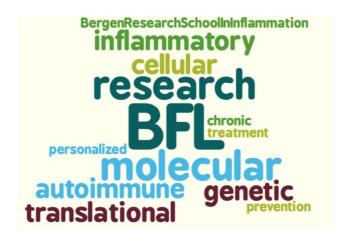
Annual Report 2022

Broegelmann Research Laboratory University of Bergen





Photos: Silke Appel, Anne Sidsel Herdlevaer, Karl A. Brokstad, Roland Jonsson, Kim E. Andreassen, Helena Erlandsson Harris, UiB. Stefan Bladh Front cover illustration by Kjerstin Jakobsen Editor: Helena Erlandsson Harris

Content

Director's comment

Vision and research areas

The Broegelmann foundation board

Our researchers

Principal investigators

Silke Appel

Yenan Bryceson

Helena Erlandsson Harris

Piotr Mydel

Marie Wahren-Herlenius

Graduations

External funding – major grants

Bergen Research School in Inflammation

Meetings, Lectures and Dissemination

International Publications

Social activities

Director's comment



Helena Erlandsson Harris, PhD. Head of Broegelmann Research Laboratory, 2022 "I am really excited about this opportunity to join the Broegelmann Research Laboratory and together with researchers there further develop BRL as a hub for research on inflammatory diseases. My goal is to ensure that the tradition of translational immunological research is continued and expanded by utilising the modern, state-of-the-art facilities available at UiB. Not only is this important in the quest to improve treatment and care of those affected by chronic inflammatory conditions, it is also the foundation for high quality doctoral education"

These were my hopes and visions when I started as the head of the Broegelmann Research Laboratory in January 2022. Today, a bit more than a year later, I am still excited. Being more familiar with the environment I do see a lot of opportunities to fulfill my goals and we have started to realize them

Thus 2022 has been a year when we have worked on strategies and visions, recruitments, initiating new projects and organisation. At the same time as moving established research projects forward successfully, securing external funding and pursuing activities within the Bergen Research school in Inflammation. 2022 was also the year when we could leave most of the digital meeting formats the pandemic compelled on us while retaining those that improve our interactions within the scientific community.

It is with pride that I look back on 2022 and how we at the Broegelmann Research Laboratory contributed with new science and knowledge, educated students and interacted with society.

With these introductory words we proudly present our annual report 2022.

Helena Erlandsson Harris

Helena Hann

Head of Broegelmann Research Laboratory

Annual report 2022 Broegelmann research laboratory

Vision and Research Areas

"To pave the way for personalized treatment and ultimately prevention of rheumatic diseases and their associated co-morbidities"

The researchers at the Broegelmann Research Laboratory have a mutual interest in translational research on chronic inflammatory and autoimmune diseases. The research performed spans from molecular and genetic studies, cellular studies and experimental models as well as studies on clinically well-characterized patient samples.

The mutual ambition is to understand the molecular mechanisms active in different inflammatory diseases and their subgroups, their underlying risk factors and triggers. This paves the way for better, personalized treatment and ultimately to prevention of disease.

Current major areas of research:

Translational studies in autoimmunity; autoantibodies and biomarkers

Immunopathogenesis of juvenile idiopathic arthritis

Functional genomics and genetic basis of the autoimmune exocrinopathy (Sjögren's syndrome)

Host microbe interactions (*P. gingivalis* – arthritis, Alzheimer)

Hyperinflammatory primary immunodeficiency syndromes

Inflammation in mental disorders

Cancer

The Board



The Broegelmann building at Strandgaten 5, downtown Bergen.

Photo: Helena Harris

The Broegelmann Foundation Board (Kjøbmand J.P. Broegelmanns Legat)



Attorney at Law

Bernt Jacob Pettersen

Chairman



Professor, MD, PhD Lars A. Akslen



Professor, MD, PhD Robert Bjerknes

The Broegelmann Research Laboratory was inaugurated in 1957, made possible by the legacy of merchant Johan Henrik Broegelmann.

JH Brogelmann had inherited a successful fabrics and finer ladies' clothing business from his father, Johan Petter Broegelmann. In his will, JH Broegelmann expressed his wish that his assets should form the basis for a foundation that in the name of his father should support humanitarian projects. He appointed his long-term co-worker Ingeborg Berg Nielsen as executer of his will. Ingeborg Berg Nielsen established the Merchant JP Broegelmann foundation with the purpose of supporting research on unexplored diseases performed at the University of Bergen; for example rheumatism, polio or cancer. Ingeborg Berg Nielsen further bequeathed her own assets to the foundation.

Thus, in 1957 the Broegelmann Research Laboratory could be inaugurated and has since then contributed the research field of immunology, including rheumatic diseases such as arthritis and Sjögren's syndrome. Even today, 65 years later, the support of the foundation forms the basis for immunology research at the University of Bergen.

Our Researchers

Head of Laboratory

Helena Erlandsson Harris, PhD (prof I) Head of Bergen research school of inflammation (inflammation/rheumatology)

Academic staff/senior researchers	Trainees
/postdoctoral fellows Silke Appel, Dr. rer. nat. (prof I) (cellular/molecular immunology)	(supervisors in parentheses) Noemi Dudzinska, PhD candidate (Mydel/Kaminska)
Yenan Bryceson, PhD (prof II) (cellular/molecular immunology)	Urszula Kalucka, PhD candidate (Mydel/Appel/Kaminska)
Richard Davies, PhD (post doc) (cellular/molecular immunology)	Dorentina Osmani, PhD candidate (Fossen/Appel/Kaminska)
Tim Holmes, PhD (post doc)	Harini Pechiappan, PhD candidate (Erlandsson Harris/Jonsson M/Aulin/Davies)
(cellular/molecular immunology)	Aleksandra Petrovic, PhD candidate (Appel/Jonsson R/Solberg)
Roland Jonsson, DMD, PhD (prof emeritus) (rheumatology/immunology)	Aina van der Meeren, PhD candidate (Berentsen Jacobsen/ Appel/ Lied)
Marta Kaminska, PhD (post doc) (biochemistry)	Ylva Bratterud Helgesen, Masters student (Appel/Bergum)
Piotr Mydel, MD, PhD (prof I) (biochemistry/immunology)	Alireza Molai, Masters student (Appel/Bergum/Borge)
,, 5,,	Technical and administrative staff
Marie Wahren-Herlenius, MD, PhD (prof II) (rheumatology/immunology)	Kate Froland, office manager
	Marianne Eidheim, laboratory manager
Janka Babickova (researcher)	Kjerstin Jakobsen, laboratory manager
(Biochemistry/Nephrology)	Richard Davies, senior lab manager



Kjerstin Jakobsen and Harini Pechiappan



Tim Holmes





Noemi Dudzinska



Urzula Kalucka and Dorentina Osmani



ADDITIONAL SCIENTISTS/ KEY COLLABORATORS AT UIB AFFILIATED WITH THE LABORATORY

Brith Bergum, PhD
Department of Clinical Science

Johan G. Brun, Professor, Department of Clinical Science

Birgitte Berentzen Jacobsen, Assoc Professor Department of Clinical Science Haukeland University Hospital

Rebecca Cox Brokstad, Professor Department of Clinical Science

Torgils Fossen, Professor Department of Chemistry

Daniel Hammenfors, MD, PhD Haukeland University Hospital

Stephanie Le Hellard, Professor, Department of Clinical Science

Erik Johnsen, Professor, Department of Clinical Medicine

Malin V. Jonsson, Professor, Department of Clinical Dentistry

Maria K. Jonsson, PhD, Haukeland University Hospital Karl-Henning Kalland, Professor, Department of Clinical Science

Rune Kroken, Assoc Professor, Department of Clinical Medicine

Hans Peter Marti, Professor Department of Clinical Medicine

Roald Omdal, Professor emeritus, Department of Clinical Science

Lene Frøyen Sandvik, Assoc Professor, Department of Clinical Medicine

Kathrine Skarstein, Professor, Department of Clinical Medicine

Jørn Skavland, PhD Department of Clinical Science

Silje Solberg, Assoc. Prof.
Department of Clinical Medicine

Principal Investigators

Silke Appel

Yenan Bryceson

Helena Erlandsson Harris

Piotr Mydel

Marie Wahren Herlenius

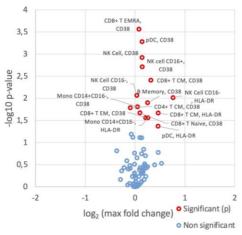
Silke Appel Professor I



Single cell network profiling of distinct immune cells allows analyzing pathway specific activity of patient samples at the single cell level. This will enable us to stratify different categories of patients and develop personalized therapies. The overall aim of our research is to further unravel the mechanisms by which dendritic cells modulate T cell responses in autoimmunity. Our results will lead to a better understanding of the cellular mechanisms involved in antigen presentation, T cell stimulation and tolerance induction, thereby revealing new tools for diagnosis and targets for therapy of patients with cancer, autoimmune and infectious diseases as well as patients undergoing stem cell transplantation. Thus, it will lead to an increased survival and better quality of life for patients.

About Silke Appel:

Head of Research, Dept Clinical Science, UiB Head of the core facility for flow cytometry, UiB During the past year, we used multi-color flow cytometry to show that regulatory T cells from patients with Sjögren's syndrome have an impaired response upon stimulation with IL-2 (Keindl et al., Arthritis Res Ther, 2022). In addition, mass cytometry analyses of patients with Sjögren's syndrome revealed differences in immune cell composition and signaling profiles between different patient subgroups (Sarkar et al., Front Immunol, 2022).



Sarkar et al., 2022

GROUP MEMBERS

Roland Jonsson, professor emeritus Alireza Molai, Master student Dorentina Osmani, MSc, PhD candidate Aleksandra Petrovic, MD, PhD candidate Aina van der Meeren, MSc, PhD candidate

Yenan Bryceson Professor II



Our laboratory studies the molecular regulation of cytotoxic lymphocyte function in the context of inflammatory disease as well as cancer. Defects in cytotoxic lymphocyte function are associated with often-fatal hyperinflammatory primary immunodeficiency syndromes in infants, but many such patients cannot be explained by current molecular insights. Furthermore, harnessing lymphocyte differentiation and function represent a promising avenue increasing the efficacy of cellular immunotherapy of cancer.



About Yenan Bryceson:

Yenan Bryceson did his undergraduate training at the University of Oslo and PhD at the National Institutes of Health, Rockville, MD, USA. He was appointed professor II at the University of Bergen in 2012. He is also a professor at Karolinska Institutet, Stockholm, Sweden.

Our laboratory studies the molecular regulation of cytotoxic lymphocyte function in inflammation. Cytotoxic lymphocytes are broadly categorized into cytotoxic CD8+ T cells and natural killer (NK) cells. Defects in cytotoxic lymphocyte function are associated with often-fatal hyperinflammatory primary immunodeficiency syndromes in infants. These hyperinflammatory syndromes are also associated with an increased risk of developing hematological malignancies.

Recently, adoptive transfer of NK cells or T cells engineered to express chimeric activating receptors (CARs) have shown efficacy in clinical trials against hematological malignancies, but improving their persistence and potential utility against solid tumors remains a major challenge.

We are working to decipher the molecular regulation of cytotoxic lymphocyte differentiation and function in blood and tissues using cutting-edge single cell techniques, including advanced flow cytometry and sorting combined with high-throughput sequencing. A particular focus of ours is on how specific transcription factors can be harnessed to manipulate NK cell differentiation and potentiate function for improved immunotherapy of cancer.

GROUP MEMBERS
Timothy Holmes, PhD (post doc)

Helena Erlandsson Harris Professor I



Juvenile idiopathic arthritis (JIA) is chronic arthritis affecting children. There is a great need for improved diagnostic and prognostics tools as well as new therapeutic options. In addition to joint inflammation, destruction of joint tissue and pain are hallmarks of the disease. In depth understanding of the molecular mechanisms driving these three hallmarks forms the basis for development of diagnostic/prognostic biomarker tests as well as development of new, specific therapy – precision medicine.

Alarmins are endogenous molecules released from stressed and dying cells with the function to initiate an inflammatory response. We study the alarmin HMGB1, how it can induce cell migration, cytokine production, cell differentiation and regeneration. All important features of chronic inflammation, including arthritis.

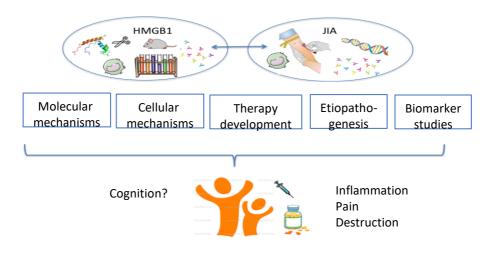


About Helena Erlandsson Harris:

Helena Erlandsson Harris did her undergraduate training at the University of Uppsala and PhD at the Karolinska Institutet, Sweden. She was appointed professor at Karolinska institutet in 2014. She became head of the Broegelmann Research Laboratory and Professor in Immunology at the University of Bergen in 2022.

Our projects are focused on expanding the molecular knowledge of the immune mechanisms active in JIA as a basis for biomarker and therapy development. To achieve this we analyse biosamples collected from children with JIA and compare generated data with information retrieved from national quality registers. In a recently started project we are investigating the possible connection of JIA, neuroinflammation and its potential influence on quality of life.

How HMGB1 is contributing to inflammation, pain and destruction is studied with a translational approach using molecular and cellular functional studies, analyses of HMGB1 in patient samples and model systems.



GROUP MEMBERS Richard Davies, Senior Lab manager Harini Pechiappan, PhD Candidate

Piotr Mydel Professor I



The group believes in interdisciplinary cooperation with medical and dental clinicians, epidemiologists, bio-scientists, industrial scientists to achieve our goals of elucidating the role of post-translational modifications in the aetiology of autoimmune diseases.



About Piotr Mydel: Piotr Mydel did his PhD education at the Jagiellonian University in Krakow, Poland and at the University of Boston, MA, USA followed by post doctoral work at Washington University, University of Gothenburg and University of Bergen. He was appointed Professor I at the University of Bergen in 2019. Piotr Mydel also holds a professorship at the Jagiellonian University.

Taking into account that up to 30% of the adult population worldwide suffers from severe periodontitis, the impact of this disease on human health is immense. Periodontitis (PD) is largely caused by infection, in which *Porphyromonas gingivalis* is a major pathogen, and is the most prevalent infectious inflammatory disease of mankind.

Mounting evidence suggests a causative link between PD and rheumatoid arthritis (RA), as well as periodontitis and cardiovascular disease. *P. gingivalis* is the only bacterium expressing the enzyme peptidylarginine deiminase (PAD) which converts arginine to citrulline, a process referred to as citrullination. Antibodies towards such citrullinated proteins, i.e ACPAs/anti-CCP, are of central importance in RA a chronic autoimmune disease which affect 0.5-1% worldwide. The presence of ACPAs or anti-CCP autoantibodies is not only highly specific for RA but the presence is also related to a more severe and destructive disease progression.

We hypothesise that anti-citrullinated protein antibodies can be generated, in genetically susceptible individuals, as a consequence of *P.gingivalis*-induced citrullination in the gingiva.

GROUP MEMBERS

Janka Babickova, PhD, Researcher Noemie Dudzińska, PhD candidate Urszula Zofia Kalucka, MSc, PhD candidate Marta Kaminska, PhD, postdoc Marie Wahren-Herlenius Professor II

The majority of autoimmune diseases are more common in women than in men, but the molecular basis for this sex-bias remains poorly understood. In our projects we focus on the autoimmune exocrinopathy Sjögren's syndrome which has among the highest observed female-to-male ratios, to dissect the genetic and hormonal contribution to sex-dependent immune regulation at single cell resolution and how these differences may lead to autoimmune disease.

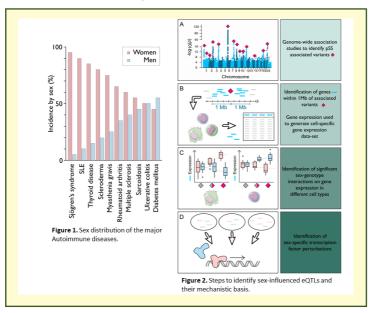


About Marie Wahren Herlenius:

Marie Wahren-Herlenius did part of her PhD training and a postdoctoral period at the Broegelmann Research Laboratory. She was appointed professor II, University of Bergen in 2020. She is also a professor at Karolinska Institutet, Stockholm, Sweden

Women are at much higher risk of developing autoimmune disease, with the most extreme numbers in systemic disorders such as SLE and Sjögren's syndrome for which more than nine out of ten patients are women. There is a clear genetic contribution to these diseases and genome-wide studies have identified polymorphisms associated with Sjögren's syndrome. Interestingly, many of the associated genetic variants lead to differential gene regulation. However, the influence of sex, or why these immune-pathways and related genes would become dysregulated specifically in women is not clear.

In our projects, we build on the observation that genetic polymorphisms associated with Sjögren's syndrome that we identify dramatically increase the likelihood for the disease to develop in women carrying these genetic traits compared to men. Consequently, the context "female sex" may lead to a different functional impact of the genetic polymorphisms associated with systemic autoimmunity than the context "male sex". Our projects aim to identify sex-influenced eQTLs, and dissect the genetic and hormonal contribution to sex-dependent immune regulation at single cell resolution and how these differences may lead to autoimmune disease.





Research Education

Bergen Research School in Inflammation (BRSI)



BRSI, coordinated by Helena Erlandsson Harris, is open to PhD candidates at UiB and encompasses activities such as courses, seminars, journal clubs – all with the aim to provide subject-specific knowledge to the participants.

Furthermore BRSI is an important platform for the interaction between PhD candiates and experienced scientists, to build networks, provide input on research projects and facilitate informal mentoring on research careers.

Courses given during 2022

- HUIMM320 Basic Immunology (5 ECTS)
- HUIMM307 Basic course in Flow Cytometry (5 ECTS)
- HUIMM901 Translational Immunology/project seminars (3 ECTS)
- HUIMM902 Journal club and watch (3 ECTS)
- HUIMM903 Human Immunobiology (10 ECTS)
- HUIMM 905 Advanced immunology course, precision medicine for inflammatory diseases (3 ECTS)
- HUIMM306A/906A Molecular and Cellular Methods in Immunology (8 ETCS)

HUIMM 905, advanced immunology, is given every third year on different topics. This year's topic was 'precision medicine for inflammatory diseases'. The course was held at Myrkdalen hotel over days and had 30 participants.



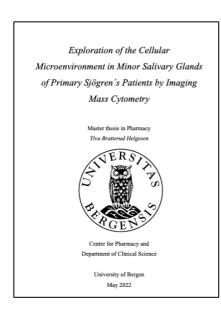
<u>Graduations</u>

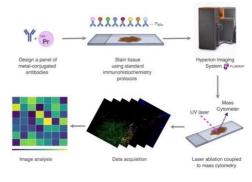
Master thesis:

Ylva Bratterud Helgesen- Master in Pharmacy

Exploration of the Cellular Microenvironment in Minor Salivary Glands of Primary Sjögren's Patients by Imaging Mass Cytometry

Supervisors: Silke Appel and Brith Begum





External funding- major grants

2019 - 2024

Piotr Mydel: National Institutes of Health (RO1 DE022597) "Bacterial peptidylarginine deiminase, a link between gums and joint disease", Total 6.4 mill NOK

2020-2023

Foundation for Research in Rheumatology (FOREUM) "Genetic variants associated with Sjögren's syndrome leading to differential gene expression in males and females and their functional impact on the immune system"; Karolinska Institutet v/Marie Wahren Herlenius as coordinator, UiB and Harvard as partners. Appel/Jonsson Total 1 500 000 NOK

2020 - 2023

Piotr Mydel: National Science Center Poland (2019/33/B/NZ4/01889) "Protein Carbamylation in Hemostatic Dysfunctions in Chronic Kidney Disease"; Total 505 000 €

2020 - 2023

Piotr Mydel: EU Joint Programme – Neurodegenerative Disease Research (EC/JPND; ES655895) "Alzheimer's disease as a comorbidity of chronic periodontitis with *Porphyromonas gingivalis* as a causative link between both diseases" Total 575 000 €

2019-2024

Roland Jonsson/Marie Wahren Herlenius: The European Commission Horizon 2020 contract NECESSITY (IMI2-JU/EU/H2020 nr. 806975) "New Clinical Endpoints in primary Sjögren's Syndrome: an Interventional Trial based on stratifying patients"

2021 - 2024

Piotr Mydel: EC/Grieg "Novel mechanisms of PAD activity regulation. Substrate specificity and activation of peptidyl arginine deiminases in the context of RA" Total 499 000 €

2021-2025

Yenan Bryceson: Norwegian Research Council "Engineering NK cells for improved functionality in immunotherapy" Total 12 mill NOK

2021-2025

Marie Wahren Herlenius: Norwegian Research Council "The molecular basis of sex differences in Sjögren's syndrome" Total 10 mill NOK

2022

Helena Erlandsson Harris: Norwegian Rheumatism Association "Molecular profiling of juvenile idiopathic arthritis, a necessary basis for implementation of precision medicine" Total 300 000 NOK

We are greatful for the financial support to our research projects from all our financiers











Meetings, Lectures and Dissemination

The 23nd Broegelmann Lecture



Broegelmann Research Laboratory



Department of Clinical Science Faculty of Medicine

invites you to

The 23rd Broegelmann Lecture

held by



Marie Wahren Herlenius, MD, PhD
Professor of Experimental Rheumatology
Division of Rheumatology
Karolinska Institutet, Stockholm

"Molecular basis for sexual dimorphism in autoimmune disease"

Monday, 30th May, 2022, at 14.00-15.00 Auditorium, Armauer Hansen building

Arranged by the research group Immunology and Rheumatology, NSI Bergen and the Bergen Research School in Inflammation http://www.uib.no/rg/broegelmann

This year's Broegelmann lecture was given by Professor Marie Wahren Herlenius, Karolinska Institutet and Broegelmann Research Laboartory.

Invited Lecturers







Prof. Gunnar Peiler Department of Medical Biochemistry and Microbiology Uppsala University

Mast cells - versatile effector cells in inflammatory conditions

12th October 2022 BBB, Auditorium 4, 10:15



About each Applicat research
that cells are testinately thought of as being the main cause of the
symptome associated with alleger centroline, including alleger asthma.
When mast cells become calculated they may record by degranisation,
which is associated with the release of a panel of inflammatory
including a second of the control of the control of the control of the control
of the control of the control of the control of the control
of the control of the control of the control of the control
of the control of the control of the control
of the control of the control of the control of the control
of the control of the control of the control of the control of the control
of the control o

Prof. Pejler will hold one-on-one meetings. Slots are limited to set up a meeting contact: marta.kaminska@uib.no









Prof. Vivianne Malmström Department of Medicine, Solna Karolinska Institute

Current understanding of T cells in rheumatoid arthritis (RA) - antigen specificities, repretoires, and effector functions

> 26th October 2022 BBB. Auditorium 4, 12:00



About prof. Milmatrion's research. These defended on these in 1997; 1 did my first postoci in Oxford These defended on these in 1997; 1 did my first postoci in Oxford These defended on the interest to the control of the control









Prof. Anna Blom

Cytoprotective roles of cytosolic forms of C3

9th November 2022 BBB, Auditorium 4, 12:00



About emif. Illiom's research
Arma Biom received Basic collision in biochemistry and inolocular
Arma Biom received Basic collisioning is chained; yellor all service
and the service of the collision of extracellular matter, After defending from Dhestin in Opposition of extracellular matter, After defending from Dhestin in Opposition of extracellular matter, After defending from Dhestin in Opposition of the collision of the collision of the complement of the Service of the Complement of the Complement of the Opposition of the Complement of the Opposition of Complement of Com

Prof. Blom will hold **one-on-one meetings**. Slots are limited! To set up a meeting contact: <u>marta.kaminska⊜uib.no</u>



K2 department conference Solstrand, 24-25 February

Broegelmann Research Laboratory was well represented at the bi-annual Department conference for K2, Dept clinical medicine 2, UiB.



Left to right: Marianne Eidsheim, Brith Bergum, Richard Davies, Silke Appel, Kjerstin Jacobsen

Hva kan vi lære av Karolinska institutt?



Kan vi laere noet av Karolinska Institutet? Can we learn something from Karolinska Institutet Karoling



Helena Erlandsson Harris gave a presentation discussing her experiences from KI and visions for BRL

Scandinavian Society for Immunology Reykjavik, Iceland. June 2022

Silke Appel, Helena Erlandsson Harris, Marta Kaminska & Dorentina Osmani participated with oral presentations and posters.



Dorentina Osmani and Silke Appel



Silke Appel

Sjögren's meeting Rome, Italy. June 2022





Roland Jonsson

'OPPLEV på Marineholmen'

a knowledge and science festival held as part of the 75th anniversary of the University of Bergen 23rd April



PhD candidate Dorentina Osmani presented her project on plant-derived substances as therapeutics against CovSars2 infection.

Annual meeting 2022



National NSI Speaker

Helena Erlandsson Harris, The Broegelmann
Chair in Translational Immunology
Broegelmann, Research Laboratory,
University of Bergen
Revealing immune mechanisms active in
juvenile arthritis by a proteomics approach

International Publications 2022

- Alim MA, Njenda D, Lundmark A, Kaminska M, Jansson L, Eriksson K, Kats A, Johannsen G, Arvidsson CK, Mydel PM, Yucel-Lindberg T. Pleckstrin Levels Are Increased in Patients with Chronic Periodontitis and Regulated via the MAP Kinase-p38α Signaling Pathway in Gingival Fibroblasts. Front Immunol. 2022 Jan 11;12:801096. doi: 10.3389/fimmu.2021.801096. eCollection 2021. IF 9.8
- Aulin C, Larsson S, Vogl T, Roth J, Åkesson A, Swärd P, Heinbäck R, Erlandsson Harris H, Struglics A. The alarmins high mobility group box protein 1 and \$100A8/A9 display different inflammatory profiles after acute knee injury. Osteoarthritis Cartilage. 2022 Sep;30(9):1198-1209. Epub 2022 Jul 6. PMID: 35809846. IF 6.6
- Binder V, Chruścicka-Smaga B, Bergum B, Jaisson S, Gillery P, Sivertsen J, Hervig T, Kaminska M, Tilvawala R, Nemmara VV, Thompson PR, Potempa J, Marti HP, Mydel P. Carbamylation of Integrin α IIb β 3: The Mechanistic Link to Platelet Dysfunction in ESKD. J Am Soc Nephrol. 2022 Aug 29;33(10):1841-56. doi: 10.1681 IF 10.1
- Brodin P, Casari G, Townsend L, O'Farrelly C, Tancevski I, Löffler-Ragg J, Mogensen TH, Casanova JL; COVID Human Genetic Effort. Studying severe long COVID to understand post-infectious disorders beyond COVID-19. Nat Med. 2022 May;28(5):879-882. doi: 10.1038/s41591-022-01766-7. IF 53.4
- Bastard P, Bizien L, Bucciol G, Lind Enoksson S, Jouanguy E, Karabela ŞN, Khan T, Kendir-Demirkol Y, Arias AA, Mansouri D, Marits P, Marr N, Migeotte I, Moens L, Ozcelik T, Pellier I, Sendel A, Şenoğlu, S, Shahrooei M, Smith CIE, Vandernoot I, Willekens K, Kart Yaşar K; COVID Human Genetic Effort, Bergman P, Abel L, Cobat A, Casanova JL, Meyts I, Bryceson YT. Respiratory viral infections in otherwise healthy humans with inherited IRF7 deficiency. J Exp Med. 2022 Jul 4;219(7):e20220202. doi: 10.1084/jem.20220202. Epub 2022 Jun 7. IF 14.3
- Dahlqvist J, Ekman D, Sennblad B, Kozyrev SV, Nordin J, Karlsson Å, Meadows JRS, Hellbacher E, Rantapää-Dahlqvist S, Berglin E, Stegmayr B, Baslund B, Palm Ø, Haukeland H, Gunnarsson I, Bruchfeld A, Segelmark M, Ohlsson S, Mohammad AJ, Svärd A, Pullerits R, Herlitz H, Söderbergh A, Rosengren Pielberg G, Hultin Rosenberg L, Bianchi M, Murén E, Omdal R, Jonsson R, Eloranta ML, Rönnblom L, Söderkvist P, Knight A, Eriksson P, Lindblad-Toh K. Identification and functional characterization of a novel susceptibility locus for small vessel vasculitis with MPO-ANCA. Rheumatology (Oxford). 2022 Aug 3;61(8):3461-3470. doi: 10.1093/rheumatology/keab912.PMID: 34888651 Free PMC article. IF 7.6

- Hernández-Molina G, Kostov B, Brito-Zerón P, Vissink A, Mandl T, Hinrichs AC, Quartuccio L, Baldini C, Seror R, Szántó A, Isenberg D, Gerli R, Nordmark G, Rasmussen A, Solans-Laque R, Hofauer B, Sène D, Pasoto SG, Rischmueller M, Praprotnik S, Gheita TA, Danda D, Armağan B, Suzuki Y, Valim V, Devauchelle-Pensec V, Retamozo S, Kvarnstrom M, Sebastian A, Atzeni F, Giacomelli R, Carsons SE, Kwok SK, Nakamura H, Fernandes Moça Trevisani V, Flores-Chávez A, Mariette X, Ramos-Casals M; Sjögren Big Data Consortium. Characterization and outcomes of 414 patients with primary SS who developed hematological malignancies. Rheumatology (Oxford). 2022 Apr 6:keac205. doi: 10.1093/rheumatology/keac205. Online ahead of print. *IF 7.0*
- Zetterberg H, Han J, Schlegel TT, Harris R, Harris HE, Eriksson LI. The Neuroimmune Response to Surgery - An Exploratory Study of Trauma-Induced Changes in Innate Immunity and Heart Rate Variability. Front Immunol. 2022 Jul 7;13:911744. eCollection 2022. PMID: 35874666. IF 7
- Jonsson R. Disease mechanisms in Sjögren's syndrome: What do we know? Scand J Immunol. 2022 Mar;95(3):e13145. doi: 10.1111/sji.13145. Epub 2022 Feb 1.PMID: 35073430 Review.
- Keindl M, Davies R, Bergum B, Brun JG, Hammenfors D, Jonsson R, Lyssenko V,
 Appel S. Impaired activation of STAT5 upon IL-2 stimulation in Tregs and elevated sIL-2R in Sjögren's syndrome. Arthritis Res Ther. 2022 May 7;24(1):101. doi: 10.1186/s13075-022-02769-y.PMID: 35526080 Free PMC article. IF 5.6
- Khatri B, Tessneer KL, Rasmussen A, Aghakhanian F, Reksten TR, Adler A, Alevizos I, Anaya JM, Aqrawi LA, Baecklund E, Brun JG, Bucher SM, Eloranta ML, Engelke F, Forsblad-d'Elia H, Glenn SB, Hammenfors D, Imgenberg-Kreuz J, Jensen JL, Johnsen SJA, Jonsson MV, Kvarnström M, Kelly JA, Li H, Mandl T, Martín J, Nocturne G, Norheim KB, Palm Ø, Skarstein K, Stolarczyk AM, Taylor KE, Teruel M, Theander E, Venuturupalli S, Wallace DJ, Grundahl KM, Hefner KS, Radfar L, Lewis DM, Stone DU, Kaufman CE, Brennan MT, Guthridge JM, James JA, Scofield RH, Gaffney PM, Criswell LA, Jonsson R, Eriksson P, Bowman SJ, Omdal R, Rönnblom L, Warner B, Rischmueller M, Witte T, Farris AD, Mariette X, Alarcon-Riquelme ME; PRECISESADS Clinical Consortium, Shiboski CH; Sjögren's International Collaborative Clinical Alliance (SICCA), Wahren-Herlenius M, Ng WF; UK Primary Sjögren's Syndrome Registry, Sivils KL, Adrianto I, Nordmark G, Lessard CJ. Genome-wide association study identifies Sjögren's risk loci with functional implications in immune and glandular cells. Nat Commun. 2022 Jul 27;13(1):4287. doi: 10.1038/s41467-022-30773-y. IF 17.7

- Kokkinou E, Pandey RV, Mazzurana L, Gutierrez-Perez I, Tibbitt CA, Weigel W, Soini T, Carrasco A, Rao A, Nagasawa M, Bal SM, Jangard M, Friberg D, Lindforss U, Nordenvall C, Ljunggren M, Haapaniemi S, Keita ÅV, Söderholm J, Hedin C, Spits H, Bryceson YT, Mjösberg J. CD45RA+CD62L-ILCs in human tissues represent a quiescent local reservoir for the generation of differentiated ILCs. Sci Immunol. 2022 Apr 15;7(70):eabj8301. doi: 10.1126/sci immunol.abj8301. Epub 2022 Apr 15. JF 17.7
- Lindahl H, **Bryceson YT**. Neuroinflammation Associated With Inborn Errors of Immunity. Front Immunol. 2022 Jan 19;12:827815. doi: 10.3389/fimmu.2021.827815. eCollection 2021. *IF 9.8*
- Lundtoft C, Pucholt P, Martin M, Bianchi M, Lundström E, Eloranta ML, Sandling JK, Sjöwall C, Jönsen A, Gunnarsson I, Rantapää-Dahlqvist S, Bengtsson AA, Leonard D, Baecklund E, Jonsson R, Hammenfors D, Forsblad-d'Elia H, Eriksson P, Mandl T, Magnusson Bucher S, Norheim KB, Auglaend Johnsen SJ, Omdal R, Kvarnström M, Wahren-Herlenius M, Notarnicola A, Andersson H, Molberg Ø, Diederichsen LP, Almlöf J, Syvänen AC, Kozyrev SV, Lindblad-Toh K; DISSECT Consortium; ImmunoArray Development Consortium, Nilsson B, Blom AM, Lundberg IE, Nordmark G, Diaz-Gallo LM, Svenungsson E, Rönnblom L. Complement C4 Copy Number Variation is Linked to SSA/Ro and SSB/La Autoantibodies in Systemic Inflammatory Autoimmune Diseases. Arthritis Rheumatol. 2022 Aug;74(8):1440-1450. doi: 10.1002/art.42122. Epub 2022 Jun 27. IF 15.2
- Lundtoft C, Sjöwall C, Rantapää-Dahlqvist S, Bengtsson AA, Jönsen A, Pucholt P, Wu YL, Lundström E, Eloranta ML, Gunnarsson I, Baecklund E, Jonsson R, Hammenfors D, Forsblad-d'Elia H, Eriksson P, Mandl T, Bucher S, Norheim KB, Johnsen SJA, Omdal R, Kvarnström M, Wahren-Herlenius M, Truedsson L, Nilsson B, Kozyrev SV, Bianchi M, Lindblad-Toh K; DISSECT consortium; ImmunoArray consortium, Yu CY, Nordmark G, Sandling JK, Svenungsson E, Leonard D, Rönnblom L. Combined genetic deficiencies of the classical complement pathway are strongly associated with both systemic lupus erythematosus and primary Sjögren's syndrome. Arthritis Rheumatol. 2022 Jun 21. doi: 10.1002/art.42270. IF 15.2

- Malicki S, Pucelik B, Żyła E, Benedyk-Machaczka M, Gałan W, Golda A, Sochaj-Gregorczyk A, Kamińska M, Encarnação JC, Chruścicka B, Marti HP, Chen TJ, Magiera-Mularz K, Zięba B, Holak TA, Dąbrowski JM, Czarna A, Kozieł J, Mydel P, Dubin G. Imaging of Clear Cell Renal Carcinoma with Immune Checkpoint Targeting Aptamer-Based Probe. Pharmaceuticals (Basel). 2022 Jun 1;15(6):697. doi: 10.3390/ph15060697. IF 4.9
- Matuozzo D, Talouarn E, Marchal A, Manry J, Seeleuthner Y, Zhang Y, Bolze A, Chaldebas M, Milisavljevic B, Zhang P, Gervais A, Bastard P, Asano T, Bizien L, Barzaghi F, Abolhassani H, Tayoun AA, Aiuti A, Darazam IA, Allende LM, Alonso-Arias R, Arias AA, Aytekin G, Bergman P, Bondesan S, Bryceson YT, Bustos IG, Cabrera-Marante O, Carcel S, Carrera P, Casari G, Chaïbi K, Colobran R, Condino-Neto A, Covill LE, El Zein L, Flores C, Gregersen PK, Gut M, Haerynck F, Halwani R, Hancerli S, Hammarström L, Hatipoğlu N, Karbuz A, Keles S, Kyheng C, Leon-Lopez R, Franco JL, Mansouri D, Martinez-Picado J, Akcan OM, Migeotte I, Morange PE, Morelle G, Martin-Nalda A, Novelli G, Novelli A, Ozcelik T, Palabiyik F, Pan-Hammarström Q, Pérez de Diego R, Planas-Serra L, Pleguezuelo DE, Prando C, Pujol A, Reyes LF, Rivière JG, Rodriguez-Gallego C, Rojas J, Rovere-Querini P, Schlüter A, Shahrooei M, Sobh A, Soler-Palacin P, Tandjaoui-Lambiotte Y, Tipu I, Tresoldi C, Troya J, van de Beek D, Zatz M, Zawadzki P, Al-Muhsen SZ, Baris-Feldman H, Butte MJ, Constantinescu SN, Cooper MA, Dalgard CL, Fellay J, Heath JR, Lau YL, Lifton RP, Maniatis T, Mogensen TH, von Bernuth H, Lermine A, Vidaud M, Boland A, Deleuze JF, Nussbaum R, Kahn-Kirby A, Mentre F, Tubiana S, Gorochov G, Tubach F, Hausfater P; COVID Human Genetic Effort; COVIDeF Study Group; French COVID Cohort Study Group; CoV-Contact Cohort; COVID-STORM Clinicians; COVID Clinicians; Orchestra Working Group; Amsterdam UMC Covid-19 Biobank; NIAID-USUHS COVID Study Group, Meyts I, Zhang SY, Puel A, Notarangelo LD, Boisson-Dupuis S, Su HC, Boisson B, Jouanguy E, Casanova JL, Zhang Q, Abel L, Cobat A. Rare predicted loss-of-function variants of type I IFN immunity genes are associated with lifethreatening COVID-19. medRxiv. 2022 Oct 25:2022.10.22.22281221. doi: 10.1101/2022.10.22.22281221. Preprint.

- Meisgen S, Hedlund M, Ambrosi A, Folkersen L, Ottosson V, Forsberg D, Thorlacius GE, Biavati L, Strandberg L, Mofors J, Ramskold D, Ruhrmann S, Meneghel L, Nyberg W, Espinosa A, Hamilton RM, Franco-Cereceda A, Hamsten A, Olsson T, Greene L, Eriksson P, Gemzell-Danielsson K, Salomonsson S, Kuchroo VK, Herlenius E, Kockum I, Sonesson SE, Wahren-Herlenius M. Auxilin is a novel susceptibility gene for congenital heart block which directly impacts fetal heart function. Ann Rheum Dis. 2022 Aug;81(8):1151-1161. doi: 10.1136/annrheumdis-2021-221714. Epub 2022 Apr 25. IF 19.1
- Midtbø H, Kringeland E, Gerdts E, Ueland PM, Meyer K, Linde A, Ulvik A, Jonsson R, Tveit KS. Biomarkers of inflammation and left ventricular remodelling in psoriasis patients treated with infliximab. Int J Immunopathol Pharmacol. 2022 Jan-Dec;36:3946320221111131. doi: 10.1177/03946320221111131. PMID: 35968808 Free PMC article. IF 3.2
- Muleviciene A, Sekine T, Zondag T, Bryceson YT, Tesi B, Rascon J. Childhood Kaposi sarcoma related to hypomorphic severe combined immunodeficiency caused by a novel CORO1A mutation. Pediatr Blood Cancer. 2022 May;69(5):e29487. doi: 10.1002/pbc.29487. Epub 2021 Dec 16. IF 3.8
- Pezoulas VC, Goules A, Kalatzis F, Chatzis L, Kourou KD, Venetsanopoulou A, Exarchos TP, Gandolfo S, Votis K, Zampeli E, Burmeister J, May T, Marcelino Pérez M, Lishchuk I, Chondrogiannis T, Andronikou V, Varvarigou T, Filipovic N, Tsiknakis M, Baldini C, Bombardieri M, Bootsma H, Bowman SJ, Soyfoo MS, Parisis D, Delporte C, Devauchelle-Pensec V, Pers JO, Dörner T, Bartoloni E, Gerli R, Giacomelli R, Jonsson R, Ng WF, Priori R, Ramos-Casals M, Sivils K, Skopouli F, Torsten W, A G van Roon J, Xavier M, De Vita S, Tzioufas AG, Fotiadis DI. Addressing the clinical unmet needs in primary Sjögren's Syndrome through the sharing, harmonization and federated analysis of 21 European cohorts. Comput Struct Biotechnol J. 2022 Jan 7;20:471-484. doi: 10.1016/j.csbj.2022.01.002. eCollection 2022.PMID: 35070169 Free PMC article. *IF 7.2*

- Qu H, Heinbäck R, Salo H, Ewing E, Espinosa A, Aulin C, Erlandsson Harris H. Transcriptomic Profiling Reveals That HMGB1 Induces Macrophage Polarization Different from Classical M1. Biomolecules. 2022 Jun 2;12(6):779. PMID: 35740904. *IF 5.9*
- Sarkar I, Davies R, Aarebrot AK, Solberg SM, Petrovic A, Joshi AM, Bergum B, Brun JG, Hammenfors D, **Jonsson R, Appel S.** Aberrant signaling of immune cells in Sjögren's syndrome patient subgroups upon interferon stimulation. Front Immunol. 2022 Aug 22;13:854183. doi: 10.3389/fimmu.2022.854183. eCollection 2022.PMID: 36072585 Free PMC article. *JF 9.8*
- Seror R, Baron G, Camus M, Cornec D, Perrodeau E, Bowman SJ, Bombardieri M, Bootsma H, Gottenberg JE, Fisher B, Hueber W, van Roon JA, Devauchelle-Pensec V, Gergely P, Mariette X, Porcher R; NECESSITY WP5 STAR development working group; Development and preliminary validation of the Sjögren's Tool for Assessing Response (STAR): a consensual composite score for assessing treatment effect in primary Sjögren's syndrome. Ann Rheum Dis. 2022 Jul;81(7):979-989. doi: 10.1136/annrheumdis-2021-222054. Epub 2022 Apr 7. IF 19.1
- Smith CIE, Zain R, Österborg A, Palma M, Buggert M, Bergman P, **Bryceson Y**. Do reduced numbers of plasmacytoid dendritic cells contribute to the aggressive clinical course of COVID-19 in chronic lymphocytic leukaemia? Scand J Immunol. 2022 Apr;95(4):e13153. doi: 10.1111/sji.13153. Epub 2022 Mar 16. *IF 3.3*
- Thorlacius GE, Björk A and **Wahren-Herlenius M.** Genetics and epigenetics of Sjögren's syndrome: implications for future therapies. Nat Rev Rheumatol. 2023 May;19(5):288-306. doi: 10.1038/s41584-023-00932-6. Epub 2023 Mar 13.PMID: 36914790. *IF 32.3*
- Trutschel D, Bost P, Mariette X, Bondet V, Llibre A, Posseme C, Charbit B, Thorball CW, Jonsson R, Lessard CJ, Felten R, Ng WF, Chatenoud L, Dumortier H, Sibilia J, Fellay J, Brokstad KA, Appel S, Tarn JR, Murci LQ, Mingueneau M, Meyer N, Duffy D, Schwikowski B, Gottenberg JE; Milieu Interieur, ASSESS investigators, and NECESSITY consortium. Variability in primary Sjögren's syndrome is driven by interferon alpha, and genetically associated with the class II HLA DQ locus. Arthritis Rheumatol. 2022 Jun 20. doi: 10.1002/art.42265. IF 15.2

- Zhang Q, Bastard P; COVID Human Genetic Effort, Cobat A, Casanova JL. Human genetic and immunological determinants of critical COVID-19 pneumonia. Nature. 2022 Mar;603(7902):587-598. doi: 10.1038/s41586-022-04447-0. Epub 2022 Jan 28. *IF 69.5*
- Zhang Q, Matuozzo D, Le Pen J, Lee D, Moens L, Asano T, Bohlen J, Liu Z, Moncada-Velez M, Kendir-Demirkol Y, Jing H, Bizien L, Marchal A, Abolhassani H, Delafontaine S, Bucciol G; COVID Human Genetic Effort, Bayhan GI, Keles S, Kiykim A, Hancerli S, Haerynck F, Florkin B, Hatipoglu N, Ozcelik T, Morelle G, Zatz M, Ng LFP, Lye DC, Young BE, Leo YS, Dalgard CL, Lifton RP, Renia L, Meyts I, Jouanguy E, Hammarström L, Pan-Hammarström Q, Boisson B, Bastard P, Su HC, Boisson-Dupuis S, Abel L, Rice CM, Zhang SY, Cobat A, Casanova JL. Recessive inborn errors of type I IFN immunity in children with COVID-19 pneumonia. J Exp Med. 2022 Aug 1;219(8):e20220131. doi: 10.1084/jem.20220131. Epub 2022 Jun 16. IF 14.3
- Zhang Q, Pizzorno A, Miorin L, Bastard P, Gervais A, Le Voyer T, Bizien L, Manry J, Rosain J, Philippot Q, Goavec K, Padey B, Cupic A, Laurent E, Saker K, Vanker M, Särekannu K; COVID Human Genetic Effort; Etablissement Français du Sang Study Group; Constances Cohort; 3C-Dijon Study; Cerba HealthCare Group; Lyon Antigrippe Working Group; REIPI INF Working Group, García-Salum T, Ferres M, Le Corre N, Sánchez-Céspedes J. Balsera-Manzanero M. Carratala J. Retamar-Gentil P, Abelenda-Alonso G, Valiente A, Tiberghien P, Zins M, Debette S, Meyts I, Haerynck F, Castagnoli R, Notarangelo LD, Gonzalez-Granado LI, Dominguez-Pinilla N, Andreakos E, Triantafyllia V, Rodríguez-Gallego C, Solé-Violán J, Ruiz-Hernandez JJ, Rodríguez de Castro F, Ferreres J, Briones M, Wauters J, Vanderbeke L, Feys S, Kuo CY, Lei WT, Ku CL, Tal G, Etzioni A, Hanna S, Fournet T, Casalegno JS, Queromes G, Argaud L, Javouhey E, Rosa-Calatrava M, Cordero E, Aydillo T, Medina RA, Kisand K, Puel A, Jouanguy E, Abel L, Cobat A, Trouillet-Assant S, García-Sastre A, Casanova JL. Autoantibodies against type I IFNs in patients with critical influenza pneumonia. J Exp Med. 2022 Nov 7;219(11): e20220514. doi: 10.1084/jem.20220514. Epub 2022 Sep 16. IF 14.3

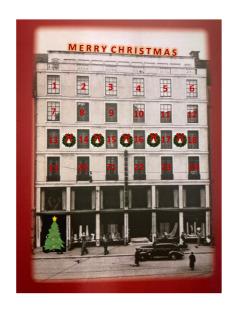
- Zhao, ZB; Marschner, JA; Iwakura, T; Li, C; Motrapu, M; Kuang, M; Popper, B; Linkermann, A; Klocke, J; Enghard, P; Muto, Y; Humphreys, BD; Harris, HE; Romagnani, P; Anders, HJ. Tubular Epithelial Cell HMGB1 Promotes AKI-CKD Transition by Sensitizing Cycling Tubular Cells to Oxidative Stress: A Rationale for Targeting HMGB1 during AKI Recovery. J Am Soc Nephrol. Online Dec 2022. 2023 Mar 1;34(3):394-411. PMID: 36857499. IF 15.0
- Zondag TCE, Torralba-Raga L, Van Laar JAM, Hermans MAW, Bouman A, Hollink IHIM, Van Hagen PM, Briggs DA, Hume AN, Bryceson YT. Novel RAB27A Variant Associated with Late-Onset Hemophagocytic Lymphohistiocytosis Alters Effector Protein Binding. J Clin Immunol. 2022 Nov;42(8):1685-1695. doi: 10.1007/s10875-022-01315-4. Epub 2022 Jul 23 IF 7.4

Social activities

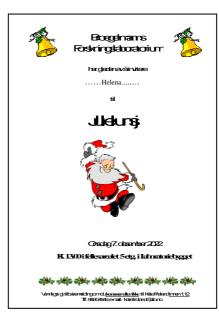




Summer dinner



The BRL advent calendar



Christmas lunch

Farewell and thank you celebration for our office manager Kate Frøland



The Broegelmann board expressed their gratitude for the excellent work Kate has provided for many years



Many colleagues wanted to say farewell to Kate and wish her happy retirement





Professor emeritus Roland Jonsson

Department head Pål Njølstad and Kate Frøland



Social Media

More about Broegelmann Research Laboratory can be found on:

https://www.uib.no/en/rg/broegelmann/105088/broegelmann-research-laboratory



Broegelmann research laboratory (BRL)



Broegelmann research laboratory (BRL)





uib.no

Broegelmann Research Laboratory
Department of Clinical Science / University of Bergen
The Laboratory Building, 5th floor / Haukeland University Hospital Jonas
Lies vei 87 / N-5021 Bergen, Norway post@uib.no / 55 58 00 00

uib.no/rg/broegelmann