-2 spike protein and nucleocapsid antibody levels are determined by electrochemiluminescence immunoassay in sera of all participants. Between February 2021 and January 2022, study questionnaires were completed for 863 pwMS. In parallel, SARS-CoV2 serology for anti Nucleocapsid and anti-Spike IgGs was performed in 1137 samples. The group is currently analyzing these data also regarding breakthrough disease, asymptomatic infections and vaccine responses.

The second study aims at cross-sectionally quantifying cellular changes linked to the development of immunosenescence by high dimensional immuno-phenotyping including multi-color flow cytometry and multiplex cytokine measurement in the blood of pwMS followed in the SMSC. In 2021 the expression of cell surface markers linked to immunosenescence was measured in various T cell subsets of 240 pwMS. These data are currently analyzed and will be integrated computationally using machine learning with results of a study organized in our MS Center that retro- and prospectively quantifies severity of infections in pwMS. The aim of this multidisciplinary research project is to

develop integrative algorithms to monitor treatment-associated immune deficiency in pwMS.

The research group “Experimental Neuroimmunology” of Prof Anne-Katrin Pröbstel at the Departments of Biomedicine and Clinical research was established in 2021 and is an interdisciplinary team of two postdocs, three PhD students, three MD doctoral students, one technician/lab manager, one bioinformatician, and two master students. The group is funded through an Eccellenza Professorship by the Swiss National Science Foundation (SNSF), the ERC (Horizon2020), the National Multiple Sclerosis Foundation, the Goldschmidt-Jacobson Foundation (Bed-to-Bench-for-Top Clinicians), the Propatient Foundation, intramural funding by the University of Basel as well as industry. The overall aim lies in understanding the functional diversity and specificity of B cells and their interaction with gut microbiota in central nervous system inflammation with a focus on MS, MOGAD, autoimmune encephalitis, and neurolupus.

The group strives to develop strategies to foster immune regulatory responses and achieve tailored depletion of immune cell subpopulations

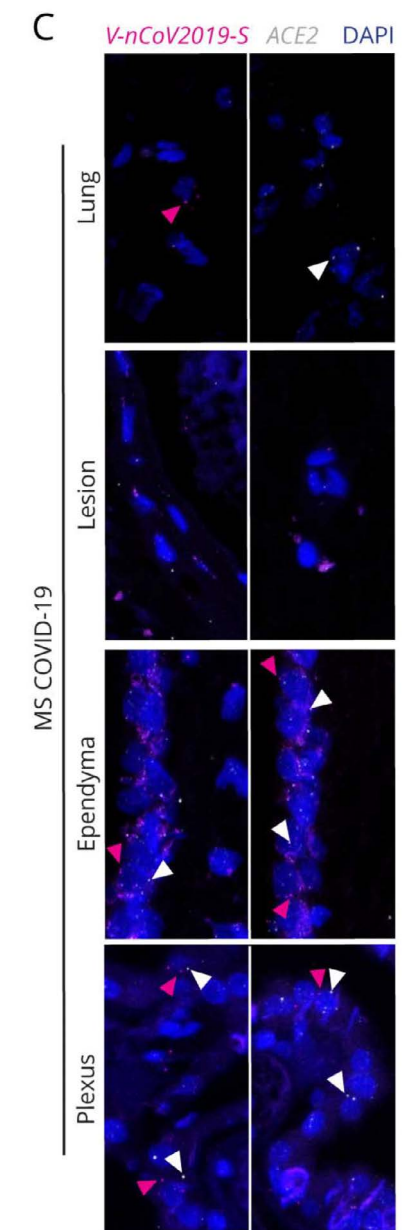


B lymphocytes in the brain in acute inflammation. The left picture shows a blood vessel (green) in the brain, with B-lymphocytes (pink) in the vessel wall and in brain tissue surrounding the vessel. The right picture shows virus-infected nerve cells (green), astrocytes (pink) and B-lymphocytes (yellow) in the brain.

through targeted manipulation of the gut microbiome. Research in the group focuses on three main topics: (I) deciphering microbial-immune cell crosstalk in MS, (II) decoding pathogenic B cell and antibody profiles in MOGAD, (III) identifying microbial and immune signatures associated with treatment (non-)response.

In 2021, microbial signatures associated with disease activity as well as with relevant side effects under immune modulating therapy in a patient subgroup were identified (Diebold et al. under review). Further, the group identified an immune trafficking signature in patients with MOGAD for which currently a therapeutic blocking antibody is evaluated in pre-clinical models (unpublished). Moreover, they discovered a novel mucosal originating autoantibody in a clinical distinct subgroup of patients with atypical demyelination (Gomes*, Kulsvehaugen* et al. in preparation). Finally, they contributed to elucidating the immune pathology of Neuro-COVID 19 in MS and non-MS patients (Ref.38; 84; Etter et al. under review). In 2021, Prof Pröbstel received the Fritz-and-Ursula-Melchers B cell immunology award of the German Society of Immunology, and the the Ingrid-zu-Solms-Prize for Medicine for her work on B cells and antibodies and their interaction with microbiota in MS. She has been appointed as member of the scientific committee of the Fondation ARSEP, representative of the Swiss Neurological Society in the Multiple Sclerosis Scientific Panel of the EAN, member of the steering committee

of the German Neurological Society for the “Guidelines on diagnosis and treatment of multiple sclerosis, neuromyelitis optica and MOGAD”. Several members of her group are supported with prestigious fellowships by international and Swiss institutions.



Sars-CoV-2 transcripts in the choroid plexus of MS and non-MS COVID 19 patients but not in MS lesion areas

4 | Financial Statement

Financial Statement	2021
Research contributions	3'750'000.00
Other income	333'176.12
Total Income	<u>4'083'176.12</u>
Technical development incl. expenses for third party services	-1'304'270.21
Personnel	-798'372.45
Administration and other expenses	-127'646.78
Total Expenses	<u>-2'230'289.44</u>
Ordinary result for the period	<u>1'852'886.68</u>

Equity	2021
Equity as of 01.01.2021	1'429'186.02
Income 2021	4'083'176.12
Expenses 2021	-2'230'289.44
Equity as of 31.12.2021	<u>3'282'072.70</u>

Expenses by Cost Centers

	Workstream 1	Workstream 2	Workstream 3
Personnel	-231'793.22	-199'764.78	-48'894.00
Consumables and other lab services	-12'833.66	0.00	0.00
Technical development	-1'293'245.21	0.00	0.00
Administration and other expenses	-40'911.00	0.00	0.00
Total Expenses	<u>-1'578'783.09</u>	<u>-199'764.78</u>	<u>-48'894.00</u>

	Data Storage and Analysis	Management/ Administration	Total
Personnel	-134'636.52	-183'283.93	-798'372.45
Consumables and other lab services	0.00	0.00	-12'833.66
Technical development	0.00	0.00	-1'293'245.21
Administration and other expenses	-25'511.00	-59'416.12	-125'838.12
Total Expenses	<u>-160'147.52</u>	<u>-242'700.05</u>	<u>-2'230'289.44</u>

5 | Main Partnering Institutions and Research Support



6 | Members and Collaborators of RC2NB by Workstreams

Workstream 1

Research Group Leaders

PD Dr Marcus D’Souza (Neurostatus-UHB)
PD Dr Johannes Lorscheider (dreaMS)
PD Dr Yvonne Naegelin (dreaMS)

Group Members and Collaborators

dreaMS and digital solution:

Melanie Lacalamita (study nurse)
Marko Obradovic, MSc (software engineer)
Vera Müller, MSc (study coordinator)
Silvan Pless, MSc (neuropsychologist, PhD candidate)
Nancy Wochnik (study nurse)
Dr Tim Wölfle, MSc (physician-scientist, PhD candidate)
(Employees of Healios Ltd. not individually mentioned)

Neurostatus-UHB:

Neurostatus-UHB:
Dr César Álvarez-González (neurologist)
Elena Börlin (IT specialist)
Lea Meret Brugger (student)
Dr Ilaria Callegari (neurologist)
Esther Maria Fricker (operations)
Dr Nuria Alicia Cerdá Fuertes (neurologist)
Eddy Angela Garcia Gorostiaga (operations)
Joel Götti (student)
Gabriel Simon Hug (student)
Dr Christian Kamm (neurologist)
Dr Giulia Mallucci (neurologist)
Svetlana Orlova (IT specialist)
Thomas Trouillet (programmer)
Colleen Waiz (operations)
Andrea Zimmer (study coordinator)

Workstream 2

Research Group Leaders

Prof Cristina Granziera (advanced neuroimaging research – ThINk Basel)
Prof Jens Kuhle (Swiss MS Cohort Study and Body Fluid Biomarker Laboratory)

Group Members and Collaborators

ThINk Basel

Prof Cristina Granziera team:

Dr Muhamed Barakovic (Postdoc)
Dr Alessandro Cagol (Postdoc)
Xinjie Chen (PhD candidate)
Riccardo Galbusera (PhD candidate)
Dr Lester Melia Garcia (senior researcher)
Osman Hatipoglu (master student)
Selina Leber (master student)
Marguerite Limberg (research assistant)
Po-Jui Lu (PhD candidate)
Dr Jannis Müller (Postdoc)
Reza Rahmanzadeh (PhD)
Dr Esther Ruberte (senior researcher)
Dr Gretel Sanabria Diaz (Postdoc)
Aida Suljakovic (personal assistant)
Dr Niena Siebenborn (neuroradiologist)
Sabine Schädelin, MSc (statistician)
Igor Schneider (master student)
Dr Alexandra Todea (neuroradiologist)
Dr Charidimos Tsagkas (Postdoc)
Dr Matthias Weigel (senior researcher)
Antonia Wengner (master student)

PD Dr Athina Papadopoulou team:

Dr Cesar Alvarez (MD)
Dr Katerina Ebner (MD)
Dr Nuria Cerde Fuertes (MD)
Dr Jenni Kuhlmann (MD)

PD Dr Regina Schläger team:

Valentina Crepulja (master student)
Eva Kesenheimer (PhD candidate)
Dr Laura Sander (MD)
Dr Janina Wendebourg (PhD candidate)

PD Dr Oezguer Yaldizli team:

Dr Jannis Müller (Postdoc)

PD Dr Katrin Parmar team:

Dr Charidimos Tsagkas (Postdoc)

Swiss MS Cohort Study and Body Fluid Biomarker Laboratory

Prof Jens Kuhle team:

Dr Pascal Benkert (statistician)
Caroline Brunner (study nurse)
Lilian Demuth (study coordinator)
Leila Develioglu (technician)
Melanie Lacalamita (study nurse)
Prof David Leppert (senior Postdoc)
Marguerite Limberg (study nurse)
Aleksandra Maleska, MSc (bioengineer)
Stephanie Meier (PhD candidate)
Dr Johanna Oechtering (senior neurologist/Postdoc)
Dr Annette Orleth (Postdoc)
Miriam Rhyner (study nurse)
Monica Röthlisberger (study nurse)
Sabine Schaedelin, MSc (statistician)
Suvitha Subramaniam, MSc (data scientist)
Dr Eline Willemse (Postdoc)
Nancy Wochnik (study nurse)
PD Dr Oezguer Yaldizli (senior neurologist/Postdoc)

Workstream 3

Research Group Leaders

Prof Tobias Derfuss (Cellular and Molecular Neuroimmunology)
PD Dr Matthias Mehling (Immunosenescence)
Prof Anne-Katrin Pröbstel (Experimental Neuroimmunology)

Group Members and Collaborators

Prof Tobias Derfuss team:

Dr Ilaria Callegari (PhD candidate)
Sebastian Holdermann, MSc (PhD candidate)
Hye-In Kim, MSc (PhD candidate)
Dr Nicholas Sanderson (Postdoc)
Mika Schneider, BSc (master student)
Dr Edoardo Galli (Postdoc)

PD Dr Matthias Mehling team:

Mali Coray (MD-PhD candidate)
Dr Varenka Epple (MD)
Annika Frentzel, BSc (master student)
Dr Jakob Fuhrmann (MD)
Dr Klara Ivanek (Postdoc)
Melanie Kaech, BSc (master student))

Prof Anne-Katrin Pröbstel team:

Prof Anne-Katrin Pröbstel team:
Tim Dürrenberger (doctoral student)
Dr Julia Flammer (resident/Postdoc)
Vidmante Fuchs (master student)
Ana Beatriz Gomes (PhD candidate)
Laila Kulsvehagen (PhD candidate)
Anne-Cathérine Lecourt (lab manager/technician)
Luc Lutz (master student)
Elisabeth Pössnecker (PhD candidate)
Laura Rieder (master student)
David Schreiner (bioinformatician)
Dr Lena Siewert (Postdoc)

7 | Publications in peer reviewed journals

1. Abboud H, Salazar-Camelo A, George N, Planchon SM, Matiello M, Mealy MA, Goodman A; **Pröbstel AK** (member of the Guthy-Jackson Foundation NMO International Clinical Consortium) On-behalf of the Guthy-Jackson Foundation NMO International Clinical Consortium. Symptomatic and restorative therapies in neuromyelitis optica spectrum disorders. J Neurol. 2021 Sep 5.

2. Allum JHJ, Rust HM, Lutz N, Schouenborg C, Fischer-Barnicol B, Haller V, **Derfuss T, Kuhle J, Yaldizli Ö**. Characteristics of improvements in balance control using vibro-tactile bio-feedback of trunk sway for multiple sclerosis patients. J Neurol Sci. 2021 Jun 15;425:117432.

3. **Barakovic M**, Tax CMW, Rudrapatna U, Chamberland M, Rafael-Patino J, **Granziera C**, Thiran JP, Daducci A, Canales-Rodríguez EJ, Jones DK. Resolving bundle-specific intra-axonal T2 values within a voxel using diffusion-relaxation tract-based estimation. Neuroimage. 2021 Feb 15;227:117617.

4. **Barakovic M**, Girard G, Schiavi S, Romascano D, Descoteaux M, **Granziera C**, Jones DK, Innocenti GM, Thiran JP, Daducci A. Bundle-Specific Axon Diameter Index as a New Contrast to Differentiate White Matter Tracts. Front Neurosci. 2021 Jun 15;15:646034.

5. Bavato F, Cathomas F, Klaus F, Gütter K, Barro C, **Maceski A**, Seifritz E, **Kuhle J**, Kaiser S, Quednow BB. Altered neuroaxonal integrity in schizophrenia and major depressive disorder assessed with neurofilament light chain in serum. J Psychiatr Res. 2021 Aug;140:141-148.

6. Benedict RHB, Tomic D, Cree BA, Fox R, Giovannoni G, Bar-Or A, Gold R, Vermersch P, Pohlmann H, Wright I, Karlsson G, Dahlke F, Wolf C, **Kappos L**. Siponimod and Cognition in Secondary Progressive Multiple Sclerosis: EXPAND Secondary Analyses. Neurology. 2021 Jan 19;96(3):e376-e386..

7. Bischof A, Jaeger VK, Hadden RDM, Luqmani RA, **Pröbstel AK**, Merkel PA, Suppiah R, Craven A, Collins MP, Daikeler T. Peripheral neuropathy in antineutrophil cytoplasmic antibody-associated vasculitides: Insights from the DCVAS study. Neurol Neuroimmunol Neuroinflamm. 2019 Sep 20;6(6):e615. Bjornevik K, O'Reilly EJ, Molsberry S, Kolonel LN, Le Marchand L, Paganoni S, Schwarzschild MA, **Benkert P, Kuhle J**, Ascherio A. Prediagnostic Neurofilament Light Chain Levels in Amyotrophic Lateral Sclerosis. Neurology. 2021 Aug 11;97(15):e1466–74.

8. Bosticardo S, Schiavi S, **Schaedelin S, Lu PJ, Barakovic M, Weigel M, Kappos L, Kuhle J**, Daducci A, Granziera C. Microstructure-Weighted Connectomics in Multiple Sclerosis. Brain Connect. 2021 Sep 8.

9. Butzkueven H, Trojano M, **Kappos L**, Spelman T, Wiendl H, Rosales K, Su R, Licata S, Ho PR, Campbell N. Clinical outcomes in patients who discontinue natalizumab therapy after 2 years in the Tysabri@Observational Program (TOP). Mult Scler. 2021 Mar;27(3):410-419.

10. Butzkueven H, Spelman T, Horakova D, Hughes S, Solaro C, Izquierdo G, Kubala Havrdová E, Grand'Maison F, Prat A, Girard M, Hupperts R, Onofrj M, Lugaresi A, Taylor B; MSBase Study Group, Giovannoni G, **Kappos L**, Hauser SL, Montalban X, Craveiro L, Freitas R, Model F, Overell J, Muros-Le Rouzic E, Sauter A, Wang Q, Wormser D, Wolinsky JS. Risk of requiring a wheelchair in primary progressive multiple sclerosis: Data from the ORATORIO trial and the MSBase registry. Eur J Neurol. 2021 Mar 16.

11. Butzkueven H, **Kappos L**, Spelman T, Trojano M, Wiendl H, Su R, Liao S, Hyde R, Licata S, Ho PR, Campbell N. No evidence for loss of natalizumab effectiveness with every-6-week dosing: a propensity score-matched comparison with every-4-week dosing in patients enrolled in the Tysabri Observational Program (TOP). Ther Adv Neurol Disord. 2021 Sep 27;14:17562864211042458.

12. Calabresi PA, **Kappos L**, Giovannoni G, Plavina T, Koulinska I, Edwards MR, Kieseier B, de Moor C, Sotirchos ES, Fisher E, Rudick RA, Sandrock A. Measuring treatment response to advance precision medicine for multiple sclerosis. Ann Clin Transl Neurol. 2021 Nov;8(11):2166-2173.

13. **Callegari I, Derfuss T, Galli E**. Update on treatment in multiple sclerosis. Presse Med. 2021 Jun;50(2):104068.

14. Centonze D, Rocca MA, Gasperini C, **Kappos L**, Hartung HP, Magyari M, Oreja-Guevara C, Trojano M, Wiendl H, Filippi M. Disease-modifying therapies and SARS-CoV-2 vaccination in multiple sclerosis: an expert consensus. J Neurol. 2021 Nov;268(11):3961-3968.

15. Chalkou K, Steyerberg E, Bossuyt P, **Subramaniam S, Benkert P, Kuhle J**, Disanto G, **Kappos L**, Zecca C, Egger M, Salanti G. Development, validation and clinical usefulness of a prognostic model for relapse in relapsing-remitting multiple sclerosis. Diagn Progn Res. 2021 Oct 27;5(1):17.

16. Chard DT, Alahmadi AAS, Audoin B, Charalambous T, Enzinger C, Hulst HE, Rocca MA, Rovira À, Sastre-Garriga J, Schoonheim MM, Tijms B, Tur C, Gandini Wheeler-Kingshott CAM, Wink AM, Ciccarelli O, Barkhof F; **Kappos L (member of MAGNIMS SC)**. Mind the gap: from neurons to networks to outcomes in multiple sclerosis. Nat Rev Neurol. 2021 Mar;17(3):173-184.

17. Chitnis T, Banwell B, **Kappos L**, Arnold DL, Gücüyener K, Deiva K, Skripchenko N, Cui LY, Saubadu S, Hu W, Benamor M, Le-Halpere A, Truffinet P, Tardieu M; TERIKIDS Investigators. Safety and efficacy of teriflunomide in paediatric multiple sclerosis (TERIKIDS): a multicentre, double-blind, phase 3, randomised, placebo-controlled trial. Lancet Neurol. 2021 Dec;20(12):1001-1011.

18. Collongues N, **Kuhle J, Tsagkas C**, Lamy J, Meyer N, Barro C, Parmar K, Amann M, Wuerfel J, **Kappos L**, Moreau T, de Seze J. Biomarkers of treatment response in patients with progressive multiple sclerosis treated with high-dose pharmaceutical-grade biotin (MD1003). Brain Behav. 2021 Feb;11(2):e01998.

19. Comabella M, Clarke MA, Schaedelin S, Tintoré M, Pareto D, Fissolo N, Pinteac R, **Granziera C**, Sastre-Garriga J, **Benkert P**, Auger C, **Kuhle J**, Montalban X, Rovira A. CSF chitinase 3-like 1 is associated with iron rims in patients with a first demyelinating event. Mult Scler. 2022 Jan;28(1):71-81.

20. Cree BA, Magnusson B, Rouyre N, Fox RJ, Giovannoni G, Vermersch P, Bar-Or A, Gold R, Piani Meier D, Karlsson G, Tomic D, Wolf C, Dahlke F, **Kappos L**. Siponimod: Disentangling disability and relapses in secondary progressive multiple sclerosis. Mult Scler. 2021 Sep;27(10):1564-1576.

21. Cree BA, Cohen JA, Reder AT, Tomic D, Silva D, Piani Meier D, Laflamme AK, Ritter S, **Leppert D, Kappos L**. Disability improvement as a clinically relevant outcome in clinical trials of relapsing forms of multiple sclerosis. Mult Scler. 2021 Dec;27(14):2219-2231.

22. **D’Souza M**, Papadopoulou A, Girardey C, **Kappos L**. Standardization and digitization of clinical data in multiple sclerosis. Nat Rev Neurol. 2021 Feb;17(2):119-125.

23. De Brouwer E, Becker T, Moreau Y, Havrdova EK, Trojano M, Eichau S, Ozakbas S, Onofrj M, Grammond P, **Kuhle J, Kappos L**, Sola P, Cartechini E, Lechner-Scott J, Alroughani R, Gerlach O, Kalincik T, Granella F, Grand'Maison F, Bergamaschi R, José Sá M, Van Wijmeersch B, Soysal A, Sanchez-Menoyo JL, Solaro C, Boz C, Iuliano G, Buzzard K, Aguera-Morales E, Terzi M, Trivio TC, Spitaleri D, Van Pesch V, Shaygannejad V, Moore F, Oreja-Guevara C, Maimone D, Gouider R, Csepány T, Ramo-Tello C, Peeters L. Longitudinal machine learning modeling of MS patient trajectories improves predictions of disability progression. Comput Methods Programs Biomed. 2021 Sep;208:106180.

24. De Brouwer E, Becker T, Moreau Y, Havrdova EK, Trojano M, Eichau S, Ozakbas S, Onofrj M, Grammond P, **Kuhle J, Kappos L**, Sola P, Cartechini E, Lechner-Scott J, Alroughani R, Gerlach O, Kalincik T, Granella F, Grand'Maison F, Bergamaschi R, Sá MJ, Van Wijmeersch B, Soysal A, Sanchez-Menoyo JL, Solaro C, Boz C, Iuliano G, Buzzard K, Aguera-Morales E, Terzi M, Trivio TC, Spitaleri D, Van Pesch V, Shaygannejad V, Moore F, Oreja-Guevara C, Maimone D, Gouider R, Csepány T, Ramo-Tello C, Peeters L. Corrigendum to Longitudinal machine learning modeling of MS patient trajectories improves predictions of disability progression: [Computer Methods and Programs in Biomedicine, Volume 208, (September 2021) 106180]. Comput Methods Programs Biomed. 2022 Jan;213:106479.

25. de Sitter A, Burggraaff J, Bartel F, Palotai M, Liu Y, Simoes J, Ruggieri S, Schregel K, Ropele S, Rocca MA, Gasperini C, Gallo A, Schoonheim MM, Amann M, Yiannakas M, Pareto D, Wattjes MP, Sastre-Garriga J, **Kappos L**, Filippi M, Enzinger C, Frederiksen J, Uitdehaag B, Guttmann CRG, Barkhof F, Vrenken H. Development and evaluation of a manual segmentation protocol for deep grey matter in multiple sclerosis: Towards accelerated semi-automated references. Neuroimage Clin. 2021;30:102659.

26. DeLuca J, Schippling S, Montalban X, **Kappos L**, Cree BAC, Comi G, Arnold DL, Hartung HP, Sheffield JK, Liu H, Silva D, Cohen JA. Effect of Ozanimod on Symbol Digit Modalities Test Performance in Relapsing MS. Mult Scler Relat Disord. 2021 Feb;48:102673. Disanto G, Ripellino P, Riccitelli GC, Sacco R, Scotti B, Fucili A, Pravata E, **Kuhle J**, Gobbi C, Zecca C. De-escalating rituximab dose results in stability of clinical, radiological, and serum neurofilament levels in multiple sclerosis. Mult Scler. 2021 Jul;27(8):1230-1239.

27. Dürr M, Nissen G, Sühs KW, Schwenkenbecher P, Geis C, Ringelstein M, Hartung HP, Fries MA, Kaufmann M, Malter MP, Madlener M, Thaler FS, Kämpfel T, Senel M, Häusler MG, Schneider H, Bergh FT, Kellinghaus C, Zettl UK, Wandinger KP, Melzer N, Gross CC, Lange P, Dreyhaupt J, Tumani H, Leipoldt F, Lewerenz J; **Pröbstel AK** (member of the German Network for Research on Autoimmune Encephalitis) German Network for Research on Autoimmune Encephalitis. CSF Findings in Acute NMDAR and LGI1 Antibody-Associated Autoimmune Encephalitis. Neurol Neuroimmunol Neuroinflamm. 2021 Oct 25;8(6):e1086.

28. Egle M, Loubiere L, Maceski A, **Kuhle J**, Peters N, Markus HS. Neurofilament light chain predicts future dementia risk in cerebral small vessel disease. J Neurol Neurosurg Psychiatry. 2021 Feb 8;92(6):582–9.

29. Fernández-Velasco JI, **Kuhle J**, Monreal E, Meca-Lallana V, Meca-Lallana J, Izquierdo G, Gascón-Giménez F, Sainz de la Maza S, Walo-Delgado PE, Maceski A, Rodríguez-Martin E, Roldán E, Villarrubia N, Saiz A, Blanco Y, Sánchez P, Carreón-Guarnizo E, Aladro Y, Brieva L, Íñiguez C, González-Suárez I, Rodríguez de Antonio LA, Masjuan J, Costa-Frossard L, Villar LM. Effect of Ocrelizumab in Blood Leukocytes of Patients With Primary Progressive MS. Neurol Neuroimmunol Neuroinflamm. 2021 Jan 6;8(2):e940.

30. Filippi M, Preziosa P, Barkhof F, Chard DT, De Stefano N, Fox RJ, Gasperini C, **Kappos L**, Montalban X, Moraal B, Reich DS, Rovira À, Toosy AT, Traboulsee A, Weinshenker BG, Zeydan B, Banwell BL, Rocca MA. Diagnosis of Progressive Multiple Sclerosis From the Imaging Perspective: A Review. JAMA Neurol. 2021 Mar 1;78(3):351-364.

31. Filippi M, Danesi R, **Derfuss T**, Duddy M, Gallo P, Gold R, Havrdová EK, Kornek B, Saccà F, Tintoré M, Weber J, Trojano M. Early and unrestricted access to high-efficacy disease-modifying therapies: a consensus to optimize benefits for people living with multiple sclerosis. J Neurol. 2021 Oct 9:1–8.

32. Filippi M, Preziosa P, Meani A, Dalla Costa G, Mesaros S, Drulovic J, Ivanovic J, Rovira A, Tintorè M, Montalban X, Ciccarelli O, Brownlee W, Miszkiel K, Enzinger C, Khalil M, Barkhof F, Strijbis EMM, Frederiksen JL, Cramer SP, Fainardi E, Amato MP, Gasperini C, Ruggieri S, Martinelli V, Comi G, Rocca MA; **Kappos L (member of MAGNIMS SC)**. Performance of the 2017 and 2010 Revised McDonald Criteria in Predicting MS Diagnosis After a Clinically Isolated Syndrome: A MAGNIMS Study. Neurology. 2022 Jan 4;98(1):e1-e14.

33. Fischer-Barnicol B, **Oechtering J, Kuhle J, Lorscheider J, Kappos L, Derfuss T**. Combination of teriflunomide and interferon as follow-up therapy after fingolimod-associated PML. Neurol Neuroimmunol Neuroinflamm. 2020 Dec 3;8(1):e927.

34. Fischi-Gomez E, Bonnier G, Ward N, **Granziera C**, Hadjikhani N. Ultrahigh field in vivo characterization of microstructural abnormalities in the orbitofrontal cortex and amygdala in autism. Eur J Neurosci. 2021 Sep;54(6):6229-6236.

35. Fisse AL, Pitarokoili K, **Leppert D**, Motte J, Pedreiturria X, **Kappos L**, Gold R, **Kuhle J**, Yoon MS. Serum neurofilament light chain as outcome marker for intensive care unit patients. J Neurol. 2021 Apr;268(4):1323-1329.

36. Fox RJ, Raska P, Barro C, Karafa M, König V, Bermel RA, Chase M, Coffey CS, Goodman AD, Klawiter EC, Naismith RT, **Kuhle J**. Neurofilament light chain in a phase 2 clinical trial of ibudilast in progressive multiple sclerosis. Mult Scler. 2021 Nov;27(13):2014-2022.

37. Francalancia J, Mavrogiorgou P, Juckel G, **Mitrovic T, Kuhle J, Naegelin Y, Kappos L**, Calabrese P. Death Anxiety and Attitudes towards Death in Patients with Multiple Sclerosis: An Exploratory Study. Brain Sci. 2021 Jul 22;11(8):964.

38. **Fuchs V**, Kutza M, Wischnewski S, Deigendesch N, **Lutz L, Kulsvehagen L**, Ricken G, **Kappos L**, Tzankov A, Hametner S, Frank S, Schirmer L, **Pröbstel AK**. Presence of SARS-CoV-2 Transcripts in the Choroid Plexus of MS and Non-MS Patients With COVID-19. Neurol Neuroimmunol Neuroinflamm. 2021 Jan 27;8(2):e957. Geis T, Brandstetter S, Toncheva AA,

Laub O, Leipold G, Wagner R, Kabesch M, Kasser S, **Kuhle J**, Wellmann S; CoKiBa Study group. Serum neurofilament light chain (sNfL) values in a large cross-sectional population of children with asymptomatic to moderate COVID-19. J Neurol. 2021 Nov;268(11):3969-3974.

39. Geys L, Parciak T, Pirmani A, McBurney R, Schmidt H, Malbaša T, Ziemssen T, Bergmann A, Rojas JI, Cristiano E, García-Merino JA, Fernández Ó, **Kuhle J**, Gobbi C, Delmas A, Simpson-Yap S, Nag N, Yamout B, Steinemann N, Seelndrayers P, Dubois B, van der Mei I, Stahmann A, Drulovic J, Pekmezovic T, Brola W, Tintore M, Kalkers N, Ivanov R, Zakaria M, Naseer MA, Van Hecke W, Grigoriadis N, Boziki M, Carra A, Pawlak MA, Dobson R, Hellwig K, Gallagher A, Leocani L, Dalla Costa G, de Carvalho Sousa NA, Van Wijmeersch B, Peeters LM. The Multiple Sclerosis Data Alliance Catalogue: Enabling Web-Based Discovery of Metadata from Real-World Multiple Sclerosis Data Sources. Int J MS Care. 2021 Nov-Dec;23(6):261-268.

40. Ghezzi A, Amato MP, Edan G, Hartung HP, Havrdová EK, **Kappos L**, Montalban X, Pozzilli C, Sorensen PS, Trojano M, Vermersch P, Comi G. The introduction of new edications in pediatric multiple sclerosis: Open issues and challenges. Mult Scler. 2021 Mar;27(3):479-482.

41. Gibiansky E, Petry C, Mercier F, Günther A, Herman A, **Kappos L**, Hauser S, Yamamoto Y, Wang Q, Model F, Kletzl H. Ocrelizumab in relapsing and primary progressive multiple sclerosis: Pharmacokinetic and pharmacodynamic analyses of OPERA I, OPERA II and ORATORIO. Br J Clin Pharmacol. 2021 Jun;87(6):2511-2520.

42. Giovannoni G, **Kappos L**, de Seze J, Hauser SL, Overell J, Koendgen H, Manfrini M, Wang Q, Wolinsky JS. Risk of requiring a walking aid after 6.5 years of ocrelizumab treatment in patients with relapsing multiple sclerosis: Data from the OPERA I and OPERA II trials. Eur J Neurol. 2021 Mar 16.

43. Goeral K, Hauck A, Atkinson A, Wagner MB, Pimpel B, Fuiko R, Klebermass-Schrehof K, Leppert D, **Kuhle J**, Berger A, Olischar M, Wellmann S. Early life serum neurofilament dynamics predict neurodevelopmental outcome of preterm infants. J Neurol. 2021 Jul;268(7):2570-2577.

44. Gold R, Arnold DL, Bar-Or A, Fox RJ, **Kappos L**, Mokliatchouk O, Jiang X, Lyons J, Kapadia S, Miller C. Long-term safety and efficacy of dimethyl fumarate for up to 13years in patients with relapsing-remitting multiple sclerosis: Final ENDORSE study results. Mult Scler. 2021 Sep 1:13524585211037909.

45. **Granziera C**, Wuerfel J, Barkhof F, Calabrese M, De Stefano N, Enzinger C, Evangelou N, Filippi M, Geurts JJG, Reich DS, Rocca MA, Ropele S, Rovira À, Sati P, Toosy AT, Vrenken H, Gandini Wheeler-Kingshott CAM, **Kappos L**; MAGNIMS Study Group. Quantitative magnetic resonance imaging towards clinical application in multiple sclerosis. Brain. 2021 Jun 22;144(5):1296-1311.

46. Guedel DS, Peters IJ, Banderet F, Eppe V, Egli S, **Mehling M**, Mayr M, Leeb A, Berger CT. Smartphone-based active vaccine safety surveillance (SmartVax) at a Swiss adult vaccination clinic - a pilot study. Swiss Med Wkly. 2021 Dec 14;151:w30090.

47. Harris S, Comi G, Cree BAC, Arnold DL, Steinman L, Sheffield JK, Southworth H, **Kappos L**, Cohen JA; Ozanimod Study Investigators. Plasma neurofilament light chain concentrations as a biomarker of clinical and radiologic outcomes in relapsing multiple sclerosis: Post hoc analysis of Phase 3 ozanimod trials. Eur J Neurol. 2021 Nov;28(11):3722-3730.

48. Hartung HP, **Derfuss T**, Cree BA, Sormani MP, Selmaj K, Stutters J, Prados F, MacManus D, Schneble HM, Lambert E, Porchet H, Glanzman R, Warne D, Curtin F, Kornmann G, Buf-fet B, Kremer D, Küry P, Leppert D, Rückle T, Barkhof F. Efficacy and safety of temelimab in multiple sclerosis: Results of a randomized phase 2b and extension study. *Mult Scler.* 2021 Jul 9;13524585211024997.

49. Hauser SL, **Kappos L**, Montalban X, Craveiro L, Chognot C, Hughes R, Koendgen H, Pasquarelli N, Pradhan A, Prajapati K, Wolinsky JS. Safety of Ocrelizumab in Patients With Relap-sing and Primary Progressive Multiple Sclerosis. *Neurology.* 2021 Oct 19;97(16):e1546-e1559.

50. Hunziker S, Quinto A, Ramin-Wright M, Becker C, Beck K, Vincent A, Tisljar K, Disanto G, **Benkert P, Leppert D**, Pargger H, Marsch S, Sutter R, Peters N, **Kuhle J**. Serum neurofila-ment measurement improves clinical risk scores for outcome prediction after cardiac arrest: results of a prospective study. *Crit Care.* 2021 Jan 20;25(1):32.

51. Ineichen BV, Moridi T, Ewing E, Ouellette R, Manouchehrinia A, Stawiarz L, Ferreira D, Muehlboeck SJ, **Kuhle J**, Westman E, Leppert D, Hillert J, Olsson T, Kockum I, Piehl F, Gran-berg T. Neurofilament light chain as a marker for cortical atrophy in multiple sclerosis without radiological signs of disease activity. *J Intern Med.* 2021 Aug;290(2):473-476.

52. **Johansson D**, Rauld C, Roux J, Regairaz C, **Galli E, Callegari I**, Raad L, Waldt A, Cuttat R, Roma G, **Diebold M**, Becher B, **Kuhle J, Derfuss T**, Carballido JM, Sanderson NSR. Mass Cytometry of CSF Identifies an MS-Associated B-cell Population. *Neurol Neuroimmu-nol Neuroinflamm.* 2021 Feb 15;8(2):e943.

53. **Kappos L**, Cohan S, Arnold DL, Robinson RR, Holman J, Fam S, Parks B, Xiao S, Castro-Borrero W. Safety and efficacy of daclizumab beta in patients with relapsing multiple sclero-sis in a 5-year open-label study (EXTEND): final results following early termination. *Ther Adv Neurol Disord.* 2021 Feb 26;14:1756286420987941.

54. **Kappos L**, Fox RJ, Burcklen M, Freedman MS, Havrdová EK, Hennessy B, Hohlfeld R, Lublin F, Montalban X, Pozzilli C, Scherz T, D'Ambrosio D, Linscheid P, Vaclavkova A, Pirozek-Lawniczek M, Kracker H, Sprenger T. Ponesimod Compared With Teriflunomide in Patients With Relapsing Multiple Sclerosis in the Active-Comparator Phase 3 OPTIMUM Study: A Randomized Clinical Trial. *JAMA Neurol.* 2021 May 1;78(5):558-567.

55. **Kappos L**. No consensus about consensus? *Neurol Res Pract.* 2021 Aug 6;3(1):46.

56. Keller CW, **Oechtering J**, Wiendl H, **Kappos L, Kuhle J**, Lünemann JD. Impact of com-plement activation on clinical outcomes in multiple sclerosis. *Ann Clin Transl Neurol.* 2021 Apr;8(4):944-950.

57. Kesenheimer EM, Wendebourg MJ, Weigel M, Weidensteiner C, Haas T, Richter L, San-der L, Horvath A, Barakovic M, Cattin P, **Granziera C**, Bieri O, Schlaeger R. Normalization of Spinal Cord Total Cross-Sectional and Gray Matter Areas as Quantified With Radially Sampled Averaged Magnetization Inversion Recovery Acquisitions. *Front Neurol.* 2021 Mar 25;12:637198.

58. Kim K, **Pröbstel AK**, Baumann R, Dyckow J, Landefeld J, Kogl E, Madireddy L, Lou-dermilk R, Eggers EL, Singh S, Caillier SJ, Hauser SL, Cree BAC; UCSF MS-EPIC Team, Schirmer L, Wilson MR, Baranzini SE. Cell type-specific transcriptomics identifies neddy-lation as a novel therapeutic target in multiple sclerosis.*Brain.* 2021 Mar 3;144(2):450-461.

59. Koini M, Pirpamer L, Hofer E, Buchmann A, Pinter D, Ropele S, Enzinger C, **Benkert P, Leppert D, Kuhle J**, Schmidt R, Khalil M. Factors influencing serum neurofilament light chain levels in normal aging. *Aging (Albany NY).* 2021 Dec 18;13(24):25729-25738.

60. **Kuhle J**, Daizadeh N, **Benkert P, Maceski A, Barro C, Michalak Z**, Sormani MP, Godin J, Shankara S, Samad TA, Jacobs A, Chung L, Rösch N, Kaiser C, Mitchell CP, Leppert D, Havari E, Kappos L. Sustained reduction of serum neurofilament light chain over 7 years by alemtuzumab in early relapsing-remitting MS. *Mult Scler.* 2021 Aug 11:13524585211032348.

61. La Rosa F, Yu T, Barquero G, Thiran JP, **Granziera C**, Bach Cuadra M. MP2RA-GE UNI translation via generative adversarial network improves the automatic tissue and le-sion segmentation in multiple sclerosis patients. *Comput Biol Med.* 2021 May;132:104297.

62. Langdon DW, Tomic D, Penner IK, Calabrese P, Cutter G, Häring DA, Dahlke F, **Kappos L**. Baseline characteristics and effects of fingolimod on cognitive performance in patients with relapsing-remitting multiple sclerosis. *Eur J Neurol.* 2021 Dec;28(12):4135-4145.

63. **Leppert D, Sutter R, Kuhle J**. Reply to: Neurofilament Light Chain in Patients with CO-VID-19 and Bacterial Pneumonia. *Ann Neurol.* 2021 Jul;90(1):175-176.

64. Liebmann M, Korn L, Janoschka C, Albrecht S, Lauks S, Herrmann AM, Schulte-Mecklen-beck A, Schwab N, Schneider-Hohendorf T, Eveslage M, Wildemann B, Luessi F, Schmidt S, Diebold M, Bittner S, Gross CC, Kovac S, Zipp F, **Derfuss T**, Kuhlmann T, König S, Meuth SG, Wiendl H, Klotz L. Dimethyl fumarate treatment restrains the antioxidative capacity of T cells to control autoimmunity. *Brain.* 2021 Nov 29;144(10):3126-3141.

65. Lin TY, Vitkova V, Asseyer S, Martorell Serra I, Motamedi S, Chien C, Ditzhaus M, Pa-padopoulou A, **Benkert P, Kuhle J**, Bellmann-Strobl J, Ruprecht K, Paul F, Brandt AU, Zim-mermann HG. Increased Serum Neurofilament Light and Thin Ganglion Cell-Inner Plexiform Layer Are Additive Risk Factors for Disease Activity in Early Multiple Sclerosis. *Neurol Neu-roimmunol Neuroinflamm.* 2021 Aug 4;8(5):e1051.

66. **Lorscheider J, Benkert P**, Lienert C, Hänni P, **Derfuss T, Kuhle J, Kappos L, Yaldizli Ö**. Comparative analysis of dimethyl fumarate and fingolimod in relapsing-remitting multiple sclerosis. *J Neurol.* 2021 Mar;268(3):941-949.

67. **Lorscheider J**. When does a heap become a heap? *Mult Scler.* 2021 Mar;27(3):329-330.

68. Louwsma J, Brunger AF, Bijzet J, Kroesen BJ, Roeloffzen WWH, Bischof A, **Kuhle J**, Drost G, Lange F, Kuks JBM, Gans ROB, Hazenberg BPC, Nienhuis HLA. Neurofilament light chain, a biomarker for polyneuropathy in systemic amyloidosis. *Amyloid.* 2021 Mar;28(1):50-55.

69. Lukas C, Bellenberg B, Prados F, Valsasina P, Parmar K, Brouwer I, Pareto D, Rovira À, Sastre-Garriga J, Gandini Wheeler-Kingshott CAM, **Kappos L**, Rocca MA, Filippi M, Yianna-kas M, Barkhof F, Vrenken H. Quantification of Cervical Cord Cross-Sectional Area: Which Acquisition, Vertebra Level, and Analysis Software? A Multicenter Repeatability Study on a Traveling Healthy Volunteer. *Front Neurol.* 2021 Aug 4;12:693333.

70. **Lu PJ**, Yoo Y, **Rahmanzadeh R, Galbusera R, Weigel M**, Ceccaldi P, Nguyen TD, Spin-cemaille P, Wang Y, Daducci A, La Rosa F, Bach Cuadra M, Sandkühler R, Nael K, Doshi A, Fayad ZA, **Kuhle J, Kappos L**, Odry B, Cattin P, Gibson E, **Granziera C**. GAMER MRI: Ga-ted-attention mechanism ranking of multi-contrast MRI in brain pathology. *Neuroimage Clin.* 2021;29:102522.

71. **Lu PJ, Barakovic M, Weigel M, Rahmanzadeh R, Galbusera R**, Schiavi S, Daducci A, La Rosa F, Bach Cuadra M, Sandkühler R, **Kuhle J, Kappos L**, Cattin P, **Granziera C**. GAMER-MRI in Multiple Sclerosis Identifies the Diffusion-Based Microstructural Measures That Are Most Sensitive to Focal Damage: A Deep-Learning-Based Analysis and Clinico-Biological Validation. *Front Neurosci.* 2021 Apr 6;15:647535.

72. Maggi P, **Kuhle J, Schädelin S**, van der Meer F, **Weigel M, Galbusera R**, Mathias A, **Lu PJ, Rahmanzadeh R, Benkert P**, La Rosa F, Bach Cuadra M, Sati P, Théaudin M, Pot C, van Pesch V, Leppert D, Stadelmann C, **Kappos L**, Du Pasquier R, Reich DS, Absinta M, **Granziera C**. Chronic White Matter Inflammation and Serum Neurofilament Levels in Multiple Sclerosis. *Neurology.* 2021 Aug 10;97(6):e543-e553.

73. Marignier R, Hacohen Y, Cobo-Calvo A, **Pröbstel AK**, Aktas O, Alexopoulos H, Amato MP, Asgari N, Banwell B, Bennett J, Brilot F, Capobianco M, Chitnis T, Ciccarelli O, Deiva K, De Sèze J, Fujihara K, Jacob A, Kim HJ, Kleiter I, Lassmann H, Leite MI, Linington C, Meinl E, Palace J, Paul F, Petzold A, Pittock S, Reindl M, Sato DK, Selmaj K, Siva A, Stankoff B, Tintore M, Traboulsee A, Waters P, Waubant E, Weinshenker B, **Derfuss T**, Vukusic S, Hemmer B. Myelin-oligodendrocyte glycoprotein antibody-associated disease. *Lancet Neurol.* 2021 Sep;20(9):762-772.

74. McComb M, Browne RW, Bhattacharya S, Bodziak ML, Jakimovski D, Weinstock-Guttman B, **Kuhle J**, Zivadinov R, Ramanathan M. The cholesterol autooxidation products, 7-keto-cholesterol and 7β-hydroxycholesterol are associated with serum neurofilaments in multiple sclerosis. *Mult Scler Relat Disord.* 2021 May;50:102864.

75. Meier AL, Bodmer NS, Wirth C, Bachmann LM, Ribi C, **Pröbstel AK**, Waeber D, Jelcic I, Steiner UC; Swiss SLE Cohort Study (SSCS). Neuro-psychiatric manifestations in patients with systemic lupus erythematosus: A systematic review and results from the Swiss lupus cohort study. *Lupus.* 2021 Sep;30(10):1565-1576.

76. Meinl E, Derfuss T, Krumbholz M, **Pröbstel AK**, Hohlfeld R. Humoral autoimmunity in multiple sclerosis. *J Neurol Sci.* 2011 Jul 15;306(1-2):180-2. Mimpfen M, Muris AH, Rolf L, Gerlach O, **Kuhle J**, Hupperts R, Smolders J, Damoiseaux J. Prognostic value of natural killer cell/T cell ratios for disease activity in multiple sclerosis. *Eur J Neurol.* 2021 Mar;28(3):901-909.

77. **Naegelin Y, Kuhle J, Schädelin S**, Datta AN, Magon S, Amann M, **Barro C**, Ramelli GP, Heesom K, Barde YA, Weber P, **Kappos L**. Fingolimod in children with Rett syndrome: the FINGORETT study. *Orphanet J Rare Dis.* 2021 Jan 6;16(1):19.

78. Niemann L, Lezius S, Maceski A, **Leppert D**, Englisch C, Schwedhelm E, Zeller T, Gerloff C, **Kuhle J**, Choe CU. Serum neurofilament is associated with motor function, cognitive decline and subclinical cardiac damage in advanced Parkinson's disease (MARK-PD). *Parkinsonism Relat Disord.* 2021 Sep;90:44-48.

79. **Oechtering J, Schaedelin S, Benkert P**, Müller S, Achtnichts L, Vehoff J, Disanto G, Findling O, Fischer-Barnicol B, **Orleth A**, Chan A, Pot C, **Barakovic M, Rahmanzadeh R, Galbusera R**, Heijnen I, Lalive PH, Wuerfel J, **Subramaniam S**, Aeschbacher S, Conen D, **Naegelin Y, Maceski A, Meier S**, Berger K, Wiendl H, Lincke T, Lieb J, **Yaldizli Ö**, Sinnecker T, **Derfuss T**, Regeniter A, Zecca C, Gobbi C, **Kappos L, Granziera C, Leppert D, Kuhle J**; Swiss Multiple Sclerosis Cohort Study. Intrathecal Immunoglobulin M Synthesis is an Independent Biomarker for Higher Disease Activity and Severity in Multiple Sclerosis. *Ann Neurol.* 2021 Sep;90(3):477-489.

80. Parmar K, Fonov VS, **Naegelin Y**, Amann M, Wuerfel J, Collins DL, Gaetano L, Magon S, Sprenger T, **Kappos L, Granziera C, Tsagkas C**. Regional Cerebellar Volume Loss Predicts Future Disability in Multiple Sclerosis Patients. *Cerebellum.* 2021 Aug 21.

81. Petzold A, Albrecht P, Balcer L, Bekkers E, Brandt AU, Calabresi PA, Deborah OG, Graves JS, Green A, Keane PA, Nij Bijvank JA, Sander JW, Paul F, Saidha S, Villoslada P, Wagner SK, Yeh EA, **Kappos L (member of INVISUAL)**; IMSVISUAL, ERN-EYE Consortium. Artificial intelligence extension of the OSCAR-IB criteria. *Ann Clin Transl Neurol.* 2021 Jul;8(7):1528-1542.

82. Portaccio E, Fonderico M, Hemmer B, **Derfuss T**, Stankoff B, Selmaj K, Tintorè M, Amato MP. Impact of COVID-19 on multiple sclerosis care and management: Results from the European Committee for Treatment and Research in Multiple Sclerosis survey. *Mult Scler.* 2022 Jan;28(1):132-138.

83. Probert F, Yeo T, Zhou Y, Sealey M, Arora S, Palace J, Claridge TDW, Hillenbrand R, **Oechtering J, Leppert D, Kuhle J**, Anthony DC. Integrative biochemical, proteomics and metabolomics cerebrospinal fluid biomarkers predict clinical conversion to multiple sclerosis. *Brain Commun.* 2021 Apr 19;3(2):fcab084.

84. **Pröbstel AK**, Schirmer L. SARS-CoV-2-specific neuropathology: fact or fiction? *Trends Neurosci.* 2021 Dec;44(12):933-935.

85. Puentes F, **Benkert P**, Amor S, **Kuhle J**, Giovannoni G. Antibodies to neurofilament light as potential biomarkers in multiple sclerosis. *BMJ Neurol Open.* 2021 Nov 11;3(2):e000192.

86. Raftopoulos R, **Kuhle J**, Grant D, Hickman SJ, Altmann DR, Leppert D, Blennow K, Zetterberg H, Kapoor R, Giovannoni G, Gnanapavan S. Neurofilament results for the phase II neuroprotection study of phenytoin in optic neuritis. *Eur J Neurol.* 2021 Feb;28(2):587-594.

87. **Rahmanzadeh R, Lu PJ, Barakovic M, Weigel M**, Maggi P, Nguyen TD, Schiavi S, Daducci A, La Rosa F, **Schaedelin S**, Absinta M, Reich DS, Sati P, Wang Y, Bach Cuadra M, Radue EW, **Kuhle J, Kappos L, Granziera C**. Myelin and axon pathology in multiple sclerosis assessed by myelin water and multi-shell diffusion imaging. *Brain.* 2021 Jul 28;144(6):1684-1696.

88. Ramanujam R, Zhu F, Fink K, Karrenbauer VD, **Lorscheider J, Benkert P**, Kingwell E, Tremlett H, Hillert J, Manouchehrinia A; BeAMS Study group. Accurate classification of secondary progression in multiple sclerosis using a decision tree. *Mult Scler.* 2021 Jul;27(8):1240-1249.

89. Rocca MA, Valsasina P, Meani A, Gobbi C, Zecca C, Rovira A, Sastre-Garriga J, Kearney H, Ciccarelli O, Matthews L, Palace J, Gallo A, Biseco A, Lukas C, Bellenberg B, Barkhof F, Vrenken H, Preziosa P, Filippi M; **KAPPOS L (part of MAGNIMS Study Group)**. Association of Gray Matter Atrophy Patterns With Clinical Phenotype and Progression in Multiple Sclerosis. *Neurology.* 2021 Mar 16;96(11):e1561-e1573.

90. Rodgers S, **Calabrese P**, Ajdacic-Gross V, Steinemann N, Kaufmann M, Salmen A, Manjaly ZM, Kesselring J, Kamm CP, **Kuhle J**, Chan A, Gobbi C, Zecca C, Müller S, von Wyl V. Major depressive disorder subtypes and depression symptoms in multiple sclerosis: What is different compared to the general population? *J Psychosom Res.* 2021 May;144:110402.

91. Rodgers S, Manjaly ZM, Calabrese P, Steinemann N, Kaufmann M, Salmen A, Chan A, Kesselring J, Kamm CP, **Kuhle J**, Zecca C, Gobbi C, von Wyl V, Ajdacic-Gross V. The Effect of Depression on Health-Related Quality of Life Is Mediated by Fatigue in Persons with Multiple Sclerosis. *Brain Sci.* 2021 Jun 5;11(6):751.

92. Rosso M, Healy BC, Saxena S, Paul A, Bjornevik K, **Kuhle J, Benkert P**, Leppert D, Guttmann C, Bakshi R, Weiner HL, Chitnis T. MRI Lesion State Modulates the Relationship Between Serum Neurofilament Light and Age in Multiple Sclerosis. *J Neuroimaging.* 2021 Mar;31(2):388-393.

93. Rübsamen N, **Maceski A, Leppert D, Benkert P, Kuhle J**, Wiendl H, Peters A, Karch A, Berger K. Serum neurofilament light and tau as prognostic markers for all-cause mortality in the elderly general population-an analysis from the MEMO study. *BMC Med.* 2021 Feb 15;19(1):38.

94. Saak A, Benkert P, Akgün K, **Willemse E, Kuhle J**, Ziemssen T, Jackson S, Schaefer J. Serum Neurofilament Light Chain: A Marker of Nervous System Damage in Myopathies. *Front Neurosci.* 2021 Dec 17;15:791670.

95. Sabatino JJ Jr, **Pröbstel AK**, Zamvil SS. Publisher Correction: B cells in autoimmune and neurodegenerative central nervous system diseases. *Nat Rev Neurosci.* 2020 Jan;21(1):56.

96. Sairanen V, Ocampo-Pineda M, **Granziera C**, Schiavi S, Daducci A. Incorporating outlier information into diffusion-weighted MRI modeling for robust microstructural imaging and structural brain connectivity analyses. *Neuroimage.* 2022 Feb 15;247:118802.

97. Saraste M, Bezukladova S, Matilainen M, Tuisku J, Rissanen E, Sucksdorff M, Laaksonen S, Vuorimaa A, **Kuhle J, Leppert D**, Airas L. High serum neurofilament associates with diffuse white matter damage in MS. *Neurol Neuroimmunol Neuroinflamm.* 2020 Dec 8;8(1):e926.

98. Saraste M, Bezukladova S, Matilainen M, Sucksdorff M, **Kuhle J, Leppert D**, Airas L. Increased serum glial fibrillary acidic protein associates with microstructural white matter damage in multiple sclerosis: GFAP and DTI. *Mult Scler Relat Disord.* 2021 May;50:102810.

99. Schilling KG, Rheault F, Petit L, Hansen CB, Nath V, Yeh FC, Girard G, **Barakovic M**, Rafael-Patino J, Yu T, Fisci-Gomez E, Pizzolato M, **Ocampo-Pineda M**, Schiavi S, Canales-Rodríguez EJ, Daducci A, **Granziera C**, Innocenti G, Thiran JP, Mancini L, Wastling S, Cocozza S, Petracca M, Pontillo G, Mancini M, Vos SB, Vakharia VN, Duncan JS, Melero H, Manzanedo L, Sanz-Morales E, Peña-Melián Á, Calamante F, Attyé A, Cabeen RP, Korobova L, Toga AW, Vijayakumari AA, Parker D, Verma R, Radwan A, Sunaert S, Emsell L, De Luca A, Leemans A, Bajada CJ, Haroon H, Azadbakht H, Chamberland M, Genc S, Tax CMW, Yeh PH, Srikanthana R, Mcknight CD, Yang JY, Chen J, Kelly CE, Yeh CH, Cochereau J, Maller JJ, Welton T, Almairac F, Seunarine KK, Clark CA, Zhang F, Makris N, Golby A, Rath Y, O'Donnell LJ, Xia Y, Aydogan DB, Shi Y, Fernandes FG, Raemaekers M, Warrington S, Michielse S, Ramírez-Manzanares A, Concha L, Aranda R, Meraz MR, Lerma-Usabiaga G, Roitman L, Fekonja LS, Calarco N, Joseph M, Nakua H, Voineskos AN, Karan P, Grenier G, Legarreta JH, Adluru N, Nair VA, Prabhakaran V, Alexander AL, Kamagata K, Saito Y, Uchida W, Andica C, Abe M, Bayrak RG, Wheeler-Kingshott CAMG, D'Angelo E, Palesi F, Savini G, Rolandi N, Guevara P, Houenou J, López-López N, Mangin JF, Poupon C, Román C, Vázquez A, Maffei C, Arantes M, Andrade JP, Silva SM, Calhoun VD, Caverzasi E, Sacco S, Lauricella M, Pestilli F, Bullock D, Zhan Y, Brignoni-Perez E, Lebel C, Reynolds JE, Nestrasil I, Labounek R, Lenglet C, Paulson A, Aulicka S, Heilbronner SR, Heuer K, Chandio BQ, Guaje J, Tang W, Garyfallidis

E, Raja R, Anderson AW, Landman BA, Descoteaux M. Tractography dissection variability: What happens when 42 groups dissect 14 white matter bundles on the same dataset? *Neuroimage.* 2021 Nov;243:118502.

100. Schindler P, Grittner U, **Oechtering J, Leppert D**, Siebert N, Duchow AS, Oertel FC, Asseyer S, Kuchling J, Zimmermann HG, Brandt AU, **Benkert P**, Reindl M, Jarius S, Paul F, Bellmann-Strobl J, **Kuhle J**, Ruprecht K. Serum GFAP and NfL as disease severity and prognostic biomarkers in patients with aquaporin-4 antibody-positive neuromyelitis optica spectrum disorder. *J Neuroinflammation.* 2021 May 1;18(1):105.

101. Selmaj KW, Cohen JA, Comi G, Bar-Or A, Arnold DL, Steinman L, Hartung HP, Montalban X, Havrdova EK, Cree BAC, Minton N, Sheffield JK, Ding N, **Kappos L**. Ozanimod in relapsing multiple sclerosis: Pooled safety results from the clinical development program. *Mult Scler Relat Disord.* 2021 Jun;51:102844.

102. Sormani MP, Freedman MS, Aldridge J, Marhardt K, **Kappos L**, De Stefano N. MAGNIMS score predicts long-term clinical disease activity-free status and confirmed disability progression in patients treated with subcutaneous interferon beta-1a. *Mult Scler Relat Disord.* 2021 Apr;49:102790.

103. Srpova B, Uher T, Hrnčiarova T, Barro C, Andelova M, Michalak Z, Vaneckova M, Krasensky J, Noskova L, Havrdova EK, **Kuhle J**, Horakova D. Serum neurofilament light chain reflects inflammation-driven neurodegeneration and predicts delayed brain volume loss in early stage of multiple sclerosis. *Mult Scler.* 2021 Jan;27(1):52-60.

104. Steinacker P, Feneberg E, Halbgebauer S, Witzel S, Verde F, Oeckl P, Van Damme P, Gaur N, Gray E, Grosskreutz J, Jardel CG, Kachanov M, **Kuhle J**, Lamari F, **Maceski A**, Del Mar Amador M, Mayer B, Morelli C, Petri S, Poesen K, Raaphorst J, Salachas F, Silani V, Turner MR, Verbeek MM, Volk AE, Weishaupt JH, Weydt P, Ludolph AC, Otto M. Chitotriosidase as biomarker for early stage amyotrophic lateral sclerosis: a multicenter study. *Amyotroph Lateral Scler Frontotemporal Degener.* 2021 May;22(3-4):276-286.

105. Sutter R, Hert L, De Marchis GM, Twerenbold R, **Kappos L, Naegelin Y**, Kuster GM, **Benkert P**, Jost J, **Maceski AM**, Rüegg S, Siegemund M, **Leppert D**, Tschudin- Sutter S, **Kuhle J**. Serum Neurofilament Light Chain Levels in the Intensive Care Unit: Comparison between Severely Ill Patients with and without Coronavirus Disease 2019. *Ann Neurol.* 2021 Mar;89(3):610-616.

106. Thaler FS, Zimmermann L, Kammermeier S, Strippel C, Ringelstein M, Kraft A, Sühs KW, Wickel J, Geis C, Markewitz R, Urbanek C, Sommer C, Doppler K, Penner L, Lewerenz J, Rößling R, Finke C, Prüss H, Melzer N, Wandinger KP, Leyboldt F, Kümpfel T, **Pröbstel AK** (member of GENERATE), German Network for Research on Autoimmune Encephalitis (GENERATE). Rituximab Treatment and Long-term Outcome of Patients With Autoimmune Encephalitis: Real-world Evidence From the GENERATE Registry. *Neurol Neuroimmunol Neuroinflamm.* 2021 Oct 1;8(6):e1088.

107. Tietz AK, Angstwurm K, Baumgartner T, Doppler K, Eisenhut K, Elisak M, Franke A, Golombeck KS, Handreka R, Kaufmann M, Kraemer M, Kraft A, Lewerenz J, Lieb W, Madlener M, Melzer N, Mojzisova H, Möller P, Pfefferkorn T, Prüss H, Rostásy K, Schnegelsberg M, Schröder I, Siebenbrodt K, Sühs KW, Wickel J, Wandinger KP, Leyboldt F, Kühlenbäumer G; **Pröbstel AK** (member of GENERATE), German Network for Research on Autoimmune Encephalitis (GENERATE). Genome-wide Association Study Identifies 2 New Loci Associated With Anti-NMDAR Encephalitis. *Neurol Neuroimmunol Neuroinflamm.* 2021 Sep 28;8(6):e1085.

108. **Tsagkas C**, Parmar K, Pezold S, Barro C, Chakravarty MM, Gaetano L, **Naegelin Y**, Amann M, Papadopoulou A, Wufer J, **Kappos L**, **Kuhle J**, Sprenger T, **Granziera C**, Magon S. Classification of multiple sclerosis based on patterns of CNS regional atrophy covariance. Hum Brain Mapp. 2021 Jun 1;42(8):2399-2415.

109. **Tsagkas C**, Schäfer S, Baumgartner A, Müller J, Sinnecker T, Mehling M, van der Stouwe JG, Felder S, Panagiotou A, Potlukova E, Winzeler B, Scholz MC, Timper K, Blackham K, Nickel CH, Christ-Crain M, Bingisser R, Lyrer P, **Kappos L**, **Yaldizli Ö**. Syndrome of inappropriate antidiuretic hormone secretion and hypothalamic hypocortisolism in neuro-myelitis optica. Lancet. 2021 Jun 5;397(10290):2194.

110. **Tsagkas C**, **Naegelin Y**, Amann M, Papadopoulou A, Barro C, Chakravarty MM, Gaetano L, Wufer J, **Kappos L**, **Kuhle J**, **Granziera C**, Sprenger T, Magon S, Parmar K. Central nervous system atrophy predicts future dynamics of disability progression in a real-world multiple sclerosis cohort. Eur J Neurol. 2021 Dec;28(12):4153-4166.

111. **Tsagkas C**, Wendebourg MJ, Mehling M, **Lorscheider J**, Lyrer P, Décard BF. Acute Polyradiculomyelitis With Spinal Cord Gray Matter Lesions: A Report of Two Cases. Front Neurol. 2021 Aug 19;12:721669.

112. Uher T, McComb M, Galkin S, Srpova B, Oechtering J, **Barro C**, Tyblova M, Bergsland N, Krasensky J, Dwyer M, Havrdova EK, Posova H, Vaneckova M, Zivadinov R, Horakova D, **Kuhle J**, Ramanathan M. Neurofilament levels are associated with blood-brain barrier integrity, lymphocyte extravasation, and risk factors following the first demyelinating event in multiple sclerosis. Mult Scler. 2021 Feb;27(2):220-231.

113. Uher T, Havrdova EK, Benkert P, Bergsland N, Krasensky J, Srpova B, Dwyer M, Tyblova M, Meier S, Vaneckova M, Horakova D, Zivadinov R, **Leppert D**, Kalincik T, **Kuhle J**. Measurement of neurofilaments improves stratification of future disease activity in early multiple sclerosis. Mult Scler. 2021 Nov;27(13):2001-2013.

114. Valsasina P, Gobbi C, Zecca C, Rovira A, Sastre-Garriga J, Kearney H, Yiannakas M, Matthews L, Palace J, Gallo A, Bisecco A, Gass A, Eisele P, Filippi M, Rocca MA; **Kappos L (part of MAGNIMS Study Group)**. Characterizing 1-year development of cervical cord atrophy across different MS phenotypes: A voxel-wise, multicentre analysis. Mult Scler. 2021 Oct 4;13524585211045545.

115. von Wyl V, **Benkert P**, Moser A, **Lorscheider J**, Décard B, Hänni P, Lienert C, **Kuhle J**, **Derfuss T**, **Kappos L**, **Yaldizli Ö**. Disability progression <i>in relapse-free</i> multiple sclerosis patients on fingolimod versus interferon-beta/glatiramer acetate. Mult Scler. 2021 Mar;27(3):439-448.

116. Vrenken H, Jenkinson M, Pham DL, Guttmann CRG, Pareto D, Paardekooper M, de Sitter A, Rocca MA, Wotschel V, Cardoso MJ, Barkhof F; **Kappos L (part of MAGNIMS Study Group)**. Opportunities for Understanding MS Mechanisms and Progression With MRI Using Large-Scale Data Sharing and Artificial Intelligence. Neurology. 2021 Nov 23;97(21):989-999.

117. Wattjes MP, Ciccarelli O, Reich DS, Banwell B, de Stefano N, Enzinger C, Fazekas F, Filippi M, Frederiksen J, Gasperini C, Hachohen Y, **Kappos L**, Li DKB, Mankad K, Montalban X, Newsome SD, Oh J, Palace J, Rocca MA, Sastre-Garriga J, Tintoré M, Traboulsee A, Vrenken H, Yousry T, Barkhof F, Rovira À; Magnetic Resonance Imaging in Multiple Sclerosis study group; Consortium of Multiple Sclerosis Centres; North American Imaging in Multiple

Sclerosis Cooperative MRI guidelines working group. 2021 MAGNIMS-CMSC-NAIMS consensus recommendations on the use of MRI in patients with multiple sclerosis. Lancet Neurol. 2021 Aug;20(8):653-670.

118. **Weigel M**, Dechent P, **Galbusera R**, Bahn E, Nair G, **Lu PJ**, **Kappos L**, Brück W, Stadelmann C, **Granziera C**. Imaging multiple sclerosis pathology at 160 µm isotropic resolution by human whole-brain ex vivo magnetic resonance imaging at 3 T. Sci Rep. 2021 Jul 29;11(1):15491.

119. Wiendl H, Spelman T, Butzkueven H, **Kappos L**, Trojano M, Su R, Campbell N, Ho PR, Licata S. Real-world disability improvement in patients with relapsing-remitting multiple sclerosis treated with natalizumab in the Tysabri Observational Program. Mult Scler. 2021 Apr;27(5):719-728.

120. Wiendl H, Gold R, Berger T, **Derfuss T**, Linker R, Mäurer M, Stangel M, Aktas O, Baum K, Berghoff M, Bittner S, Chan A, Czaplinski A, Deisenhammer F, Di Pauli F, Du Pasquier R, Enzinger C, Fertl E, Gass A, Gehring K, Gobbi C, Goebels N, Guger M, Haghighia A, Hartung HP, Heidenreich F, Hoffmann O, Hunter ZR, Kallmann B, Kleinschnitz C, Klotz L, Leussink V, Leutmezer F, Limmroth V, Lünemann JD, Lutterotti A, Meuth SG, Meyding-Lamadé U, Platten M, Rieckmann P, Schmidt S, Tumani H, Weber MS, Weber F, Zettl UK, Ziemssen T, Zipp F; die Multiple Sklerose Therapie Konsensus Gruppe (MSTKG). Multiple Sklerose Therapie Konsensus Gruppe (MSTKG): Positionspapier zur verlaufsmodifizierenden Therapie der Multip-len Sklerose 2021 (White Paper) [Multiple sclerosis treatment consensus group (MSTCG): position paper on disease-modifying treatment of multiple sclerosis 2021 (white paper)]. Nervenarzt. 2021 Aug;92(8):773-801. German.

121. Wiendl H, Gold R, Zipp F; **Derfuss T** (member of Multiple Sclerosis Therapy Consensus Group.), Multiple Sclerosis Therapy Consensus Group. Multiple sclerosis therapy consensus group (MSTCG): answers to the discussion questions. Neurol Res Pract. 2021 Aug 6;3(1):44.

122. Wiendl H, Gold R, Berger T, **Derfuss T**, Linker R, Mäurer M, Aktas O, Baum K, Berghoff M, Bittner S, Chan A, Czaplinski A, Deisenhammer F, Di Pauli F, Du Pasquier R, Enzinger C, Fertl E, Gass A, Gehring K, Gobbi C, Goebels N, Guger M, Haghighia A, Hartung HP, Heidenreich F, Hoffmann O, Kallmann B, Kleinschnitz C, Klotz L, Leussink V, Leutmezer F, Limmroth V, Lünemann JD, Lutterotti A, Meuth SG, Meyding-Lamadé U, Platten M, Rieckmann P, Schmidt S, Tumani H, Weber F, Weber MS, Zettl UK, Ziemssen T, Zipp F; ‘Multiple Sclerosis Therapy Consensus Group’ (MSTCG). Multiple Sclerosis Therapy Consensus Group (MSTCG): position statement on disease-modifying therapies for multiple sclerosis (white paper). Ther Adv Neurol Disord. 2021 Aug 18;14:17562864211039648.

123. Wilke C, Reich S, van Swieten JC, Borroni B, Sanchez-Valle R, Moreno F, Laforce R, Graff C, Galimberti D, Rowe JB, Masellis M, Tartaglia MC, Finger E, Vandenberghe R, de Mendonça A, Tagliavini F, Santana I, Ducharme S, Butler CR, Gerhard A, Levin J, Danek A, Otto M, Frisoni G, Ghidoni R, Sorbi S, Bocchetta M, Todd E, **Kuhle J**, **Barro C**; Genetic Frontotemporal dementia Initiative (GENFI), Rohrer JD, Synofzik M. Stratifying the Presymptomatic Phase of Genetic Frontotemporal Dementia by Serum NfL and pNfH: A Longitudinal Multicentre Study. Ann Neurol. 2022 Jan;91(1):33-47.

124. **Woelfle T**, **Pless S**, **Wiencierz A**, **Kappos L**, **Naegelin Y**, **Lorscheider J**. Practice Effects of Mobile Tests of Cognition, Dexterity, and Mobility on Patients With Multiple Sclerosis: Data Analysis of a Smartphone-Based Observational Study. J Med Internet Res. 2021 Nov 18;23(11):e30394.

125. Yeo T, Probert F, Sealey M, Saldana L, Gerald R, Höeckner S, Schiffer E, Claridge TDW, **Leppert D**, DeLuca G, **Kuhle J**, Palace J, Anthony DC. Objective biomarkers for clinical relapse in multiple sclerosis: a metabolomics approach. Brain Commun. 2021 Oct 12; 3(4):fcab240.

126. Yu T, Canales-Rodríguez EJ, Pizzolato M, Piredda GF, Hilbert T, Fisch-Gomez E, **Weigel M**, **Barakovic M**, Bach Cuadra M, **Granziera C**, Kober T, Thiran JP. Model-informed machine learning for multi-component T2< relaxometry. Med Image Anal. 2021 Apr;69:101940.

127. Zecca C, Disanto G, Sacco R, MacLachlan S, **Kuhle J**, Ramagopalan SV, Gobbi C. Increasing cancer risk over calendar year in people with multiple sclerosis: a case-control study. J Neurol. 2021 Mar;268(3):817-824.

128. Zhong M, van der Walt A, Stankovich J, Kalincik T, Buzzard K, Skibina O, Boz C, Hodgkinson S, Slee M, Lechner-Scott J, Macdonell R, Prevost J, **Kuhle J**, Laureys G, Van Hiffte L, Alroughani R, Kermod AG, Butler E, Barnett M, Eichau S, van Pesch V, Grammond P, McCombe P, Karabudak R, Duquette P, Girard M, Taylor B, Yeh W, Monif M, Gresle M, Butz-kueven H, Jokubaitis VG. Prediction of multiple sclerosis outcomes when switching to ocrelizumab. Mult Scler. 2021 Oct 8:13524585211049986.

129. Zuber P, Gaetano L, Griffa A, Huerbin M, Pedullà L, Bonzano L, Altermatt A, **Tsagkas C**, **Parmar K**, Hagmann P, Wuerfel J, **Kappos L**, Sprenger T, Sporns O, Magon S. Additive and interaction effects of working memory and motor sequence training on brain functional connectivity. Sci Rep. 2021 Nov 29;11(1):23089.

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