

# ANNUAL REPORT 2021



**Josep Carreras**<sup>®</sup>  
LEUKAEMIA  
Research Institute



**ANNUAL  
REPORT**  
2021



**Josep Carreras**<sup>®</sup>  
LEUKAEMIA  
Research Institute

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## FOREWORD

Dr. Manel Esteller  
Director

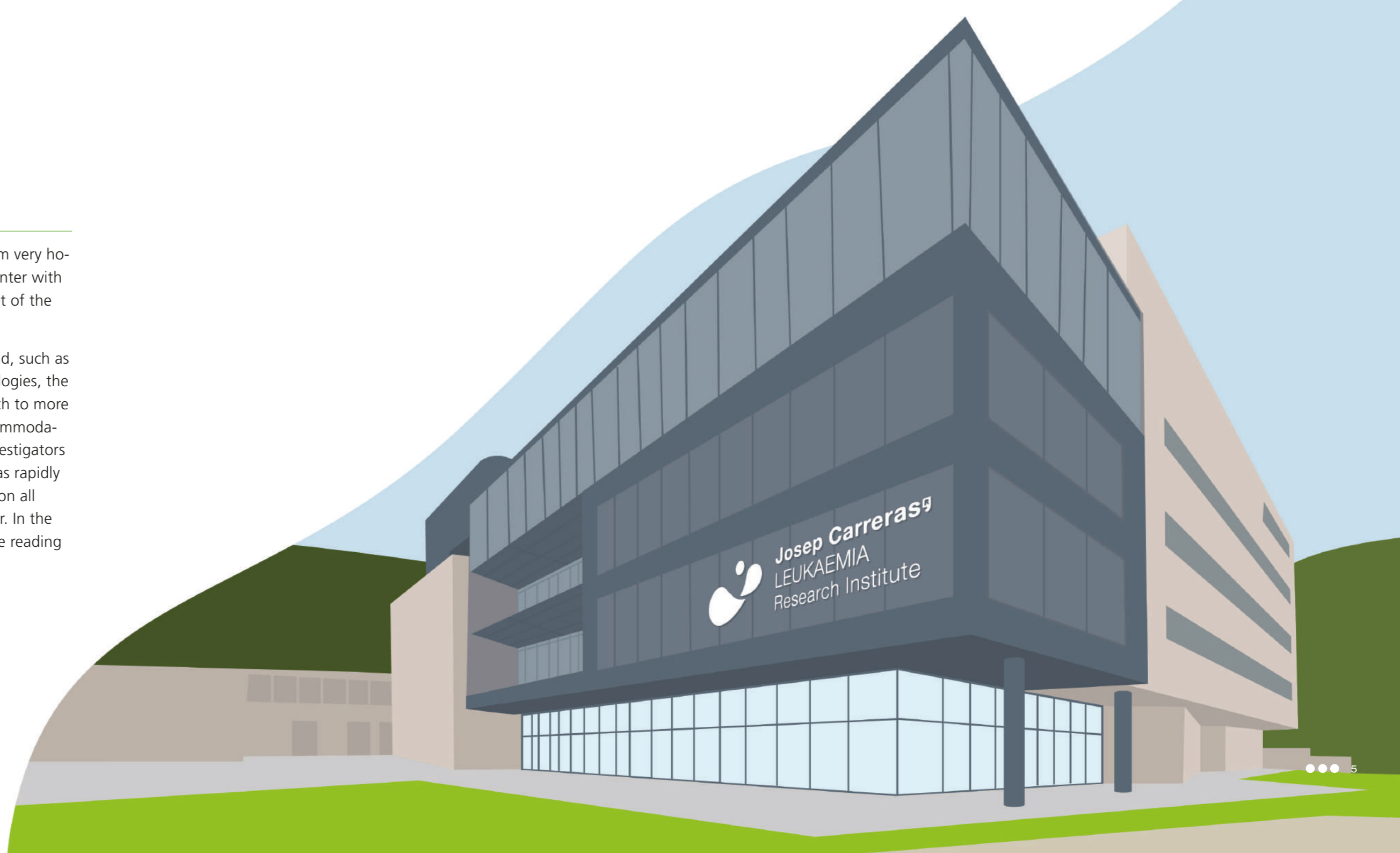
2021 has been a great year for the Institute. Entering the second and half year as Director of the center, I feel proud of all our personnel. In the fight against terrible diseases, such as some forms of leukemia, lymphoma and other blood disorders, the effort of all is required. This endeavor is possible because modern research involves not only devoted scientists, but also hard-working technicians and management staff.

Our Institute has experienced an excellent growth thanks to all the discoveries made, which are translated into many and highly relevant publications, but also into a significant increase in the competitive funding received

and greater outreach to society. I am very honored to serve as Director of the center with all of you as the most precious asset of the institution.

We will have many challenges ahead, such as the implementation of new technologies, the trend to move from wet-lab research to more bioinformatic approaches, the accommodation of newly recruited talented investigators and the goal to bring our findings as rapidly as possible to the patients. I count on all of you in the combat against cancer. In the meantime, take a rest and enjoy the reading of this annual report.

With gratitude,





## ABOUT US WHO WE ARE

The **Josep Carreras Leukaemia Research Institute** is a non-profit research institute based in Badalona (Barcelona), **dedicated to biomedical research and personalized medicine in leukemia and other malignant blood diseases.** It conducts excellent research into the basic, epidemiological, preventive, clinical and translational aspects of leukemia and other hematopoiesis malignancies.

The Josep Carreras Leukaemia Research Institute, **directed by Dr. Manel Esteller, was launched in 2010 by the Josep Carreras Foundation,** together with the Catalan government, **and is the first European research center devoted exclusively to leukemia and other malignant blood diseases.**

Research efforts are imperative to provide **patients with high-quality healthcare,** and our specialists combine their extensive knowledge to focus on patients' needs. **Our**

**aim is to understand the origins and development of leukemia and other malignant hematopoiesis pathologies with a view to their prevention, and our efforts are directed towards identifying new therapeutic targets and developing effective treatments** with fewer side effects through cutting-edge research.

Given our efficient governance, the Josep Carreras Leukaemia Research Institute is a Catalan Research Center of Excellence and forms part of the Catalan government's current research center network. It is also accredited by the Spanish Ministry of Health as a Health Research Center of Excellence and by the Spanish Association Against Cancer. The Institute also holds the HR Excellence in Research Award from the European Commission.

The ultimate goal of our interdisciplinary **team is to ensure that leukemia is a curable disease in all cases,** and we won't stop until we have achieved this.

The Josep Carreras Institute is a collaborative hub for basic and translational researchers who work together on the fundamental biological and clinical aspects of leukemia at our state-of-the-art facilities, which provide an excellent work environment and serve as a magnet for outstanding researchers from all over the world.

**It is home to 35 research groups and an increasing number of associated clinicians from five independent, coordinated scientific campuses: Hospital Clínic-UB Campus, Sant Pau Campus, Can Ruti Campus, Mar Campus and the Trueta Campus.** Our laboratories on those clinical campuses allow us to collaborate closely with clinicians from the five associated hospitals: Hospital Clínic, Hospital de Sant Pau, Hospital Germans Trias i Pujol, Hospital del Mar and Dr. Josep Trueta Hospital.



# MISSION, VISION AND VALUES

## MISSION

The Josep Carreras Leukemia Research Institute's mission is to carry out research into the epidemiological, preventive, clinical, translational and basic aspects of leukemia and other malignant blood diseases through innovation, with the ultimate aim of finding a cure.

## VISION

The vision of the Josep Carreras Leukaemia Research Institute is to be a world-class reference and excellent research multicampus that contributes to the improvement of results, and the cure of patients affected by leukemia and other malignant hemopathies, through innovation, sustainability, social responsibility, talent, and professional experience.

## VALUES

- Altruism
- Patient orientation
- Professional commitment
- Coexistence
- Respect
- Campus - Foundation alignment
- Scientific Leadership
- Collaboration
- Research - Assistance Integration
- Perseverance
- Continuous Improvement
- Methodological, scientific, and technological Innovation
- Environmental sustainability
- Austerity
- Transparency
- Social commitment.

## ABOUT US GOVERNING BODIES

The highest governing body is the Board of Trustees, which is represented by the Josep Carreras Foundation, the Catalan government's Ministry of Business and Knowledge, the Catalan government's Ministry of Health, the Autonomous University of Barcelona (UAB), the University of Barcelona (UB), Badalona Town Council, the Directorate General for Health Research and Planning, the Catalan Institute of Oncology, the Northern Metropolitan Territorial Area Administration, the Catalan Foundation for Research and Innovation, Hospital Clínic / the August Pi i Sunyer Biomedical Research Institute (IDIBAPS), the UB Hospital Coordination Committee and the Research Centres of Catalonia Institution Foundation (iCERCA).



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Ombudsman at the Josep Carreras Institute

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Dean for Research and Technology at the Universitat Autònoma de Barcelona (UAB)

**Mr. Robert Fabregat i Fuentes**  
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Hospital de la Santa Creu de Sant Pau

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Hospital Clínic de Barcelona

**Dr. Josep M<sup>a</sup> Campistol Plana**  
Hospital Clínic de Barcelona

**Mr. Josep Lluís Lafarga Traver**  
Lawyer



## SCIENTIFIC ADVISORY BOARD

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Head of Division of Cancer Epigenomics in German Cancer Research Center (DKFZ)

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Pathology Department of the NYU School of Medicine

**Prof. Pura Muñoz-Cánoves**  
ICREA Research Professor and Cell Biology Professor in the Department of Experimental and Health Sciences at the UPF

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Internal Scientific Committee Coordinator

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Campus Coordinator

**Dr. Armando López Guillermo**

**Dr. Joan Bladé Creixentí**

**Dr. Francisco Cervantes Requena**

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**Dr. Juan Manuel Sancho Cía**

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**Dr. Javier Briones Mejide**  
Campus Coordinator

**Dr. Josep Nomdedeu Guinot**

**Dr. Joan Carles Souto Andrés**

**Dr. Ramon Mangués Bafalluy**

**Dr. Carol Moreno Anastasio**





## ABOUT US

# DIRECTOR

**Dr. Manel Esteller** is Chairman of Genetics at the University of Barcelona's Faculty of Medicine and an ICREA Research Professor. Since May 2019, Dr. Esteller has been the Director of the Josep Carreras Leukaemia Research Institute. Dr. Esteller is considered to be among the top 0.1% of world scientists based on impact by Stanford University (METRICS). He is also a member of numerous international scientific societies and his work has been recognized by numerous awards, including the World Health Summit Award (2010), the Rey Jaime I Research Award (2013), the National Award in Oncology (2014), the Dr. Josep Trueta Medal from the Catalan government (2015), the National Research Award from the Catalan government (2015), the Gold Medal from the Parliament of Catalonia (2016), the International Award of Catalonia (2016), the Innovation in Healthcare Oncology Award (2018), the Narcís Monturiol Medal from the Catalan government (2020) and the Fernández-Cruz Award for excellence in biomedical research (2021).



Prof. Manel Esteller  
**Director**



Prof. Evarist Feliu  
**President of the Delegate Committee**



Dr. Josep Maria Ribera  
**Clinical Research Deputy Director**



Dr. Albert Oriol  
**Applied Research Director**



Dr. Jordi Esteve Reiner  
**UB Clinical Campus Coordinator**



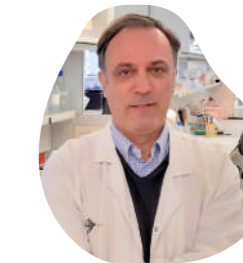
Mrs. Ana Garrido Anglada  
**Strategy Director and Acting Managing Director**



Dr. Anna Bigas  
**Basic Research Deputy Director**



Dr. Rafael Marcos  
**Epidemiological Research Director**

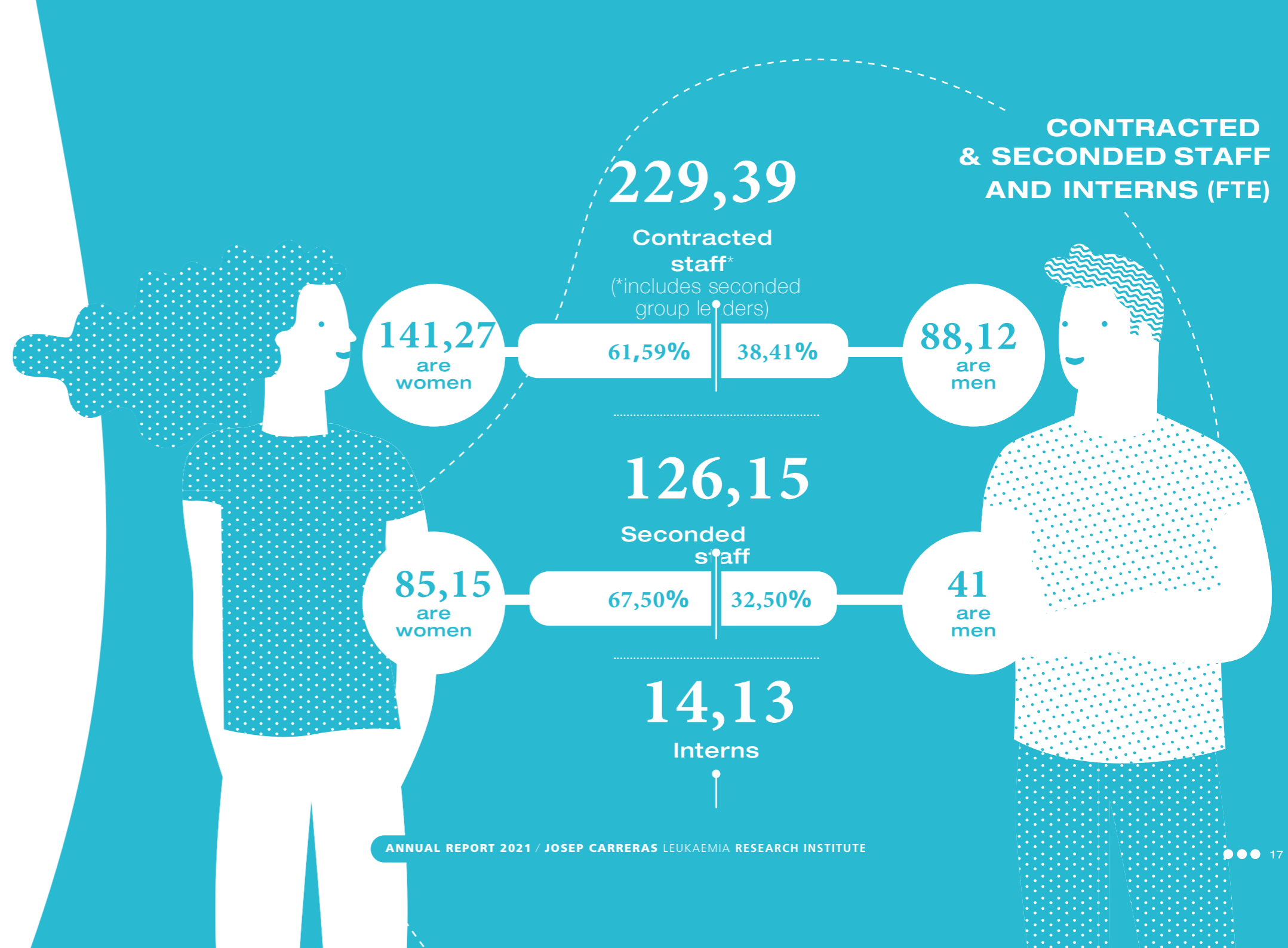
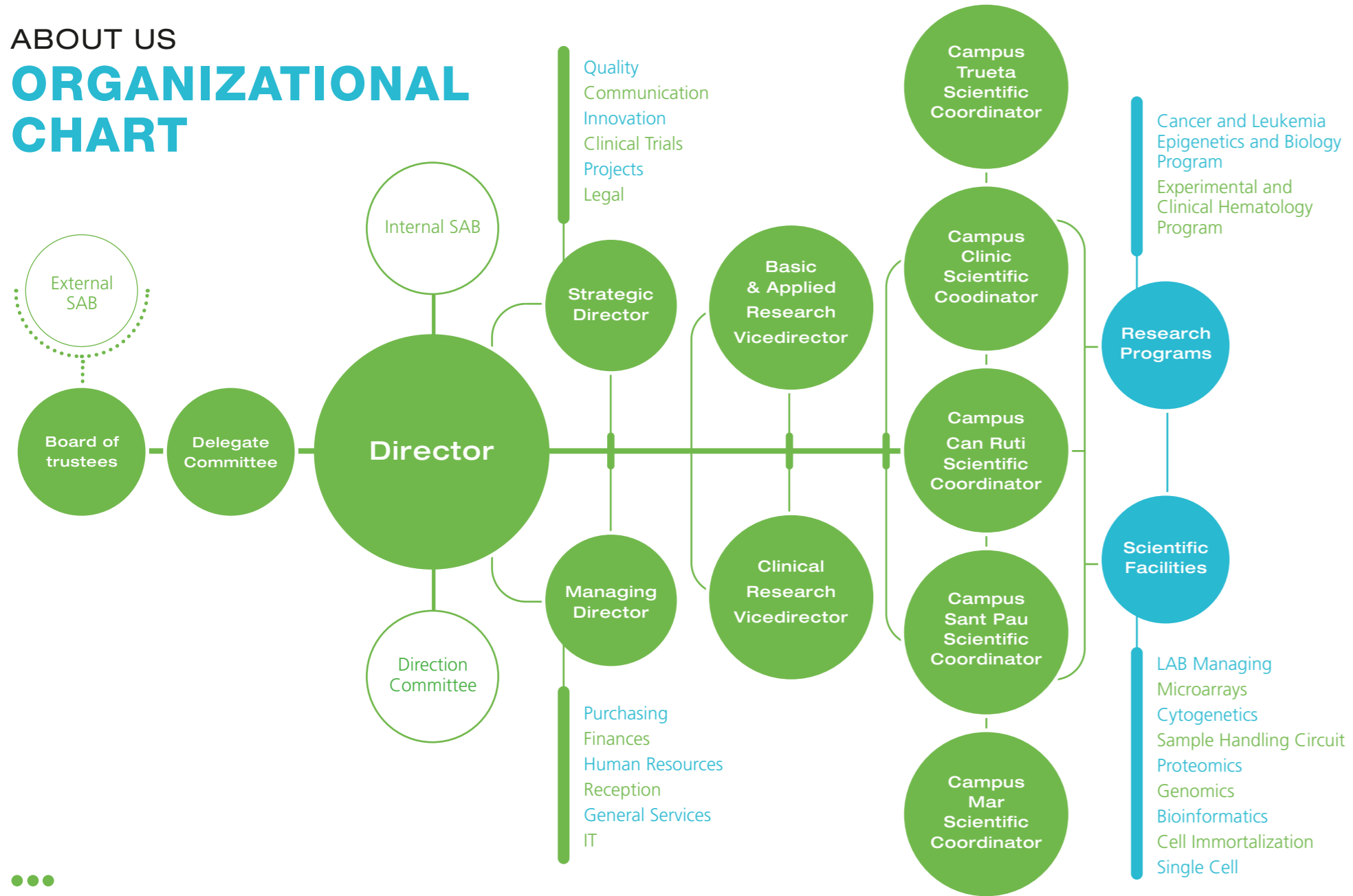


Dr. Javier Briones Mejide  
**Sant Pau Campus Coordinator**



Dr. Francesc Solé Ristol  
**Can Ruti Campus Coordinator**

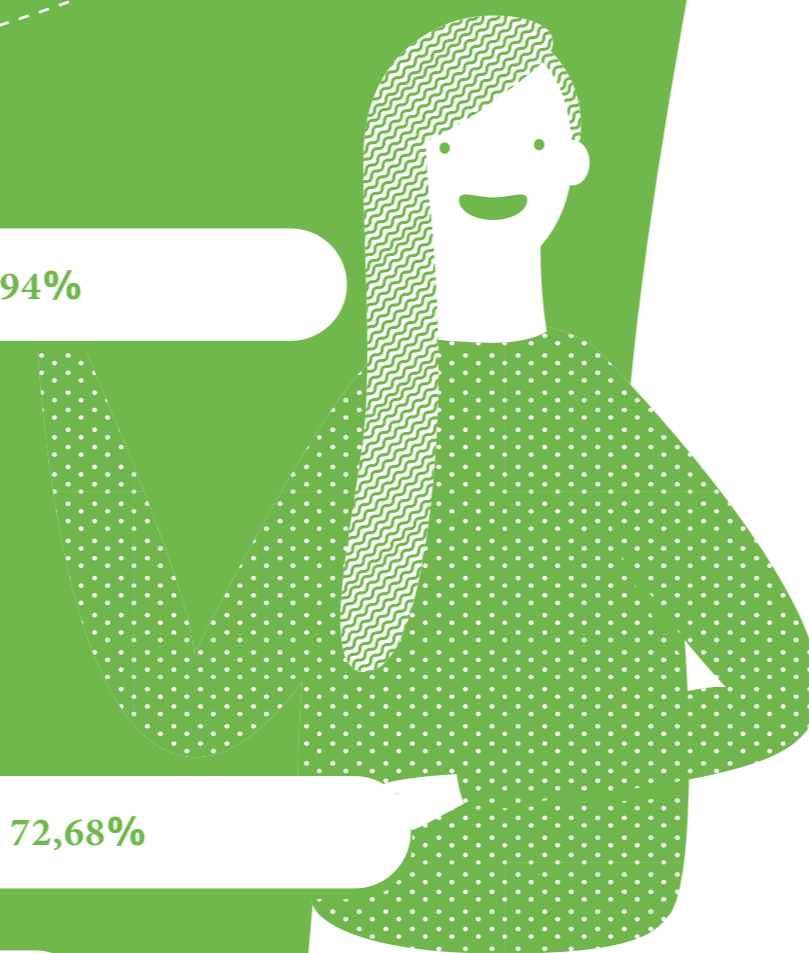
ABOUT US  
**ORGANIZATIONAL CHART**



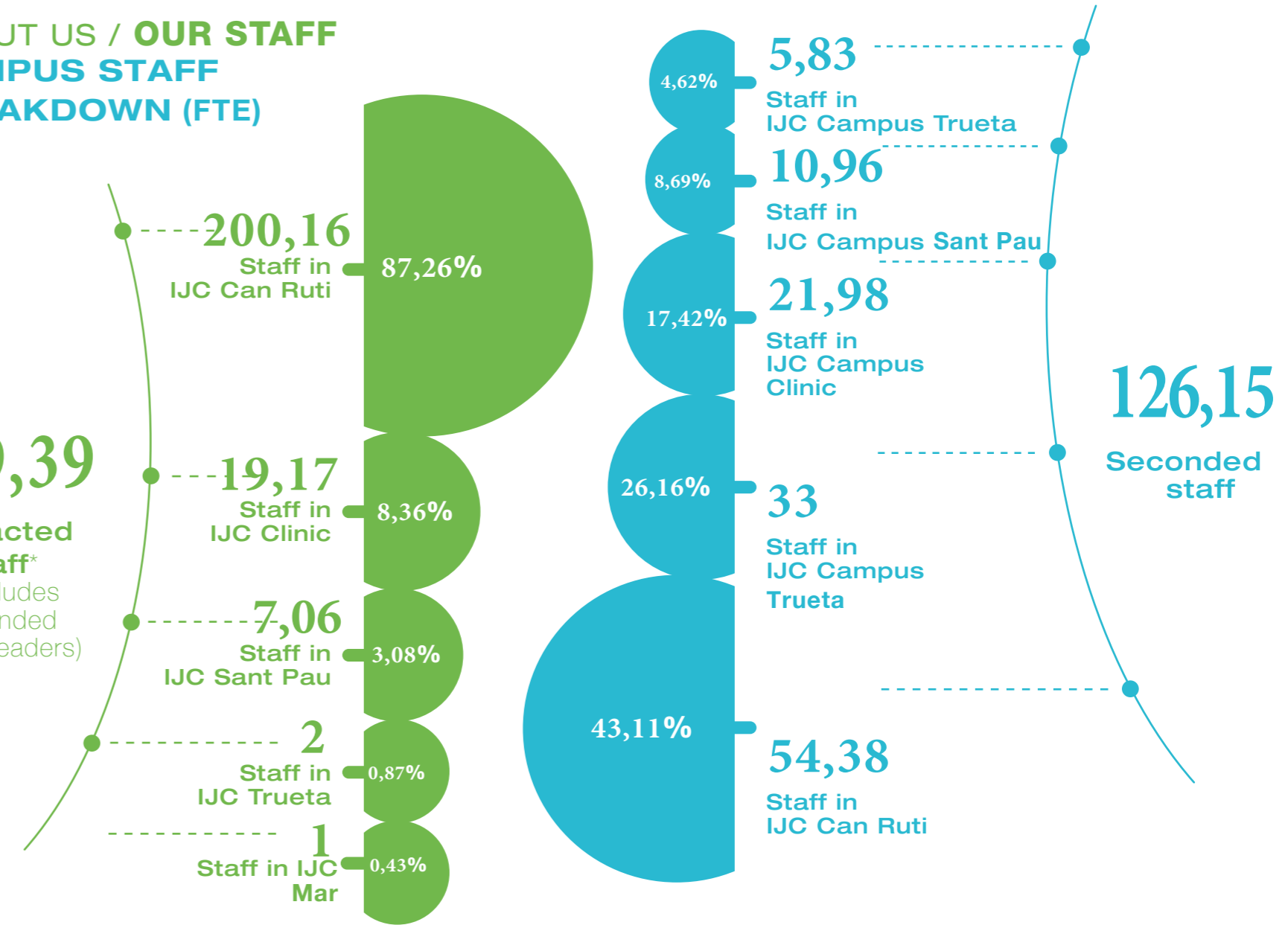


ABOUT US  
**OUR STAFF**

**ADMINISTRATION  
VS. RESEARCH  
BREAKDOWN  
(FTE)**



ABOUT US / **OUR STAFF**  
**CAMPUS STAFF  
BREAKDOWN (FTE)**





## ABOUT US

# RESEARCH PROGRAMS

### CANCER AND LEUKEMIA EPIGENETICS AND BIOLOGY PROGRAM (PEBCL)

1. Cancer Epigenetics led by Manel Esteller
2. Cancer Genetics led by Montse Sanchez-Cespedes
3. Chromatin Biology Laboratory led by Alex Vaquero
4. Chromatin, Metabolism and Cell Fate led by Marcus Buschbeck
5. 3D Chromatin Organization led by Biola M. Javierre
6. Epigenetics and Immune Disease led by Esteban Ballestar
7. Lymphocyte Development and Disease led by Maribel Parra
8. Regulatory Genomics led by Tanya Vavouri
9. Regulatory RNA and Chromatin led by Sònia Guil
10. Epigenetic Control of Hematopoiesis led by José Luis Sardina
11. Transcriptional Dynamics in Leukemia led by Sergi Cuartero
12. Cancer Immunogenomics led by Eduard Porta
13. Cancer Heterogeneity and Hierarchies led by Verónica Rodilla
14. Leukemia and Immuno-Oncology led by Laura Belver
15. Cellular Systems Genomics led by Elisabetta Mereu
16. Stem Cells and Cancer led by Anna Bigas
17. Endothelial Pathobiology and Microenvironment led by Mariona Graupera
18. T-Cell Lymphoma led by Laura Mondragón

### EXPERIMENTAL AND CLINICAL HEMATOLOGY PROGRAM (PHEC)

19. Acute Lymphoblastic Leukemia (ALL) led by Josep M<sup>a</sup> Ribera
20. Barcelona Endothelium Team (BET) led by Enric Carreras
21. Functional Cytomics led by Jordi Petriz
22. Myeloid Neoplasms led by Lurdes Zamora and Blanca Xicoy
23. Immunohematology and Glycobiology led by Fumiichiro Yamamoto
24. Leukemia Stem Cell led by Ruth Risueño
25. Lymphoid Neoplasms led by Tomás Navarro
26. Multiple Myeloma led by Albert Oriol
27. Myelodysplastic Syndromes led by Francesc Solé
28. Stem Cell Biology, Developmental Leukemia and Immunotherapy led by Pablo Menéndez
29. Cellular Immunotherapy and Gene Therapy led by Javier Briones
30. Stem Cell Transplantation and Cellular Immunotherapy led by Álvaro Urbano-Ispizua
31. Epigenetic Therapies led by María Berdasco
32. Lymphoma Translational led by Gaël Roué
33. Descriptive and Analytical Epidemiology of Cancer led by Rafael Marcos Gragera
34. Oncogenesis and Antitumor Drugs led by Ramon Mangués
35. Chronic Lymphocytic Leukemia led by Carolina Moreno
36. Hematology Research led by David Gallardo
37. Myeloid Neoplasms (Clínic) led by Jordi Esteve
38. Hematological Diseases, Transplant and Cell Therapy led by Jordi Sierra
39. Hematological Diagnosis led by Josep Nomdedéu



## ABOUT US **RESEARCH GROUPS**

1

### **CANCER EPIGENETICS LED BY MANEL ESTELLER**

#### **GROUP MEMBERS**

**ESTELLER BADOSA, MANEL**

Group Leader

**SETIÉN BARANDA, ESTEBAN**

FERNANDO

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**DÁVALOS VEGA, MARIA VERÓNICA**

Associate Researcher

**MUSULÉN PALET, EVA**

Associate Researcher

**BLECUA CARRILLO ALBORNOZ,  
PEDRO**

Associate Researcher

**FERRER AGUILAR, GERARDO**

Postdoctoral Researcher

**PONTEL, LUCAS BLAS**

Postdoctoral Researcher

**JANIN, MAXIME HENRI**

Postdoctoral Researcher

**VILLANUEVA LEGARDA, LOREA**

Postdoctoral Researcher

**JOSHI, RICKY**

Postdoctoral Researcher

**CAMPILLO MARCOS, IGNACIO**

Postdoctoral Researcher

**ORŠOLIC, INES**

Postdoctoral Researcher

**CARRIER, ARNAUD**

Postdoctoral Researcher

**ROSSELLÓ TORTELLA, MARGALIDA**

PhD Student

**BUENO COSTA, ALBERTO**

PhD Student

**ORTIZ BARAHONA, VANESSA**

PhD Student

**MARTINEZ VERBO, LAURA**

PhD Student

**GARCIA PRIETO, CARLOS ANTONIO**

PhD Student

**POPOV, ANTON**

PhD Student

**CASADO PELAEZ, MARTA**

PhD Student

**PARRA PARRA, JERONIMO**

PhD Student

**GOMEZ PEREIRA, CRISTINA**

PhD Student

**COLL SAN MARTÍN, LAIA**

Lab Technician

**SOLER RIERA, MARTA**

Lab Technician

## OVERVIEW

Malignant cells behave differently to the rest of the tissue; they present capacities reserved for stem cells, such as proliferation. Since all cells in an organism share the same genetic information, the difference between any cell type, including malignant cells, lies in the subset of information to which they have access. The term epigenetics refers to the many control layers that limit a cell's access to only those parts of the genome relevant to its organic function.

## OUR RESEARCH

Our group aims to understand the fundamental epigenetic and epitranscriptomic mechanisms that act in living cells, such as DNA methylation and histone modification patterns and gene expression regulation through microRNAs. These mechanisms do not affect the cell's genetic information, only its availability, and its alteration may mark the beginning of abnormal behaviour. Malfunction of epigenetic control is one of the most poorly understood causes of human tumors and knowledge of this phenomenon is key to developing new strategies against tumor formation and cancer.

## OUR GOALS

To shed light on this emerging field, the Cancer Epigenetics group focuses on:

### 1. Defining the epigenome of cancer cells.

The ability to identify the epigenetic differences, either on DNA or RNA, between a healthy cell and a transformed cell represents the starting point for our research. Knowledge of this area is fairly limited and our lab is currently carrying out intense research in this field.

### 2. Study of the epigenetic machinery and mechanisms.

We are interested in understanding the role and function of DNA methyltransferases, the large group of proteins directly responsible for interacting with DNA and shaping the open or closed transcriptional state.

### 3. Use of epigenetic markers to predict response to antitumor therapies.

Our group has a long-standing interest in translating the use of epigenetic knowledge gained from research into biomarkers to predict clinical outcomes.

### 4. Preclinical testing of epigenetic compounds.

We are interested in the development and study of new epigenetic drugs that target DNA methylation and histone modification writers, readers and erasers, and that could exert an anti-cancer effect.

## OUR CHALLENGES

Cancer remains a horrible disease, and all new biomarkers and innovative approaches to treat the disorder are necessary and welcome additions. Our research has had an impact on the clinical outcome of several tumor types, from brain tumors to cancer of unknown primary, including subsets of leukemia and lymphoma patients. Discovering the particular epigenetic and epitranscriptomic characteristics of each cancer type may help deliver personalized therapies to oncology patients.

Through our research, we hope to answer the following questions:

1

Will the epigenome map lead to a better understanding of cancer and guide global research to make this disease curable?

2

How can we transfer epigenetics research to prevent cancer and improve clinical management with a view to improving patients' quality of life?

3

Is there an ultimate epigenetic signalling pathway and context that can explain any cancer?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Cancer epigenetics; DNA methylation; RNA epitranscriptomics; histone modification; epigenetic gene silencing





ABOUT US  
**RESEARCH  
GROUPS**

2

**CANCER GENETICS  
LED BY MONTSE SANCHEZ-CESPEDES**

**GROUP  
MEMBERS**

**SANCHEZ-CESPEDES, MONTSE**  
Group Leader

**ROMERO FERRARO, OCTAVIO  
ALFREDO**  
Senior Researcher

**SAIGÍ MORGUÍ, MARIA**  
Postdoctoral Researcher

**ALBURQUERQUE BEJAR, JUAN JOSÉ**  
Postdoctoral Researcher

**FERRERO ANDRÉS, ANA**  
Postdoctoral Researcher

**VILARRUBÍ PORTA, ANDREA**  
PhD Student

**NAVAJAS CHOCARRO, PABLO**  
PhD Student

**MORILLAS VIÑUALES, JUAN**  
PhD Student

**PROS SIMÓN, EVA**  
Lab Technician

**BARTOLESSIS ARIAS, ISABEL**  
Lab Technician

## OVERVIEW

Lung cancer causes over 1.3 million deaths annually and remains the deadliest type of cancer worldwide. Although efforts in recent years to fully characterize human cancer on a genetic and molecular level have provided important insights to increase our understanding of the gene alteration profile underlying the development of Lung Cancer, the impact of this knowledge in the survival of patients remains modest. Our group is devoted to the genetic, epigenetic and molecular study of the mechanisms that drive LC development. Ultimately, our purpose is to implement the clinical management of cancer patients and to design novel therapeutic strategies.

## OUR RESEARCH

The complete genetic characterization of tumors is important to understand cancer development, promote the discovery of new drugs and improve the selection of patients that may benefit from a given cancer therapy. Our research uses the latest high-throughput sequencing technologies to create profiles and catalogues of the recurrently altered genes in cancer. We also have a keen interest in understanding the mechanisms by which the abnormal function of these genes contributes to cancer development.

## OUR GOALS

Our laboratory is currently engaged in a number of important projects:

### 1. **Screening for factors that determine tumor immunoescape and the response to immunotherapy.**

We have become increasingly interested in the study of those biological factors, which allow tumors to escape control of the immune system and determine the response to immunotherapy.

### 2. **Genomic and genetic profiling of lung tumors to identify novel targets for therapeutics and determinants for the primary and acquired response to tyrosine kinase inhibitors (TKIs).**

We use high-throughput genomic sequencing technolo-

gies, such as whole exome and RNA-sequencing, to gather information about the genetic background and gene expression profiles of lung tumors from both smokers and non-smokers.

### 3. **Genetic alterations at epigenetic factors: biological understanding and opportunity for novel therapeutics.**

Over the past 15 years, our group has provided key information to understanding cancer biology. Currently, we are using high-throughput technologies to understand tumor development and to identify molecular vulnerabilities that can be used therapeutically.

## OUR CHALLENGES

Recent epidemiological data point to a worrying increase in the incidence of LC in those who have never smoked, particularly women. The reasons are not well understood, a fact that limits the design of prevention measures.

Through our research, we hope to answer the following questions:

1

What are the genetic and molecular abnormalities that trigger the development of cancer, particularly Lung Cancer?

2

How can we use genetic/molecular information to identify novel targets to implement Lung Cancer therapeutics?

3

What is the molecular basis for the lack of response to immunotherapy?

4

How can we predict and prevent acquired resistance to targeted therapeutics?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Targeted therapeutics; immunotherapy; epigenetic regulation; SWI/SNF-complex; MYC/MAX-pathway



ABOUT US  
**RESEARCH  
GROUPS**

**3** CHROMATIN BIOLOGY LABORATORY  
LED BY ALEX VAQUERO

**GROUP  
MEMBERS**

**VAQUERO GARCÍA, ALEJANDRO**  
Group Leader

**LUIS PAÑOS MOLERO**

**VÁZQUEZ PRAT, BERTA NIEVES**  
Postdoctoral Researcher

**ESPINOSA ALCANTUD,  
MARIA DOLORES**  
Postdoctoral Researcher

**MARAZUELA DUQUE, ANA**  
Postdoctoral Researcher

**FERNÁNDEZ DURAN, IRENE**  
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**IANNI, ALESSANDRO**  
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PhD Student

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PhD Student

**GÁMEZ GARCÍA - CERVIGÓN,  
ANDRES**  
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**SOLÉ GRÍFOL, MARINA**  
PhD Student

**MORILLAS VIÑUALES, JUAN**  
PhD Student

**GUITART SOLANES, ANNA**  
PhD Student



## OVERVIEW

The members of the sirtuin family of NAD<sup>+</sup>-dependent enzymes are key coordinators of this response, as they play an important role in the crosstalk between the environment and the genome, at both cellular and physiological level. In particular, they play a key role in the maintenance of genome stability, epigenetics, metabolic homeostasis, and cell differentiation and development. The relevance of sirtuin function is highlighted by their involvement in some of the most common human pathologies, including cancer (such as blood malignancies), diabetes and other endocrine-related diseases, neurodegenerative diseases and ageing.

## OUR RESEARCH

The chromatin biology lab's primary purpose is understanding the mechanisms of the stress response and their impact on cancer and ageing. Specifically, the group focuses its efforts on defining the contribution of sirtuins to this response in the maintenance of genome stability, epigenetics and metabolic homeostasis.

To fulfil this main objective, the group's work encompasses a wide range of research areas, from basic aspects of sirtuin biology to their contribution in the development of human pathologies such as leukemia and ageing.

## OUR GOALS

We aim at the identification of novel mechanisms and factors involved in the onset and development of blood malignancies, and the creation of tools that could be helpful for its diagnosis and treatment. In this regard, the group's main objectives are:

**1.**  
**To understand the enzymatic duality of sirtuins and their specific contribution to sirtuin function.**

In particular, we focus our efforts on the poorly understood topic of ADPRT activity.

**2.**  
**To characterize sirtuin-dependent mechanisms of genomic stability,** including constitutive heterochromatin integrity, DNA damage

signalling and repair, and cell cycle checkpoint control.

**3.**  
**To define the role of sirtuins in B-cell differentiation and characterize their functional implication in cancer,** particularly in the context of hematopoietic pathologies such as leukemia and lymphoma. Our main efforts are currently focused on two types of leukemia, pediatric B-ALL and AML.

**4.**  
**To understand the involvement of sirtuin function in the beneficial effects of nutrient restriction on ageing development.**

**5.**  
**To develop a new methodology to measure the activity of sirtuins *in vivo*.**

## OUR CHALLENGES

Through our research, we seek to answer the following questions:

**1**  
What is the physiological mechanism associated with the genotoxic and metabolic stress response?

**2**  
What is the contribution of the sirtuin family of enzymes to the maintenance of genome stability after stress?

**3**  
What is the implication of these mechanisms in the onset and development of blood cancers and ageing?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Stress response; sirtuins; epigenetics; leukemia; ageing





ABOUT US  
**RESEARCH  
GROUPS**

4

**CHROMATIN, METABOLISM AND CELL FATE  
LED BY MARCUS BUSCHBECK**

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**CASTANYER COSTA, JOAN**  
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**VALERO LÁZARO, VANESA**  
Lab Technician

## OVERVIEW

We focus on understanding the molecular aspects of chromatin regulation and have a long-standing interest in the study of histone variants. We want to find ways to translate knowledge about chromatin regulation into therapeutic tools for the management of diseases such as blood cancers.

## OUR RESEARCH

We seek to bridge the gap between basic molecular research and translational research by exploring chromatin regulation, in particular the molecular biology of histone variants. We aim to exploit this knowledge for the identification of novel intervention strategies for the treatment of blood cancers. We focus on the continuum of myeloid diseases, ranging from the premalignant expansion of altered clones to chronic myelodysplastic syndromes and acute myeloid leukemia.

## OUR GOALS

Through our research, we aim to gain a better understanding of the epigenetic mechanisms that contribute to the development of blood cancers. By functionally mining the chromatin regulatory space, we further aim to provide new starting points by identifying novel drug targets. In this regard, our research focuses on two main lines:

**1.**

To mine the chromatin regulatory space to identify novel drug targets that can either help improve current treatments or intercept disease at an early asymptomatic stage.

**2.**

We study histones from the protein core of the nucleosome, particularly the variant macroH2A that led to two major discoveries: its major role in nuclear organization and its ability to bind metabolites through its mostly understood macrodomain, establishing a direct link between chromatin and metabolism.

## OUR CHALLENGES

Through our research, we hope to answer the following questions:

**1**

How do epigenetic mechanisms operate on the molecular level?

**2**

How do chromatin and, in particular, histone variants contribute to cell fate transitions?

**3**

How can we exploit this knowledge for the development of novel therapeutic strategies?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

myelodysplastic syndrome, acute myeloid leukemia, chromatin, nuclear organization, histone variants



ABOUT US  
**RESEARCH  
GROUPS**

5

**3D CHROMATIN ORGANIZATION  
LED BY BIOLA M. JAVIERRE**

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**LÓPEZ MARTÍ, PAULA**  
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PhD Student

**ALCALDE MERINO, ÁLVARO**  
PhD Student



## OVERVIEW

Our group combines cutting-edge experimental and bioinformatics approaches to understand the specific 3D chromatin organization of normal and malignant hematopoiesis and its interaction with non-coding determinants and trans-regulatory elements. Our long-term goal is to keep making progress in the fight against cancer. We will not stop until a cure is found.

## OUR RESEARCH

Since 2018, our work is focused on the study of the molecular mechanisms driving healthy and malignant hematopoiesis under a three-dimensional perspective. We are deeply interested in the unexplored noncoding DNA, which encompasses DNA that does not contain genes but that regulate the expression of remote genes, and the alterations of this. For that, we are developing new methods to study the spatial genome architecture and integrating this data with different methods in epigenetics, biochemistry and functional genomics to provide fundamental insight on normal and malignant hematopoiesis.

## OUR GOALS

Our lab's main research goals are as follows:

- 1. To define the cell type-specific 3D chromatin organization in human hematopoietic cells.**  
we aim to investigate whether the dynamic changes in chromatin interactions between gene promoters and regulatory elements can shape transcription decisions controlling hematopoiesis.
- 2. To identify the altered DNA topology in blood cancer.**  
Genome architecture plays a key role in genome expression regulation. Chromatin interactions are therefore crucial for cellular health, and errors in these interactions can give rise to the development of a broad range of diseases, including blood cancer.

- 3. To prioritize new candidate genes and pathways related to blood cancer.**  
By studying the physical interactions between gene promoters and regulatory elements, we will be able to connect blood cancer determinants to putative target genes, thereby prioritizing new candidate genes and pathways and offering an insight into the genomic regulatory mechanisms underlying cancer.

## OUR CHALLENGES

As in other complex diseases, most of mutations and epimutations associated with blood cancers lie in non-coding regions, frequently at enhancers, and potentially exert their roles by altering the regulation of the target genes. However, these non-coding determinants remain unexplored because the vast majority of regulatory elements that control transcriptional activity of each gene in each cell type are unknown.

Through our research, we hope to answer the following questions:

- 1** Can the dynamic changes in chromatin interactions shape the transcription decisions controlling hematopoiesis?
- 2** How does the altered genome architecture drive malignant transformation?
- 3** What is the role of non-coding determinants in cancer predisposition and development?

## ABOUT US

# RESEARCH GROUPS

## KEYWORDS

Genome architecture, spatial-temporal chromatin organization, hematopoiesis, blood cancer, cis non-coding determinants, enhancer-promoter interactions



ABOUT US  
**RESEARCH  
GROUPS**

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**EPIGENETICS AND IMMUNE DISEASE  
LED BY ESTEBAN BALLESTAR**

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Lab Technician

## OVERVIEW

We aim at understanding the mechanisms underlying the deposition and removal of epigenetic modifications in immune cells, the influence of genetic and environmental determinants, and the acquisition of epigenetic alterations in immune-mediated disease including primary immunodeficiencies, autoimmune and autoinflammatory diseases. We also investigate the impact of the epigenetic regulation of immune cells in the microtumor environment.

## OUR RESEARCH

We started these lines of research more than 10 years ago, by studying the occurrence of DNA methylation alterations in the context of systemic lupus erythematosus (SLE), an archetypical systemic autoimmune disease. Later on, we performed new studies with MZ twins discordant for common variable immunodeficiency (CVID), the most prevalent symptomatic primary immunodeficiency.

More recently, our team also demonstrated the occurrence of DNA methylation alterations in monocytes in representative autoinflammatory syndromes. We have shown that alterations in the DNA methylome of peripheral blood monocytes reflect the disease activity in rheumatoid arthritis mediated by the elevated levels of inflammatory cytokines present in such state.

## OUR GOALS

Our main lines of research and specific goals are:

### 1. To understand the role of epigenetic control and its upstream determinants in relation with immune function.

We aim at understanding how immune cell-cell crosstalk, cytokines and other factors, cell signalling pathways and transcription factors determine epigenetic control and impact immune cell function.

### 2. To identify epigenetic alterations in immune-mediated diseases and investigate their clinical relevance.

Our studies focus on different diseases including primary immunodeficiencies, such as common variable immunodeficiency (CVID)

and hyper IgM type 2 syndrome, and autoimmune diseases, such as rheumatoid arthritis, systemic sclerosis and systemic lupus erythematosus.

### 3. To investigate the effects of immunomodulators and epigenetic compounds in shaping the epigenome and responses of immune cells.

We dissect the molecular consequences of different immunomodulators as well as inhibitors of epigenetic enzymes in immune cells.

## OUR CHALLENGES

The study of epigenetic dysregulation can help understand the determinants of immune dysregulation and can have an impact in the treatment of these diseases. Therefore, with our research we want to answer:

- 1 How do immune cells translate the surrounding information provided by the direct contact with other cells or the cytokines and other molecules into epigenetic profiles that determine their responses?
- 2 What is the relevance of the epigenetic alterations that are found in different immune mediated diseases in relation to the aberrant function of these cells?

- 3 How can we apply the knowledge on the epigenetic dysregulation in immune-mediated disease to the clinics?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Epigenetics, DNA methylation, Immune-mediated disease, autoimmune disease, primary immunodeficiency





ABOUT US  
**RESEARCH  
GROUPS**

7

**LYMPHOCYTE DEVELOPMENT AND DISEASE  
LED BY MARIBEL PARRA**

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Lab Technician



## OVERVIEW

B-cell lymphopoiesis is a complex developmental process that involves several cellular transitions, including cell commitment and early and late cellular differentiation. Proper transcriptional control at each cellular transition is essential for the correct development of B lymphocytes. How specific gene expression programmes are selected and maintained, thus resulting in the proper generation of B cells, remains a fundamental question in biology. Conversely, how the aberrant establishment of cell- and lineage-specific gene transcriptional programmes leads to the development of B-cell malignancies such as leukemia and lymphoma also requires extensive research.

## OUR RESEARCH

Our current research focuses on four main lines:

- **Understanding the entire role of HDAC7 in early and terminal B-cell development.** HDAC7 is an epigenetic modulator that represses functional or

lineage-inappropriate gene expression in B lymphocytes

- **Establishing HDAC7 as a novel biomarker and potential therapeutic target in pro-B acute lymphoblastic leukemia (pro-B-ALL) and diffuse large B-cell lymphoma (DLBCL).** We found that the deregulation of HDAC7 may be involved in the pathogenesis of acute lymphoblastic leukemia.
- **Working towards precision medicine against DLBCL heterogeneity using organoid culture systems.** We are investigating additional epigenetic regulators in normal and aberrant B-cell generation and implementing 3D organoid cultures from DLBCL sample patients.
- **Improving immunotherapy combinatorial therapy in DLBCL.** R-CHOP is the gold standard treatment for DLBCL patients. R-CHOP therapy combines anti-CD20 antibody (immunotherapy) with cyclophosphamide, doxorubicin, vincristine and chemotherapy.

## OUR GOALS

Acute lymphoblastic leukemia (ALL) is the most common type of cancer in children under a year old. Even though the chances of survival in infants suffering from ALL have improved significantly in recent years, an exhaustive study of the mechanisms underlying this disease is still required to make further therapeutic advances. Therefore, our group focus on:

- 1.** To understand how gene silencing is established during normal and aberrant B-cell differentiation.
- 2.** To transfer our basic knowledge in the epigenetics and transcriptional control of B-cell development to the clinical setting for infant B-ALL and DLBCL patients.

- To identify small molecules aimed at HDAC7 targeted modulation for combinatorial and precision medicine in infant pro-B-ALL with MLL-AF4 rearrangement.
- To identify novel targets for the design of next-generation immunotherapies in DLBCL.

## 3.

To implement a 3D organoid platform for DLBCL patient samples to perform compound library screenings aimed at unveiling new drugs for use in combinatorial therapy with current immunotherapy in a personalized manner.

## OUR CHALLENGES

Through our research, we aim to answer the following questions:

- 1** How do B lymphocytes decide their identity? How is gene silencing established?
- 2** Why does HDAC7 expression improve the prognosis of some hematopoiesis diseases?
- 3** Why is HDAC7 underexpressed in pro-B-ALL and DLBCL?
- 4** How can we restore HDAC7 expression in pro-B-ALL and DLBCL to impair disease progression?
- 5** Can we implement 3D organoids from DLBCL patients aimed at drug screening towards a precision medicine strategy and immunotherapy improvement?

## KEYWORDS

B lymphocyte development, Epigenetics and transcriptional regulation, HDAC7, B cell acute lymphoblastic leukemia (B-ALL), Diffuse large B-cell lymphoma (DLBCL)

## ABOUT US RESEARCH GROUPS



ABOUT US  
**RESEARCH  
GROUPS**



**REGULATORY GENOMICS  
LED BY TANYA VAVOURI**

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## OVERVIEW

Regulation of gene expression is the fine-tuning of the synthesis of the functional product of genes and is one of the most fundamental processes in life. It is the process that makes different cell types have different properties and differentiates unhealthy from healthy cells. Gene expression is regulated by internal signals (the activity of other genes, mutations, etc.) and by external signals (diet, temperature, pharmacological therapies, etc.).

## OUR RESEARCH

Our research focuses on three main areas:

- Firstly, we study **the effect of the environment on gene expression** changes that are transmitted from parents to their offspring. We want to understand how information about our exposure to different environments may be encoded in molecules - other than DNA - inside germ cells that are transmitted between generations.

- Secondly, we work on **non-coding RNAs and other non-coding elements that influence gene expression**. We are interested in which non-coding elements affect gene expression and how.
- Finally, we want to **understand how epigenetic drugs affect gene expression and chromatin** in different genomic contexts. Epigenetic drugs currently used in the clinic include those for the treatment of patients with acute myeloid leukemia and myelodysplastic syndrome. A more in-depth understanding of the effects of these drugs and how they work may lead to improved or more personalized medicine in the future.

## OUR GOALS

Our aim is to contribute to a better understanding of gene regulation and the consequences of drug treatments and inter-individual genetic variation in gene expression. Although most of our research is based on data from animal model organisms or cell lines, we hope that, in the long term, the knowledge acquired will increase our understanding about humans. Extensive aberrant gene expression and genome deregulation are extremely common in cancer, especially hematopoiesis forms, and treatments targeting gene regulation pathways are being used for hematopoiesis malignancies. Last, but not least, we hope that the data we generate and the analysis methods we develop serve as useful tools for the wider research community.

## OUR CHALLENGES

We hope that our research sheds light on the following questions:

- 1 Which epigenetic mechanisms are involved in the transmission of acquired or variable traits between generations in humans and other animals?
- 2 Which non-coding DNA elements affect gene expression and therefore potentially phenotype?
- 3 How drugs (such as those used for the treatment of blood cancers) affect gene expression and the function of the non-coding parts of our genome?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Bioinformatics, gene regulation, epigenetic inheritance, germline, genomics





ABOUT US  
**RESEARCH  
GROUPS**

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**REGULATORY RNA AND CHROMATIN  
LED BY SÒNIA GUIL**

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**SRINIVAS, TARA**  
PhD Student

## OVERVIEW

We study the emerging roles of noncoding RNAs as key regulators of gene expression in physiological cellular programmes and at the onset or during progression of human diseases, with a major focus on tumorigenesis and neurodevelopmental diseases. The research carried out by our group combines biochemical, cellular and global genomic approaches to dissect mechanisms of gene expression regulation with the participation of ncRNAs, with the ultimate aim of revealing molecules of therapeutic/biomarker interest for clinical translation.

## OUR RESEARCH

The lab focuses on a variety of RNA and RNA-binding protein functions in the context of changing cellular conditions. Our interest concentrates on the noncoding transcriptome, with the main aim of separating the wheat from the chaff to reveal true biologically relevant molecules and to understand how they are connected to broader gene regulatory networks.

Cancer research has led the way in the study of noncoding RNAs, but the abundance and key roles of the noncoding transcriptome in the human brain are being increasingly recognized. Importantly, common dysregulated mechanisms in different pathological contexts and with the involvement of ncRNAs, are emerging.

## OUR GOALS

Our research aims to gain a better understanding of the biological relevance of ncRNAs for an informed use in therapeutic strategies. Recently, our group's research has taken advantage of state-of-the-art global transcriptomic approaches to identify ncRNA candidates that act as master regulators of oncofoetal genes, thereby revealing their validity as biomarkers in human cancer.

In addition to our work related to cancer, the group has been developing new experimental tools for research into Rett syndrome, a neurodevelopmental disorder usually caused by loss-of-function mutations in the epigenetic regulator MeCP2.

## OUR CHALLENGES

Through our research, we hope to answer the following questions:

- 1 What is the precise contribution of the non-coding transcriptome to tumor biology?
- 2 How can we use RNA tools to improve treatment or diagnosis of human disease?

- 3 How can we better model neurodevelopmental diseases such as Rett syndrome to understand key initial changes in gene expression programmes?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Noncoding RNAs, cancer epigenetics, gene expression regulation, stem cells, Rett syndrome.



ABOUT US  
**RESEARCH  
GROUPS**

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**EPIGENETIC CONTROL OF HEMATOPOIESIS  
LED BY JOSÉ LUIS SARDINA**

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## OVERVIEW

DNA methylation-related genes are among the most frequently mutated in blood malignancies. Traditionally, studies aimed at understanding the effect of aberrant DNA methylation in cancer patients have focused on gene promoters. However, recent findings focus on enhancers as the most important regions in dynamic DNA methylation studies. We aim to understand how aberrant DNA methylation dynamics impact on the chromatin structure at enhancers during blood cancer onset and progression.

## OUR RESEARCH

We study how aberrant DNA methylation at distal gene regulatory regions poisons the chromatin to trigger corrupted gene expression signatures in cells, thus eventually leading to the onset and progression of hematopoiesis neoplasms. This line of research has implications for a broad spectrum of patients suffering from blood diseases sharing an abnormal genome-wide DNA methylation landscape.

## OUR GOALS

Our research aims to gain a better understanding of the biological relevance of ncRNAs for an informed use in therapeutic strategies. Recently, our group's research has taken advantage of state-of-the-art global transcriptomic approaches to identify ncRNA candidates that act as master regulators of oncofoetal genes, thereby revealing their validity as biomarkers in human cancer.

In addition to our work related to cancer, the group has been developing new experimental tools for research into Rett syndrome, a neurodevelopmental disorder usually caused by loss-of-function mutations in the epigenetic regulator MeCP2.

## OUR CHALLENGES

There is an urgent need for novel therapies for acute myeloid leukemia, since barely any drugs introduced in the last decades have increased the overall survival of its patients. Hence, our research aims to shed light on the following questions:

- 1** What is the interplay between DNA (hydroxy) methylation and chromatin dynamics at distal gene regulatory regions during hematopoietic cell fate decisions?
- 2** What are the molecular mechanisms underlying the role of TET2 in the epigenetic control of the chromatin at distal gene regulatory regions during leukemia onset and progression?
- 3** What is the role of mRNA methylation-mediated post-transcriptional control in myeloid cell differentiation?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

NA methylation; TET enzymes; Chromatin; Hematopoiesis malignancies; Stem cells





ABOUT US  
**RESEARCH  
GROUPS**

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**TRANSCRIPTIONAL DYNAMICS IN LEUKEMIA  
LED BY SERGI CUARTERO**

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**LORENZI FARÍAS, LUCÍA**  
Lab Technician

## OVERVIEW

Hematopoietic differentiation is a tightly regulated process that maintains blood production throughout life. The transcriptional changes that hematopoietic cells undergo during differentiation are controlled at multiple levels and an accurate integration of all of them is essential to ensure the production of sufficient numbers of blood cells at all stages of differentiation. However, most of the acute myeloid leukemia (AML) cases have mutations in transcriptional regulators and chromatin modifiers. These mutations alter transcriptional dynamics and can impair normal differentiation.

## OUR RESEARCH

We study the mechanisms that regulate transcription during hematopoietic differentiation and investigate the leukemogenic potential of mutations in transcriptional regulators and epigenetic modifiers. We are also looking into the role of mutations in proteins that drive the three-dimensional organization of the genome.

## OUR GOALS

Our main goals are:

**1.**  
**To understand the role of mutations in hematopoietic transcription factors and chromatin regulators in acute myeloid leukemia (AML).**

Using genetic models to mimic these mutations, we aim to dissect their impact on gene expression and thus understand how they promote a selective advantage.

**2.**  
**To characterize the impact of inflammatory signalling on normal hematopoietic differentiation and during leukemic progression.**

We want to understand the impact of inflammation on the progression of myeloid malignancies and how are they linked to the most common mutations.

## OUR CHALLENGES

Acute myeloid leukemia (AML) is the one of the most aggressive forms of leukemia, and there is an urgent need to find new treatment options. While we now know **what** genes are recurrently mutated in AML, we still do not understand **why** these mutations are malignant. Through our research, we aim to answer the following questions:

**1**

What transcriptional mechanisms are deregulated in acute myeloid leukemia?

**2**

How do inflammatory signals influence leukemic progression?

**3**

Can we use inflammatory modulation to attenuate the severity of myeloid malignancies?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Hematopoiesis, chromatin, AML, MDS, cohesin, inflammation



ABOUT US  
**RESEARCH  
GROUPS**

**12.** **CANCER IMMUNOGENOMICS  
LED BY EDUARD PORTA**

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**CERVILLA GARCÍA, SERGI**  
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**GRASES MENDOZA, DANIELA**  
Lab Technician



## OVERVIEW

Our research lies at the interface of artificial intelligence, molecular biology and medical oncology, and we bring together experts from all three fields. We use computational approaches to study the interaction between genetic variants in cancer genomes and multiple aspects of cancer, ranging from the immune response against tumors to the susceptibility of cancer cells to different treatments.

## OUR RESEARCH

Over the last two decades, there has been an explosion of three main types of big data in cancer research. We use three types of data:

- Germline genotypes from cancer patients, this is, the base non-altered genome of individual people being diagnosed with cancer.
- Somatic tumor genomes, being the particular genomes of individual tumors, with its unique set of alterations.
- The amount and composition of cells in tumors coming from single-cell sequencing.

Until now, these three different aspects of tumor immunobiology have mostly been studied on an individual basis. However, it is now evident that the three factors are inextricably linked and should be studied as a whole.

## OUR GOALS

Our main goal is to understand how genetic variation influences the immune response against cancer cells and vice versa. Specifically, we are working on the following lines:

- 1.** Understanding how inherited genetic variants change the immune response against cancer cells.
- 2.** Understanding how inherited genetic variants interact with biological sex to influence cancer predisposition. We are using the MareNostrum supercomputer to analyze genetic data from hundreds of thousands of cancer patients from this biological sex perspective, to identify genetic variants that predispose to cancer differently depending on gender.
- 3.** Integration of protein structure and genetic data to identify new cancer-associated mutations.
- 4.** Creating a molecular and cellular map of the tumor microenvironment in bladder cancer.

## OUR CHALLENGES

Through our research, we hope to answer the following questions:

- 1** Is it possible to use a person's genetic data to predict whether he/she will develop cancer?
- 2** Which genes play a role in the development of cancer?
- 3** How do genetic variants change the immune response against cancer cells?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Computational biology, cancer genomics, big data, GWAS, bladder cancer



ABOUT US  
**RESEARCH  
GROUPS**

**13.** **CANCER HETEROGENEITY AND HIERARCHIES  
LED BY VERÓNICA RODILLA**

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Lab Technician

## OVERVIEW

Our laboratory studies the key signals governing stem cell and cell fate specification during malignant progression and the mechanisms by which different signaling pathways control cell plasticity in cancer. Specifically, we use in vivo lineage tracing, live imaging, cytometry and expression profile analysis as experimental tools to achieve our goals. Our group combines murine transgenic models, patient-derived xenografts and 3D organoids to unravel cellular hierarchies within tumors, to gain a better understanding of cancer heterogeneity and drug resistance.

## OUR RESEARCH

Cancer is a heterogeneous disease with a cellular hierarchical organization that is largely unexplored in many tumor subtypes. Moreover, in some cases hierarchical relationships among stem cells, progenitors and differentiated cells remain unsolved due to the high degree of cellular plasticity, which allows cells to switch between different cellular stages.

## OUR GOALS

We are a newly created group passionate about cellular hierarchies and tumor heterogeneity. Our main lines of research and specific goals are:

### 1. To illustrate cellular hierarchies within tumors.

We use a well-established hierarchical model to study multipotency in tumors. Now, we are separately monitoring three mammary epithelial compartments to measure the presence of multipotency within breast tumors. Our hypothesis is that breast tumors.

### 2. To discover cytotoxic agents for specific cellular subpopulations.

A therapy based on a combination of several drugs to target different cellular populations could eradicate primary tumors, thereby preventing relapse and metastasis. We want to

screen for natural compounds that selectively kill specific subsets of cells that are responsible for tumor maintenance and/or intrinsically resistant to current therapies.

### 3. To target the tumor niche to prevent the spread of cancer.

One of our main objectives is to generate in vivo tools that will allow us to study new therapeutic targets to prevent relapses in hematological cancer. To that end, our lab works on different strategies, which include murine and human models, to test a panel of drugs currently used as a standard of care for non-Hodgkin's lymphoma (NHL) and explore the role of senescence in tumors cells, as well as in their microenvironment.

Our ultimate mission is to understand the tumor heterogeneity between different patients with a view to improving their treatment of choice by searching for novel and personalized therapeutic strategies.

## OUR CHALLENGES

We hope to answer the following questions through our research:

1. How can cellular plasticity improve treatment for cancer patients?

2. Can we achieve truly personalized medicine by identifying single or combinatorial therapies to target different cellular populations at the same time?

3. Can we prevent metastasis and/or relapses by targeting the most frequently colonized tissues?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Cellular hierarchies, heterogeneity, cell plasticity, senescence, tumor microenvironment





ABOUT US  
**RESEARCH  
GROUPS**

**14** | **LEUKEMIA AND IMMUNO-ONCOLOGY  
LED BY LAURA BELVER**

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## OVERVIEW

Juvenile myelomonocytic leukemia (JMML) is a rare and aggressive type of pediatric blood cancer that affects mainly children under two years of age and occurs when bone marrow production of white cells becomes severely dysregulated. Hematopoietic stem cell transplantation (HSCT) is currently the only available treatment for patients, although just two out of three children affected by the disease survive.

## OUR RESEARCH

Since 2020, our work has focused on the study of the molecular mechanisms driving JMML and the exploration of alternative therapeutic strategies specifically designed for these patients. To achieve this, we incorporate different methods into epigenetics, systems biology, functional genomics and biochemistry, to help address critical questions about the origin and progression of JMML and to identify new therapeutic targets for the treatment of this disease.

## OUR GOALS

The specific goals of our research program are as follows:

### 1. **To create a centralized JMML sample repository and patient-derived xenograft (PDX) collection.**

In collaboration with clinical teams at different hospitals around the country and with the endorsement of the Spanish Society of Pediatric Hematology and Oncology (SEHOP), we are creating a national JMML patient sample repository and PDX collection, that will be instrumental for the development of our research program.

### 2. **To develop a comprehensive molecular analysis of JMML patients to define accurate diagnostic and stratification criteria.**

We aim to explore the non-coding genome of JMML

patients to identify new genetic alterations that can drive JMML or contribute to the pathogenesis of the disease by other means. This information will help improve our knowledge of JMML and develop more accurate criteria for the diagnosis and management of JMML patients.

### 3. **To identify new potential therapeutic targets and develop specific therapies for the treatment of JMML.**

HSCT is currently the only effective therapy for the treatment of JMML. However, only two out of three children with this disease survive. Thus, new therapies specifically designed to treat JMML patients are needed. To achieve this, we are collaborating with other research groups and with biotechnology companies to identify specific JMML therapeutic targets and explore strategies for their clinical use for the treatment of JMML patients.

We are confident that our results will have an important impact on the diagnosis and treatment of JMML by increasing knowledge of the disease and expanding the therapeutic options open to these patients. Moreover, our close collaboration with clinical teams in the development of this project will guarantee that our most promising discoveries have a rapid and direct impact on JMML patients.

## OUR CHALLENGES

We hope to answer the following questions through our research:

1

What is the relevance of non-coding somatic mutations in the generation and development of JMML?

2

Can non-coding mutations predict the prognosis of JMML patients?

3

What are the best therapeutic targets for the development of JMML-specific treatments?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Leukemia, JMML, PTPN11, experimental therapeutics, CAR-T cells, rare diseases, pediatric diseases, sequencing, diagnosis, therapeutic targets, preclinical models, drug discovery



ABOUT US  
**RESEARCH  
GROUPS**

**15.** CELLULAR SYSTEMS GENOMICS  
LED BY ELISABETTA MEREU

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## OVERVIEW

In the interface between genomics, digital pathology and artificial intelligence the **Cellular Systems Genomics** group aims to define the spatiotemporal organization of complex tissues in health and disease, by the identification of key regulatory mechanisms driving heterogeneity in cellular identity and function, particularly in the context of inflammation, inflammatory disorders and autoimmune diseases.

To address these questions, we will adopt a single-cell perspective, enabling the fine-grained and spatially resolved molecular profiling of tissues. We will develop new machine learning approaches and open source tools in order to unlock molecular mechanisms hidden in large-scale datasets. In a short-term

perspective, these methods will help understand disease mechanisms, allowing the stratification of patients based on their molecular and cellular characteristics, ultimately providing new therapeutic targets for their treatments.

## OUR RESEARCH

Single cell sequencing allows to profile thousands of individual cells per experiment, enabling the unbiased analysis of tissues, organs and even entire organisms at an unprecedented resolution. These data represent a powerful tool for cell biology, with relevant clinical applications including diagnosis and treatment of diseases. Despite the many advantages of this approach, data are noisy and sparse, making

the computational analysis challenging. To address these challenges, we apply machine learning and other statistical methods to develop new analytical frameworks and open source tools to analyze, interpret and integrate data coming from single-cell and spatial genomics experiments.

As part of the Human Cell Atlas (HCA) consortium, which aims to create a catalogue of all cell types in our body, we have extensive experience on the systematic comparison of protocols in single cell RNA sequencing (scRNA-seq). In conjunction with the new Single Cell Unit of the Institute, which is equipped with the Chromium controller to perform the single-cell analysis, we will provide support to design new experiments and generate high-quality data and computational

analysis.

Beyond transcriptomic profiling with scRNA-seq, different cellular modalities can now be measured, including single-cell epigenetics (scATAC-seq), spatial transcriptomics as well as the joint profiling of chromatin accessibility and transcription on the same cell.

However, the integration of multimodal data poses new analytical challenges and new benchmarking are needed to assess reproducibility and integrity of these methods. We are working on new mathematical frameworks for the integration of multimodal data, enabling the comprehensive characterization of cells in their identity and function.

## OUR GOALS

In the European Pancreas Atlas consortium (ESPACE, <https://www.espace-h2020.eu>), we are working to build a first version of the Human Cell Atlas of the Pancreas, by profiling the transcriptome and epigenome of cells from distinct anatomical regions of the adult pancreas. The integration of distinct single-cell and spatial data types will allow the comprehensive transcriptional and epigenetic landscape of pancreas cell types within their spatial context.

Our experience in single-cell data analysis on healthy and diseased tissues allowed us to build a deep understanding of cell-type structure and plasticity in different research contexts. To accelerate biological discovery and advance science, our group will share user-friendly computational solutions, by promoting open science, diversity and supporting an inclusive and collaborative environment. We welcome proposals for interdisciplinary research collaborations, from both industry and academia.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Genomics, inflammation, autoimmunity, single cell, machine learning, computational analysis



ABOUT US  
**RESEARCH  
GROUPS**

**16** | **STEM CELLS AND CANCER  
LED BY ANNA BIGAS**

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## OVERVIEW

Our research group investigates how to generate and maintain the stem cells in the hematopoietic system under physiological conditions but also how these processes are mimicked by the tumors for their perpetuation. We constantly improve our research by implementing novel technology to understand the process of normal and malignant hematopoietic development. Our research includes basic studies at the molecular level to understand cellular processes in the context of mouse models and human patients.

## OUR RESEARCH

Our research comprises from basic biochemical research to the analysis of animal models that reproduce the pathologies of interest and allow us to study the functional relevance of new hypothesis. The ultimate goal is to confirm the importance of the findings and study possible therapeutic applications through the analysis of patient samples. In this sense we have devoted special efforts to understand the regulation of hematopoietic stem cells, as a tool to understand the mechanisms that regulate leukemia initiation and maintenance.

## OUR GOALS

The specific goals of our research program are as follows:

### 1. Generation of hematopoietic stem cells.

Our current studies are focused on understanding the signals that the embryo uses to form these self-renewing cells that maintain the hematopoietic system throughout the life of the organism.

### 2. Understanding T Acute Lymphoblastic Leukemia (T-ALL) development and T-cell lymphoma.

We study the signals that regulate the generation and maintenance of normal and leukemic cells, as well as leukemic stem cells (LSCS). With this aim we have developed in vitro and in vivo experimental models that complement the analysis of patient samples.

### 3. GATA2 deficiency syndrome.

We are collaborating in an international consortium to understand the contribution of GATA2 mutations to pediatric Myelodysplastic syndrome and transformation to Acute Myeloid Leukemia (AML). We are developing humanized blood animal models of this syndrome.

### 4. Understanding cell transformation.

We work closely with the Research Group for Molecular Mechanisms of Cancer and Stemness directed by Dr. Lluís Espinosa, and we take advantage of our discoveries in hematopoietic cells to understand epithelial tissues and vice versa.

## OUR CHALLENGES

Through our research, we aim to understand the following:

1. What signals are imposed in embryonic HSCs that affect the adult hematopoietic system?

2. What are the molecular mechanisms that impose resistance to treatment in T-ALL cells?

3. What are the basic mechanisms that control cell transformation?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Embryonic hematopoiesis, T-ALL, CTCL, GATA2, Notch, NFkB, hematopoietic stem cell, leukemic stem cells





ABOUT US  
**RESEARCH  
GROUPS**

17

**ENDOTHELIAL PATHOBIOLOGY AND MICROENVIRONMENT  
LED BY MARIONA GRAUPERA**

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## OVERVIEW

Our research is devoted to study the biology of the endothelium and its role in disease towards the development of therapeutic strategies to target this compartment. Specifically, we aim to untackel the fundamental insights of vessel growth and function in developmental setting as well as to identify the pathological contexts in which the vasculature plays a critical role either intrinsically, as in vascular anomalies, or extrinsically as in cancer.

## OUR RESEARCH

Blood vessels are crucial components of every organ, as they maintain tissue homeostasis by ensuring: (i) transport of gases, nutrients, waste products and circulating cells, (ii) blood coagulation, and (iii) vascular tone and barrier. The endothelium lines the lumen of blood vessels and regulates the dynamic passage of materials and cells, whereas mural cells adhere to the abluminal surface of the endothelium and regulate vessel growth, permability and function. Both excessive and insufficient vascular network is deleterious for organisms and lead to a broad spectrum of pathologies. The overall aim of the Graupera lab is to understand the mechanisms that regulate the vasculature in development, homeostasis and disease. Most of our research has focused on the the endothelium that plays an active role in important physiological processes and diseases such congenital disorders, obesity

and cancer. Over the past decade, we have taken advantage of the PI3K pathway as a paradigm to understand how intracellular roads regulate vessel morphogenesis, and how this knowledge can be translated into therapeutic opportunities for diseases with aberrant angiogenesis. For our research, our lab develops unique animal models including, established cell lines, and patient-derived samples. We apply a holistic approach utilising state-of-the-art techniques as high-throughput analysis, next-generation sequencing, single cell RNA sequencing, phospho/proteomics, and high-resolution imaging. Our lab closely collaborates with clinicians to translate our research into the clinic at both the diagnostic and therapeutic levels.

## OUR GOALS

The Graupera lab is devoted to 5 main research lines:

- 1.** Insights on developmental vessel growth and function.
- 2.** Understanding oncoproteins-related developmental disorders.
- 3.** To study tumor-stroma interaction.
- 4.** Identify vascular therapies to treat metabolic disorders.
- 5.** To study endothelial and hematopoietic cell interface.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Endothelium, vascular compartment, homeostasis, next generation sequencing, single cell, high-resolution imaging



ABOUT US  
**RESEARCH  
GROUPS**

**18.** T-CELL LYMPHOMA  
LED BY LAURA MONDRAGÓN

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BARBERA FERRANDO, LAURA  
Lab Technician



## OVERVIEW

Our research is focused on the better understanding of the molecular mechanisms leading to T cell lymphomas appearance. We will develop our research by determining possible defective mechanisms during thymopoiesis and, by developing preclinical mice models for the study of T cell lymphomas, such as angioimmunoblastic T cell lymphoma.

With this knowledge we expect to design and validate new therapeutic treatments more specific and effective than the ones currently available in order to improve patient's survival and quality of life.

## OUR RESEARCH

T cell lymphomas can be defined as a group of malignancies caused by the uncontrolled proliferation of T cells. They constitute less than 15% of all Non-Hodgkin's

lymphomas and within this group frequency can vary enormously. Despite all being caused by T cell defective cell growth little is known about its specific origin. Besides, they present a wide variety of symptoms and clinical characteristics ranging from highly aggressive (fast-growing) lymphomas to subtypes that can develop for years without endangering the patient's life (indolent); presence of enlarged spleen, liver and/or lymph nodes; eczema and skin rash appearance; age appearance and higher incidence in men than in women. As a result, it is often difficult to establish a correct diagnosis of the disease and even more difficult to design an appropriate therapy for its specific treatment.

In this sense, our line of research aims at improving our understanding of the molecular mechanisms leading to the

defective behaviour of the T cells originating this type of lymphoma. To develop our objective we will employ immunology, functional genomics, molecular biology and medicinal chemistry techniques.

Specifically, we will make use of potential and already described genetically modified mice models as pre-clinical models for the study of different subtypes of T cell lymphomas and we will:

- Characterize their phenotype once the disease is developed in order to find the specific T cell population inducing its appearance.
- Study thymocytes maturation processes and mature T cells response to antigens to try to determine if lymphoma appearance can be already settled during thymopoiesis or once the T cells leave the thymus.

- Compare by genomic techniques like single cell sequencing the characteristics of defective thymocytes and T cells in mice models and try to find similarities in human patient's samples and databases.

With the information obtained, we aim at finding new therapeutic targets in order to:

- Develop chemical libraries screening assays for drug discovery to modulate the activity of these therapeutic targets.
- Perform structure-activity assays and possible administration in form of nanomedicines to optimize their biological activity and cellular uptake in vitro.
- Validate their use as new therapeutic strategies alone or in combination with other chemotherapeutics employing mice models of the disease.

## OUR GOALS

To make available new therapies to treat angioimmunoblastic T cell lymphoma and reduce mortality in those patients. The most important one would be to improve life expectancy of patient's suffering from this type of disease. Although, we would like to apply our research to different types of T cell lymphomas, we are initially focused on the study of molecular mechanisms leading to angioimmunoblastic T cell lymphoma. This type of disease has no specific treatment and all the strategies chosen so far have not improved patient's survival in the last 3 decades. Finding new strategies will for sure improve their chances to recover from this disease and it will significantly improve their quality of life. To unveil the molecular mechanisms leading to T cell lymphoma appearance and to provide new

therapeutic targets to design more specific and effective therapeutic treatments to fight these group of hematopoiesis diseases.

## OUR CHALLENGES

In summary, our line of research will have the final objective to provide more specific and effective therapies to treat T cell lymphoma in order to find a cure or, if not, to improve the prognosis and quality of life of patients.

- Disease Knowledge deepening
- Drug development

There are some questions we are trying to answer with our research:

## ABOUT US RESEARCH GROUPS

1

When does defects in T cells leading to T cell lymphoma appearance start?

2

Which are the specific T cell populations responsible for T cell lymphoma induction?

3

Can we design more specific and effective therapeutic treatments for this type of disease?

## KEYWORDS

thymopoiesis, t cell lineage selection, T cell receptor, T cell activation, lymphoma



ABOUT US  
**RESEARCH  
GROUPS**

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**ACUTE LYMPHOBLASTIC LEUKEMIA (ALL)  
LED BY JOSEP M<sup>a</sup> RIBERA**

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## OVERVIEW

Our research focuses on analyzing the genomic and epigenomic landscape of patients with adult ALL (acute lymphoblastic leukemia) to find out genetic alterations that predict patients' response to treatment and to identify new alternative (targeted) therapies to apply to those patients. In this way, we aim to design more personalized treatments to increase the probability of survival of ALL patients.

## OUR RESEARCH

The group's current research is divided into two main areas, according to the two main subtypes distinguished in ALL:

- **Precursor B-cell acute lymphoblastic leukemia (BCP-ALL):** BCP-ALL is the most prevalent ALL subtype and accounts for 75% of ALL cases. Although it is a highly heterogeneous disease at genetic level, different cytogenetic subtypes have been identified and, more importantly, their prognosis has been clearly established in many clinical trials. This has allowed clinicians to stratify patients according to their genetic profile to schedule intensive or less intensive treatments.
- **T-cell acute lymphoblastic leukemia (T-ALL):** T-ALL is the least common ALL subtype (25% of adult ALL cases), and the most complex and heterogeneous at genetic level, with a dismal prognosis. To improve the survival rate of patients with T-ALL, we first need to obtain detailed and relevant molecular information to accurately define the risk and thus decide on the treatment.

## OUR GOALS

We are convinced that new treatments for ALL patients can be obtained only through basic research. Therefore, our goals are:

- 1.** To identify the genetic alterations leading to treatment resistance and disease recurrence in adult ALL.
- 2.** To accurately define the risk of ALL by genetic analysis at diagnosis and relapse in order to decide on the most appropriate treatment.

## OUR CHALLENGES

Although ALL is a rare form of cancer, it has a huge impact on patients, their relatives and the health system. To find new therapies and provide new knowledge, our research hopes to:

- 1** Decipher the genetic complexity of ALL at both diagnosis and relapse.
- 2** Identify critical genetic lesions in ALL cells that could be targetable with new drugs.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Acute lymphoblastic leukemia, adults, genomic analyses, minimal residual disease, treatment resistance





ABOUT US  
**RESEARCH  
GROUPS**

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**BARCELONA ENDOTHELIUM TEAM (BET)  
LED BY ENRIC CARRERAS**

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## OVERVIEW

Blood vessel endothelium crosses each and every tissue and presents unique structural and functional properties in each vascular bed. This cellular heterogeneity is reflected by the structural and functional heterogeneity of the whole endothelium as a complex system. Due to its location, the endothelium is exposed to all kinds of physiological and pathological stimuli and constitutes the first barrier to many drug interventions.

This organ has the capacity to constantly adapt to environmental changes by modulating vasomotor tone, haemostatic balance and inflammatory reactions, among other responses. Endothelial activation could lead to an irreversible state known as endothelial dysfunction that, in a broad sense, leads to various non-adaptive alterations of the

functional phenotype that constitute a net liability to the host.

## OUR RESEARCH

Our group has extensive experience in the study of the endothelial dysfunction that develops in association with different vascular pathologies, such as the early complications associated with hematopoietic cell transplantation, obesity, chronic kidney disease, thrombotic microangiopathies and sepsis.

We also explore strategies for the protection of this endothelial dysfunction to improve patient health. In this regard, one of our main interests is to evaluate different compounds that potentially exhibit the capacity to protect the endothelium and to decipher their mechanisms of action.

## OUR GOALS

Our main lines of research are:

- 1.** To characterize the endothelial activation and dysfunction associated with cardiometabolic diseases through in vitro models.
- 2.** To elucidate the mechanisms that lead to endothelial dysfunction.
- 3.** To investigate agents with potential protective effects on the endothelium to prevent complications.
- 4.** To find soluble markers with prognostic and diagnostic value for vascular complications.

- 5.** To study complement pathways and complement deficiencies in thrombotic microangiopathies.

- 6.** To assess platelet physiology and alterations of haemostasis by using perfusion devices to explore adhesive and cohesive properties of platelets under flow conditions.

## OUR CHALLENGES

Hematopoietic cell transplantation (HCT) has been the major curative therapy for several hematological, metabolic and neoplastic disorders. However, the efficacy of this procedure is limited by life-threatening complications, the most important of which is graft versus host disease (GvHD), which has a high mortality rate. Through our research, we aim to answer the following questions:

- 1** What are the pathophysiologic mechanisms that characterize endothelial dysfunction?
- 2** How can we avoid the vascular complications associated with hematopoietic cell transplantation?
- 3** Which is the role of the complement system in vascular complications?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Endothelium, Inflammation, Diagnostic and prognostic markers, Thrombotic microangiopathies (TMA), Drugs



ABOUT US  
**RESEARCH  
GROUPS**

**21** | **FUNCTIONAL CYTOMICS  
LED BY JORDI PETRIZ**

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## OVERVIEW

The ability to bridge large-scale and single-cell approaches at a functional level is key to identifying biomarkers expressed in rare cells, and particularly in cancer stem cells (CSCs). Our interdisciplinary group investigates and develops key experimental approaches to understand the principles underlying the emergence and prevention of tumorigenesis and cancer therapy resistance, with the aim of translating biomedical science to integrated clinical practice and public health through certified transfer processes in cooperation with trusted allies and partners.

## OUR RESEARCH

Our current research projects use innovative approaches to study the expression of primitive stem cell markers during the origin, progression and maintenance of cancer and the management of cancer; the quality and safety of hematopoietic blood progenitor and stem cell grafts; the role of myeloid-derived suppressor cells in immunotherapy and targeted therapy for clinical decision-making; new cytomic strategies for whole blood and marrow immunostaining; the use of natural compounds for cancer treatment; and the accurate detection and significance of minimal residual disease in acute leukemia.

## OUR GOALS

Our goal is to provide patients with a wide range of scientific support strategies, through precision, oversight and accuracy, to achieve:

**1. Clinical implementation of functional cytomic assays,** and precision/personalized high-quality assays for individual patients by integrating functional cytomics.

**2. Translation of functional screening to novel clinical strategies.** Measurement of the impact of exogenous interventions such as drug exposure on tumor cell phenotype.

**3. An understanding of drug resistance** and prediction of effective drug combinations.

**4. A reduction in costs.** by obtaining specialized instrumentation and personnel for the execution of cytomic screening in partnership with stakeholders and biotechnological partners.

**5. Functional and immunophenotyping datasets** aimed at understanding complex functional-to-phenotype correlations, thereby accelerating discovery of the biology of leukaemogenesis and the clinical implementation of novel therapies.

## OUR CHALLENGES

Since personalized medicine aims to provide the right therapy, in the right dose, thereby improving patient stratification and disease prediction, we expect to achieve:

**1 Better diagnosis and earlier intervention.** Cytomic analysis could determine precisely whether patients are susceptible to drug toxicities and make it possible to choose the optimal treatment strategy.

**2 Individualized drug selection.** Here we consider which molecular and functional models best predict how a patient will respond to a therapy to develop accurate and cost-effective tests.

**3 Drug development challenges.** A better understanding of the clinical observations made during individualized drug development and conventional therapy will help identify new disease subtypes and their associated molecular pathways, and design drugs that target them with more efficient trials.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Cytometry, cancer stem cells, cytome, human cytome project.



ABOUT US  
**RESEARCH  
GROUPS**

22

**MYELOID NEOPLASMS  
LED BY LURDES ZAMORA AND BLANCA XICOY**

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## OVERVIEW

Genetic profiling for hematopoiesis malignancies involves chasing a moving target. Not so long ago, leukemias were stratified based on karyotype abnormalities. In recent years, however, knowledge of molecular genetics in hematology has increased significantly, thus offering new clinical opportunities. It has now been shown that gene expression, mutations and other genetic and epigenetic abnormalities also have diagnostic, prognostic and therapeutic implications.

## OUR RESEARCH

Since 2004, our group has been studying MN, with a particular focus on characterizing genetic and epigenetic lesions to find new diagnostic, prognostic and therapeutic markers that could help us better diagnose and treat patients with these diseases. First, we started with karyotype and single nucleotide polymorphism arrays (SNP-A) to help us detect alterations at chromosome level, and we are currently performing studies at gene level (mutational profile studies) and analysing the impact that telomere size could have on the development of the disease.

## OUR GOALS

The aim of our group is to apply our research to myeloid neoplasms (MN), specifically three distinct hematopoiesis diseases: chronic myelomonocytic leukemia, myelodysplastic syndromes and myeloproliferative neoplasms (PV, ET, MFP and CML), with a view to finding better tools for diagnosis and prognosis stratification and achieving individualized targeted therapies (personalized medicine). Therefore, our research focuses mainly on the following areas:

### 1. Chronic myelomonocytic leukemia (CMML).

The aim of our research is to characterize the type, frequency and prognostic impact of mutations and cytogenetic alterations detected by SNP arrays in patients with low-risk CMML and, then, study its epigenetic changes (DNA methylation and miRNAs expression).

### 2. The classification and prognosis of the group of diseases termed myelodysplastic syndromes (MDS).

This knowledge will also contribute to a better understanding of MDS biology and a better stratification of the prognosis of these patients, which would also help with the selection of the most appropriate treatment for each one.

### 3.

#### Chronic myeloid leukemia (CML).

The aim of our research is to determine whether or not we can find any genetic marker at CML diagnosis that could explain a patient's toxicity to tyrosine kinase inhibitors, or identify which patients will achieve a molecular response. We are also seeking a technique with higher sensitivity than QRT-PCR.

### 4.

#### BCR-ABL1 negative classic myeloproliferative neoplasms (MPNs).

We study several genomic changes in an attempt to associate them with cytological subtypes, laboratory parameters, clinical complications and probability of transformation to either MF or AML.

## OUR CHALLENGES

Our research is highly socially relevant because we promote capacity building, advancing knowledge, help in making informed decisions and improve the health in general terms, with economic benefits for the whole society. Through our research, we aim to answer the following questions:

### 1

How can we improve diagnosis, prognosis stratification, treatment response and life expectancy in chronic myelomonocytic leukemia?

### 2

How can we improve diagnosis, prognosis stratification, treatment response and life expectancy in myelodysplastic syndromes?

### 3

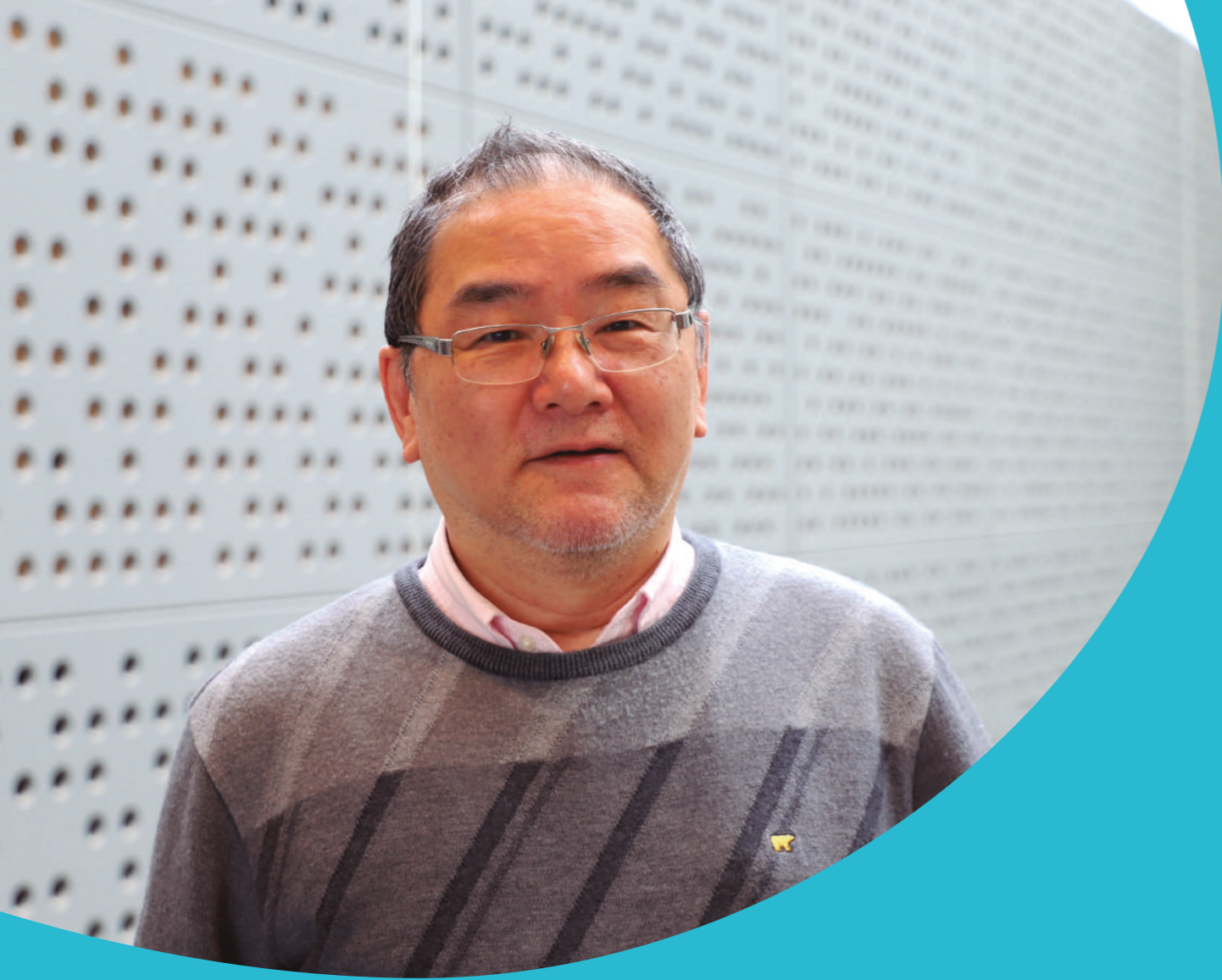
How can we improve diagnosis, prognosis stratification, treatment response and life expectancy in myeloproliferative neoplasms?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Myeloproliferative neoplasms, chronic myeloid leukemia, myelodysplastic syndromes, MPN/MDS, acute myeloid leukemia





ABOUT US  
**RESEARCH  
GROUPS**

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**IMMUNOHEMATOLOGY AND GLYCOBIOLOGY  
LED BY FUMIICHIRO YAMAMOTO**

**GROUP  
MEMBERS**

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## OVERVIEW

The ABO blood group system is one of the most important blood group systems in transfusion medicine. The ABO polymorphism consists of A and B glycan antigens on red blood cells (RBC) and antibodies against A and B antigens in the sera of individuals who do not express these antigens. In humans, A and B antigens are also present on epithelial and endothelial cells, depending on the ABO genotype/phenotype of the individual. Because of this, ABO matching is also crucial in cell, tissue and organ transplantation.

## OUR RESEARCH

We study the molecular genetic mechanisms for the expression of genetically incompatible

glycan antigens and have thus far revealed several potential mechanisms, including the appearance of FORS1 induced by the deletion of exon 3 or 4 of the AT mRNA. Because altered splicing is a hallmark of cancer, this mechanism may be responsible, at least partially, for FORS1 expression in group A and AB individuals.

We also investigate the potential mechanism by which incompatible A antigens appear in group O individuals through complementation by recombination of DNA or trans-splicing of RNA and also expression of FORS1 due to changes in specificity resulting from incorrect intra-Golgi localization of modified glycosyltransferases.

## OUR GOALS

Cancer growth indicates that the cancer cell-killing activities of natural immunity against genetically incompatible and/or cryptic glycans are ineffective and insufficient. However, they can be improved through active and/or passive immunization. Therefore, our goals are:

- 1.** To investigate the use of genetically incompatible and/or cryptic glycan antigens as molecular targets for medical intervention.
- 2.** To explore the possibility of using forced expression of genetically incompatible glycans to make cancer cells susceptible to natural immunity.

## OUR CHALLENGES

If successful, the active immunization we advocate for could revolutionize the cancer treatment landscape, thereby dramatically reducing the financial burdens on patients and society. Through our research, we aim to answer the following questions:

- 1** What is the molecular genetic/epigenetic basis of glycan alterations in cancer?
- 2** Can we use cancer-specific glycans as molecular targets for cancer detection and immunotherapy?.
- 3** Does the minitransfusion/injection of mismatched red blood cells expressing genetically incompatible and/or cryptic glycans improve humoral and cellular immunity against cancer cells expressing cancer-specific glycans?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Genetically incompatible glycan antigens, cryptic glycan antigens, cancer immunotherapy, disease susceptibility, ABO polymorphism



ABOUT US  
**RESEARCH  
GROUPS**

**24** | **LEUKEMIA STEM CELL  
LED BY RUTH RISUEÑO**

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**HUGUET I PRADELL, JULIA**  
PhD Student



## OVERVIEW

Many leukemias originate in a population of leukemic stem cells, which are responsible for initiating and maintaining the disease. Our group investigates the biology and sensitivity of stem cells that sustain tumors such as acute myeloid leukemia (AML), myelodysplastic syndromes (MDS), chronic myelomonocytic leukemia (CMML) and acute lymphoblastic leukemia T (ALL-T).

## OUR RESEARCH

Our work focuses on searching new therapeutic targets for blood disorders enabling us to identify new biomarkers that can be used for prognostic and/or diagnostic purposes, thereby gaining greater insight into the biology behind leukemic processes. New knowledge will help us develop new drugs that specifically attack the population of leukemic stem cells.

Due to the similarities between leukemic stem cells and healthy hematopoietic stem cells, our research group is striving to develop therapies that trigger the terminal differentiation of the

population of leukemic stem cells, which eliminates their capacity to initiate and maintain the disease and enhances its chemosensitivity.

## OUR GOALS

Our main goal is to understand the biology of leukemic stem cells and identify pharmacological mechanisms that will enable us to modulate their functionality and eliminate them selectively. To achieve this, we adopt a dual approach: on the one hand, we study the underlying biological mechanisms responsible for these neoplasms and, on the other, we develop new therapeutic approaches with the potential for clinical application.

## OUR CHALLENGES

Due to the biological properties of leukemic stem cells, understanding how they work and identifying the differences between these and healthy hematopoietic stem cells is essential to design new more efficient and selective therapies that can be personalized for each patient. Therefore, our research aims to:

- 1 Understand how leukemic stem cells work.
- 2 Identify therapeutic targets and develop new drugs for leukemic stem cells.
- 3 Determine the prognostic and diagnostic value of these therapeutic targets as biomarkers for leukemia.
- 4 Describe the differences between leukemic stem cells and healthy haemopoietic stem cells.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Leukemia, leukemic stem cell, drug development, hematopoiesis, differentiation therapies



ABOUT US  
**RESEARCH  
GROUPS**

**25** | **LYMPHOID NEOPLASMS  
LED BY TOMÁS NAVARRO**

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**CAÑAMERO GIRÓ, ELOI**  
Administrative

## OVERVIEW

Genetic profiling for hematopoiesis malignancies involves chasing a moving target. Not so long ago, leukemias were stratified based on karyotype abnormalities. In recent years, however, knowledge of molecular genetics in hematology has increased significantly, thus offering new clinical opportunities. It has now been shown that gene expression, mutations and other genetic and epigenetic abnormalities also have diagnostic, prognostic and therapeutic implications.

## OUR RESEARCH

Our group focuses mainly on the research of AIDS-related lymphomas (ARLs). The most frequent ARLs are diffuse large B-cell lymphoma (DLBCL), Burkitt lymphoma (BL) and Hodgkin lymphoma (HL). Plasmablastic lymphoma (PBL) and primary effusion lymphoma (LEP) are less frequent, but typically present in immunosuppressed individuals. We also study other hematopoiesis disorders with an increased incidence in the HIV-positive population such as Castleman disease (CD).

Our main areas of research are:

- **Genetic studies on HIV-related lymphomas.** Although HIV-infected patients are treated with the same regimens as HIV-negative individuals, their survival rate is lower due to the higher susceptibility to infections and secondary neoplasms.
- **Liquid biopsy in aggressive lymphomas.** This technique could be useful to diagnose DLBCL earlier, and in a more comprehensive and accurate manner than with tissue biopsy alone.
- **Genetic studies on plasmablastic lymphoma.** Plasmablastic lymphoma (PBL) is a rare B-cell lymphoid neoplasm that especially affects immunocompromised individuals and has a poor.

## OUR GOALS

We believe that genetic and epigenetic profiles will help clarify the mechanisms involved in lymphomagenesis and identify potential biomarkers, thus allowing cases to be classified more effectively. The possible diagnostic and/or prognostic impact of these markers could pave the way for the design of new targeted therapies, thus leading to new treatment approaches and improving the outcome of patients suffering from the lymphomas on which our research focuses, i.e. AIDS-related lymphomas, plasmablastic lymphoma and Castleman disease.

## OUR CHALLENGES

We expect that the results of our studies will lead to changes in the management of these rare lymphoid neoplasms and improve the poor prognosis of some lymphoid malignancies, such as plasmablastic lymphoma. Through our research, we hope to answer the following questions:

1

What genetic and epigenetic mechanisms are involved in the development of HIV-related lymphomas?

2

Which biomarkers can be used for an earlier diagnosis of lymphoid neoplasms in populations at high risk of developing these disorders, such as HIV-infected and transplanted patients?

3

How can we apply liquid biopsy in the diagnosis and follow-up of aggressive lymphomas?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Non-Hodgkin's lymphoma, Hodgkin lymphoma, HIV, Epstein-Barr virus, diagnosis, prognosis, marker, treatment, targeted therapy, early detection





ABOUT US  
**RESEARCH  
GROUPS**

**26** | **MULTIPLE MYELOMA  
LED BY ALBERT ORIOL**

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## OVERVIEW

Multiple myeloma is a cancer of plasma cells, a type of white blood cell that accumulates in the bone marrow and interferes with normal blood precursors and bone remodelling, thus causing anemia, bone lesions, renal insufficiency and recurrent infections. Between four and five out of 100,000 people are diagnosed every year. Despite the fact that treatments and prognosis have greatly improved in recent years, multiple myeloma is not yet curable. It is a recurrent disease that can leave important sequelae after each relapse.

## OUR RESEARCH

Our clinical research team participates in the main international collaborative phase I to phase III trials establishing the current standards of care, with a particular focus on the optimal combinations of agents with clinically relevant synergies.

Active trials are already focusing on the efficacy of next-generation combinations, including antibody-drug conjugates, T-cell engagers and CAR-T cells. We are interested in the identification of subjects unlikely to respond to optimized first-line strategies and, therefore, of ideal candidates for such trials with novel immunotherapeutic approaches.

## OUR GOALS

We believe that the drug combinations currently being evaluated can cure a proportion of patients with multiple myeloma. Furthermore, we believe that it should be possible to predict patients in whom such combinations are not curative so that we can promote early interventions with alternative agents, mainly based on immunotherapeutic approaches to prevent the clinical consequences of full-blown relapse and maintain a symptom-free response in patients.

On this regard, our main goals are:

- 1.** To define standards of treatment that provide a long-lasting response in most individuals.
- 2.** To identify patients who will probably be cured and will safely remain treatment-free.
- 3.** To identify patients who are unlikely to be disease-free for long with current treatments and search for alternative treatment options that can be applied before recurring disease causes organic damage.

## OUR CHALLENGES

Full-blown multiple myeloma has devastating consequences that severely reduce patients' quality of life and autonomy and represent a huge burden for caregivers and families. Therefore, the diagnosis of multiple myeloma has a dramatic impact on individuals and society. Through our research, we hope to answer the following questions:

- 1** What patients are unlikely to obtain prolonged benefits from current standards?
- 2** Would they benefit from early intervention with alternative agents?
- 3** Can we identify patients who will potentially be cured or are unlikely to relapse and safely spare them the burden of continuous therapy?

## ABOUT US RESEARCH GROUPS

### KEYWORDS

Multiple myeloma, synergistic combinations, immuno-drug conjugates, T-cell engagers, CAR-T cells.



ABOUT US  
**RESEARCH  
GROUPS**

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**MYELODYSPLASTIC SYNDROMES  
LED BY FRANCESC SOLÉ**

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## OVERVIEW

Myelodysplastic syndromes (MDS) are a heterogeneous group of hematopoiesis stem cell disorders that result in ineffective hematopoiesis, blood cytopenia, myelodysplasia and a significant risk of progression to acute myeloid leukemia (AML). MDS is one of the most common hematopoiesis malignancies in the elderly and the severity of the disease depends on a variety of biological factors that translate into a spectrum of symptoms with a profound impact on the patient's quality of life and survival; a third of MDS patients will progress to AML and the remaining two thirds will suffer from a combination of chronic anemia, recurrent infections and bleeding episodes.

## OUR RESEARCH

Our research focuses on unravelling the heterogeneity of myelodysplastic syndromes (MDS), mainly using genomic techniques. We study MDS patients who harbor a specific cytogenetic alteration: the deletion of the long arm of chromosome 5. Our aim is to improve the genetic characterization of these patients by studying the impact of adjunct cytogenetic abnormalities on their prognostic stratification; how cytogenetics and mutations can influence the response to lenalidomide treatment; the molecular landscape of MDS through next-generation sequencing techniques; and, finally, intratumoral heterogeneity before and after lenalidomide treatment using single-cell techniques.

## OUR GOALS

Through our research, we intend to contribute to a better understanding of MDS from a genomic point of view, contributing to refine the current criteria to diagnose this disease and predict patient outcomes to select the best possible treatment. Hence, our research addresses the following lines:

- 1.** Evaluating the feasibility of using peripheral blood samples to perform genetic analyses (SNP-A and NGS) in MDS.
- 2.** Monitoring mutational burden in low-risk MDS patients through the use of sequential peripheral blood samples to minimize invasive techniques on these patients.
- 3.** Genetic characterization of myelodysplastic syndromes / myeloproliferative neoplasms (MDS/MPN) to define the genetic changes that could contribute to the differential diagnosis and prognostic stratification of these patients.
- 4.** Genetic characterization of therapy-related myeloid neoplasms.
- 5.** Mechanisms of progression from clonal hematopoiesis to MDS.

## OUR CHALLENGES

Our research can translate into a more efficient use of public healthcare resources and improve the quality of life for patients. Therefore, we want to shed light on the following questions:

- 1** How might genomic techniques contribute to refining the current criteria for MDS diagnosis, prognostic stratification and treatment response?
- 2** Can peripheral blood samples be useful to monitor MDS patients through next-generation sequencing?
- 3** Could single-cell studies help us better understand intratumoral heterogeneity and clonal evolution from CHIP to MDS and TRMN (therapy-related myeloid neoplasms)?

## ABOUT US RESEARCH GROUPS

### KEYWORDS

Myelodysplastic syndromes, chronic myelomonocytic leukemia, intratumoral heterogeneity, myelodysplasia, cytopenias, CHIP, TRMN



ABOUT US  
**RESEARCH  
GROUPS**

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**STEM CELL BIOLOGY, DEVELOPMENTAL LEUKEMIA  
AND IMMUNOTHERAPY LED BY PABLO MENÉNDEZ**

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**MENÉNDEZ, PABLO**  
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**MOLINA CAMPOY, OSCAR**  
Postdoctoral Researcher

**LOPEZ MILLAN, MARIA BELEN**  
Postdoctoral Researcher

**SÁNCHEZ MARTÍNEZ, DIEGO**  
Postdoctoral Researcher

**PETAZZI, PAOLO**  
Postdoctoral Researcher

**ZANETTI, SAMANTA ROMINA**  
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**VINYOLES VERGES, MERITXELL**  
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**TIAGO CUNHA, LUIS**  
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**ROMECIN DURAN, PAOLA ALEJANDRA**  
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Lab Technician

**MARTINEZ MORENO, ALBA**  
Lab Technician

## OVERVIEW

Our group is interested in understanding the cellular origin, etiology and pathogenesis of childhood leukemia. We aim to ascertain the cell in which mutations occur and we strive to discover which cells are responsible for triggering relapses. Furthermore, we work to identify new therapeutic targets and develop more targeted, less toxic therapies. To achieve this, our laboratory uses various approaches, including genetic studies, epigenetic techniques and animal models, as well as adoptive cell immunotherapy tools.

## OUR RESEARCH

Given that acute childhood leukemia (including the B, T and myeloid variants), and childhood cancer in general, are relatively uncommon illnesses, with around 500 cases in Spain each year, it does not represent a priority target for the pharmaceutical industry. As a result, there is a serious lack of active programs that aim to identify medicines to target childhood cancer. Our group has been investigating the origin of this diseases in utero, as well as its etiological causes and physiopathological mechanisms. In 2016, we began researching non-toxic, targeted adoptive cellular immunotherapies for these children with the aim of preventing the long-term effects of current chemotherapy.

## OUR GOALS

Our group is currently involved in various lines of research in pursuit of the following objectives:

### 1. To understand the aetiology and pathogenesis of leukemia in breastfeeding infants.

To do so, we use primary samples taken from patients and develop different animal and cellular models based on prenatal (embryonic, foetal) and postnatal (neonatal and adult) stem cells.

### 2. To gain a better understanding of the role of bone marrow (BM) stroma in chemoresistance in acute myeloid leukemia (AML)

and identify new therapeutic targets for AML, which is the most common form of leukemia in adults and whose prevalence increases with age.

### 3. To improve adoptive cellular immunotherapies against ALL-B, ALL-T and AML.

To achieve this, we are searching for new therapeutic targets and developing new CARs (chimeric antigen receptors) for the different types of acute leukemia.

Our overall goal is to contribute towards curing 100% of childhood leukemias or convert them into chronic conditions, without generating lifelong toxicities.

## OUR CHALLENGES

Childhood diseases have an enormous emotional impact on the patient's whole family and everybody around them. Moreover, we must not forget that children are the future of our society, so investing in their health will benefit the future of our society enormously. Through our research, we aim to:

1 Identify the cellular origin, cellular and molecular mechanisms, and the genetic and epigenetic composition of ALL-B in breastfeeding infants.

2 Contribute to the development of new therapeutic strategies in AML targeted towards reducing the resistance mediated through the BM microenvironment and that are particularly effective against LICs.

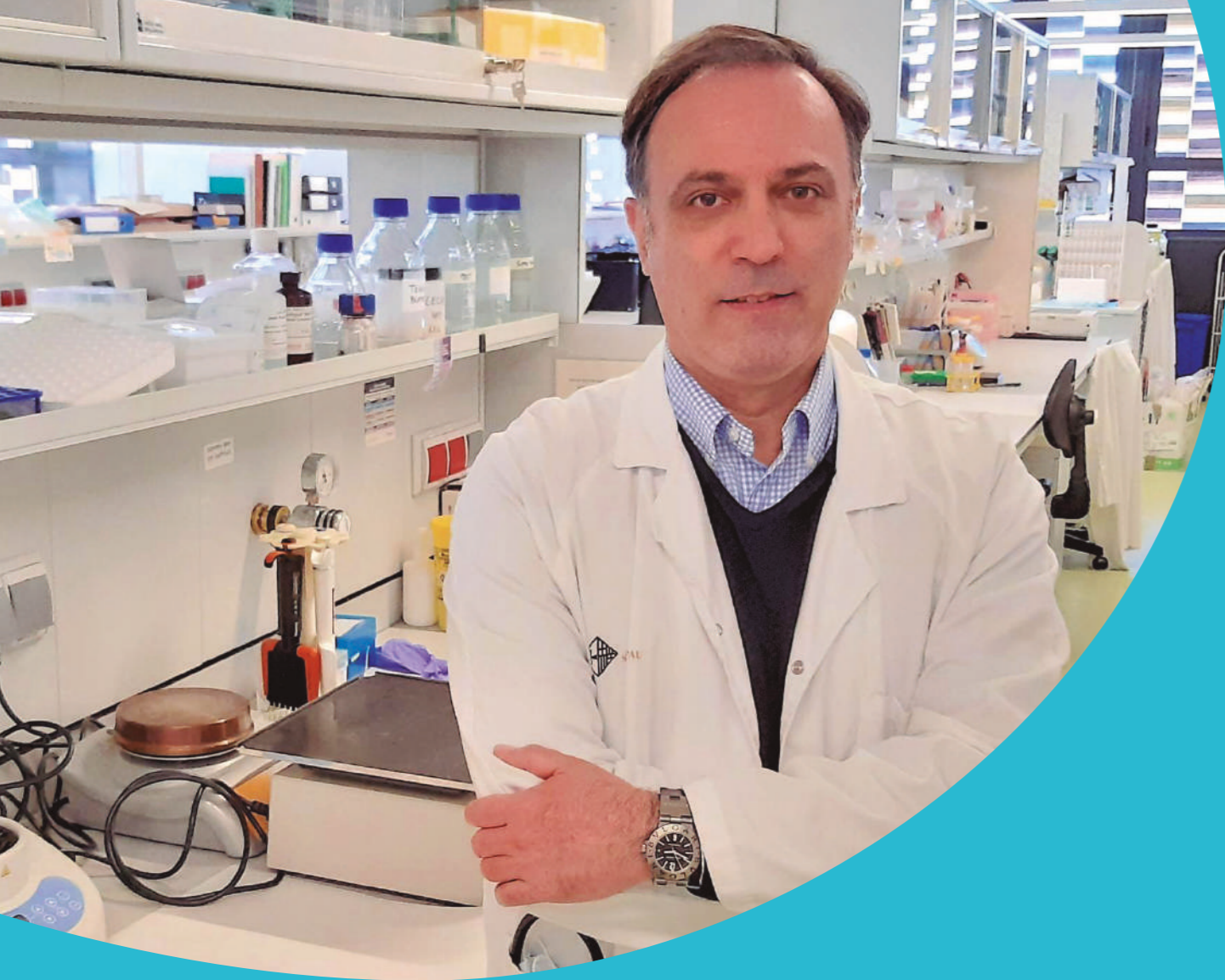
3 Develop adoptive cellular immunotherapies against ALL-B, ALL-T and AML using allogeneic T-cells without genome editing to eliminate TCR, CD3 and other molecules that play a role in immunological synapse.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Pediatric leukemia, stem cells, immunotherapy, MLL rearrangements, PDX models





ABOUT US  
**RESEARCH  
GROUPS**

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**CELLULAR IMMUNOTHERAPY AND GENE THERAPY  
LED BY JAVIER BRIONES**

**GROUP  
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Group Leader

**CABALLERO GONZÁLEZ, ANA CAROLINA**

PhD Student

## OVERVIEW

The Cellular Immunotherapy and Gene Therapy Group is focused on the study of genetically modified T-cells expressing chimeric antigen-specific receptors (CARs) and their clinical application in patients with blood cancer.

## OUR RESEARCH

Blood malignancies, like leukemia and lymphoma, are challenging diseases affecting one or more blood cell lineages. Traditional treatments include chemotherapy to deplete altered cells as much as possible, followed by bone marrow transplantation from a compatible donor, when appropriate.

The group currently focuses on studying T-cells genetically modified with chimeric antigen-specific receptors (CARs) and their clinical application in patients with blood cancer. Its current lines of research concentrate on the following aspects of cellular immunotherapy:

- Functional antitumor research into subtypes of memory T-cells.
- Study of the antitumor efficacy of memory stem T-cells genetically modified with CARs.
- Development of new CARs targeted against hematopoiesis malignancies.
- Development of clinical immunotherapy trials with CAR T-cells on patients with lymphoid neoplasms.

## OUR GOALS

Recently, a new and powerful immune cell type has been discovered called memory stem T cells. These are scarce but very special and, despite being roughly a 1% of the cells in our blood, they can find, attack and destroy cancer cells very efficiently. We have developed mechanisms to expand memory stem T cells in the lab and grow them in greater numbers, to make their action more potent and long-lasting.

Also, the genetic modification of patient-derived memory stem T cells to make them express CAR receptors, the so-called CAR-T therapy, is proving an enhanced anti leukemic potential in the clinical trials.

## ABOUT US RESEARCH GROUPS

### KEYWORDS

PCAR-T; T-Cells; Lymphoid Neoplasms



ABOUT US  
**RESEARCH  
GROUPS**

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STEM CELL TRANSPLANTATION AND CELLULAR  
IMMUNOTHERAPY LED BY ÁLVARO URBANO-ISPIZUA

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Associate Researcher



## OVERVIEW

We conduct research into cell immunotherapy treatments for patients with advanced malignant blood disorders, who tend to have a very short life expectancy. To treat such patients, we develop CAR-T and CAR-NK therapies based on adding chimeric antigen receptors (CAR) to cells of the immune system, such as T-lymphocytes and NK cells, respectively. CARs help recognize and attack tumor cells exclusively, specifically and effectively, thereby preventing an autoimmune response and reducing secondary effects on healthy cells.

## OUR RESEARCH

Former research from our group focused on the cytotoxicity mechanisms of CB-NK cells when they come into contact with MM tumor cells, in order to fight them. We discovered that they were able to regulate the cytotoxic or attack mechanisms depending on the characteristics of each tumor cell.

Later on, we started developing CAR-T cells against BCMA to treat MM patients and achieved good results. We are currently improving cell immunotherapy treatments by combining CAR-T with CB-NK cells. Moreover, we are studying how and why some tumor cells develop resistance to these therapies, with the aim of suppressing this resistance capacity.

## OUR GOALS

We are studying what happens at a molecular level between CAR-T and CB-NK cells throughout the process of recognizing, contacting and attacking tumor cells in order to identify which proteins and defence strategies are used by CAR-T, CB-NK and other cells in the immune system. Moreover, we are examining what happens within the environment of the cells when they meet tumor cells. This knowledge will help us develop better strategies to improve the efficacy of these therapies.

Through our research, we aim to achieve the best possible scenario: to cure patients and ensure that they do not relapse.

## OUR CHALLENGES

If we manage to enhance the efficacy of the CAR-T therapy and its permanence in patients to protect them from relapses, this breakthrough could be applied to patients with types of cancer other than MM. Therefore, through our research we hope to answer the following questions:

- 1 How are tumor cell resistance mechanisms against immune cells developed?
- 2 How can these tumor cell resistance mechanisms against immune cells be avoided?
- 3 How can the persistence and efficacy of CAR-T cell treatment be increased?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Multiple myeloma, B-cell malignancies, chimeric antigen receptors, T lymphocytes, NK cells, cord blood-derived NK cells, hematopoiesis malignancies, B-cell maturation antigen



ABOUT US  
**RESEARCH  
GROUPS**

**31** ■ **EPIGENETIC THERAPIES  
LED BY MARÍA BERDASCO**

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**LÓPEZ PATO, MIGUEL**  
Lab Technician

## OVERVIEW

Epigenetic therapies aim to modify the epigenome, the set of molecular processes that regulate gene expression without altering the DNA sequence and can change the course of a disease and its phenotype. There are now examples of epigenetic drugs for treating hematopoiesis malignancies approved by the United States Food and Drug Administration (FDA). However, the volume of promising preclinical evidence far exceeds the number of epigenetic research projects that have resulted in clinical applications to patients. Therefore, more translational studies that may lead to the development of more specific epigenetic drugs and more robust biomarkers are required.

## OUR RESEARCH

Our research aims to ascertain the therapeutic benefit of targeting epigenetic alterations in cancer together with the epigenetic-based stratification of patients to predict therapy response. To achieve this, we develop research based on three specific aims:

- Identification of the epigenetic alterations that act as drivers of tumor progression (“druggable epigenetic alterations”).
- Validation of epidrugs that can efficiently revert aberrant epigenomes in cancer.
- Stratification of patients based on their epigenetic profile to predict response to immunotherapy.

## OUR GOALS

We aim to develop a translational research line that focuses on elucidating the epigenetic alterations that are druggable targets in a tumor, and the means to exploit them therapeutically within the framework of precision medicine.

## OUR CHALLENGES

Through our research, we aim to help answer the following questions:

- 1 Which epigenetic alternations represent targets for drugs to treat cancer?
- 2 How can we efficiently treat tumors caused by epigenetic alternations?

- 3 Who could benefit from therapeutic strategies based on epigenetics?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Epigenetic drug, epigenetic editing, epidrug, hematopoiesis malignancies, targeted therapies





ABOUT US  
**RESEARCH  
GROUPS**

**32** | **LYMPHOMA TRANSLATIONAL  
LED BY GAËL ROUÉ**

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**GORJON DE PABLO, GEMA**  
Lab Technician

## OVERVIEW

Our research is centered on the development of innovative preclinical models of B-cell lymphoma that can be used to unravel the complex role of tumor-lymphoma crosstalk during the development of the disease and the acquisition of refractoriness in current regimens. To that end, we intend to reproduce the original composition and architecture of tumors in the laboratory to carry out a complete transcriptomic and proteomic analysis and develop new pharmacological entities in collaboration with academic experts and clinical-level pharmaceutical companies, all with a view to fostering the bench-to-bedside transfer of new and tailored therapeutic strategies.

## OUR RESEARCH

We develop in vitro and animal models with the intention of more accurately recreating the context of onset and progression of the lymphoma in each patient. In this regard, we take account of fundamental parameters that until now have been largely unexplored in preclinical research, such as the architecture of the original tumor, the components of the immune system that accompany it and the defects in the regulation of the proteins – not only of the genes – that characterize it.

## OUR GOALS

Our main areas of research are

### 1. **Development of a patient-derived xenograft platform for the evaluation of new targeted therapies in aggressive B-cell lymphomas.**

To confirm the efficacy, safety and translationality of these studies, we propose that standard in vitro assays be associated with innovative in vivo models (PDX) to allow us, firstly, to work directly with primary tumor cells to validate the most effective therapies and the most relevant biological effects while taking into account the role of the tumor microenvironment; and, secondly, to guarantee that these therapies can be translated into clinical trials with molecular and genetic determination of the factors that condition the response to the best therapies tested in each patient.

### 2. **Modulation of the lymphoid microenvironment by intrinsic protein homeostasis in aggressive B-cell lymphoma.**

We will characterize the ubiquitome of malignant B cells through proteomic profiling and correlate these intracellular complexes with the immunological pattern of each tumor model to validate the impact of tumor protein homeostasis on the development of B-cell lymphoma and the intratumoral infiltration of immune cells.

## OUR CHALLENGES

Through our research, we aim to understand the following:

- 1** To what extent intrinsic protein homeostasis can regulate the complex tumor-stroma crosstalk in different models of aggressive B-cell lymphoma.
- 2** How germinal center-derived lymphoma can be sensitized to immune checkpoint blockade therapy.

- 3** How multiomics analysis of paired treatment-naïve and therapy-refractory B-cell lymphoma can help in the design of efficient and personalized therapies.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

B-cell non-Hodgkin's lymphoma (NHL), tumor modelling, proteostasis, tumor microenvironment, immunotherapy



ABOUT US  
**RESEARCH  
GROUPS**

33

**DESCRIPTIVE AND ANALYTICAL EPIDEMIOLOGY OF CANCER  
LED BY RAFAEL MARCOS-GRAGERA**

**GROUP  
MEMBERS**

**MARCOS-GRAGERA, RAFAEL**  
Group Leader

**SOLANS MARGALEFF, MARTA**  
Postdoctoral Researcher

**AUÑÓN SANZ, CARME**  
Postdoctoral Researcher

**OSCA GELIS, GEMMA**  
Postdoctoral Researcher

**VILLAVICENCIO OBANDO, ALICIA**  
Postdoctoral Researcher

**PUIGDEMONT GUINART, MONTSE**  
Lab Technician

**SANVISSENS BERGE, ARANTZA**  
Lab Technician

**VIDAL VILA, ANNA**  
Lab Technician



## OVERVIEW

One of the main lines of research of the group is the epidemiology of hematological neoplasms, with the aim of determining the incidence, prevalence and survival of this type of cancer. The results obtained aim to provide useful and reliable information to design and/or improve the appropriate health resources and describe the population trends of this group of diseases.

## OUR RESEARCH

Hematological neoplasms are a large group of diseases with a wide variability derived mainly from the type of cells that cause it. Also, the evolution and prognosis of each histological subtype of neoplasia is very diverse and it is necessary to deepen the knowledge of each of them. In this sense, population cancer records are a reference tool for establishing the incidence and survival of each type.

## OUR GOALS

Specifically, our research objectives aim at:

- 1.** Establishing the prevalence, incidence and survival of myeloid, lymphoid and histiocytosis neoplasms globally and according to the respective subtypes.
- 2.** To analyze the temporal trend of the incidence and survival of hematological neoplasms in the context of the evolving therapeutic background.
- 3.** Determine epidemiological parameters based on sex and age.
- 4.** Carry out etiological studies of hematological neoplasms according to each of the histological subtypes.
- 5.** To study the genetic and environmental risk factors related to hematological neoplasms.
- 6.** Describe the risk factors and epidemiology of multiple myeloma based on its precursor cells.
- 7.** Analyze the associations between comorbidity and the survival of lymphoid and myeloid neoplasms.

- 8.** Evaluate the population effectiveness of new therapies in a real population and the impact on survival.

- 9.** Identify changes in the classification, definition and coding of hematological neoplasms and establish working protocols to have homogeneous tools that allow epidemiological comparisons at the international level.

## OUR CHALLENGES

Through our research, we aim to understand the following:

- 1** What is the incidence of hematological neoplasms in the territory?
- 2** What is the survival of each of the histological subtypes of neoplasia?
- 3** How have changes in the coding of hematological neoplasms over time affected the epidemiological determinants of this group of diseases?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Incidence, survival, mortality, lymphoid neoplasms, myeloid neoplasms, histiocytosis, hematological neoplasms



ABOUT US  
**RESEARCH  
GROUPS**

34

**ONCOGENESIS AND ANTITUMOR DRUGS  
LED BY RAMON MANGUES**

**GROUP  
MEMBERS**

**MANGUES BAFALLUY, RAMON**  
Group Leader

**CASANOVA RIGAT, ISOLDA**  
Postdoctoral Researcher

**UNZUETA ELORZA, UGUTZ**  
Postdoctoral Researcher

**FALGAS COMAMALA, AIDA**  
Postdoctoral Researcher

**PALLARES LOPEZ, VICTOR**  
Postdoctoral Researcher

**ALBA CASTELLÓN, LORENA**  
Postdoctoral Researcher

**SALA FAIG, RITA**  
Postdoctoral Researcher

**MEDINA GUTIERREZ, ESPERANZA**  
PhD Student

**RIOJA BLANCO, ELISA**  
PhD Student

**NUÑEZ AMELA, YAIZA**  
PhD Student

**CARRASCO DIAZ, LUIS MIGUEL**  
PhD Student

**GARCIA LEON, ANNABEL**  
Lab Technician

**NAVAS JIMENEZ, LUIS CARLOS**  
Lab Technician

## OVERVIEW

Current treatments lack selectivity towards cancer cells, which induces insufficient anticancer activity and produces severe adverse effects that limit their dosage. We are developing self-assembling protein-based nanoparticles for the treatment of hematological and solid cancers that are highly selective in targeting receptors overexpressed in cancer stem cells. They display a wide therapeutic window by avoiding renal clearance while internalizing into and selectively eliminating cancer target cells and enhancing the uptake of the payload drug into cancer tissues, with negligible uptake or toxicity in normal tissues.

We have achieved high antitumor and antimetastatic effects using apoptotic, genotoxic or

microtubule inhibitor drugs as payloads, and we are now testing novel payloads that use non-apoptotic cell death mechanisms.

## OUR RESEARCH

We generate novel protein-based nanomedicines with a high therapeutic window that tackle unmet treatment needs in acute myeloid leukemia (AML), diffuse large B-cell lymphoma (DLBCL) and colorectal cancer (CRC) using preclinical models and clinical translation, through:

- The development of animal cancer models that are resistant to current therapy or disseminated for the study of the molecular mechanisms of cancer stem cell involvement in these processes, especially in

cancer cells that overexpress the chemokine receptor CXCR4.

- The development of drug nanoconjugates or protein-only nanoparticles for intravenous injection that use receptor-mediated targeted delivery of cytotoxic agents to cancer stem cells and incorporate novel drugs or polypeptide domains that exploit the higher capacity for apoptotic induction, or the triggering of cell death mechanisms as an alternative to apoptosis.
- The development of artificial amyloid bodies for subcutaneous injection with capacity for the sustained release of therapeutic protein nanoparticles in the bloodstream that reach cancer tissues.

## OUR GOALS

Our aim is to develop nanomedicines that can effectively render cancers that have disseminated or relapsed sensitive to therapy by acquiring resistance to current therapy. In doing so, we expect to increase the cure and complete response rates, thereby leading to longer survival times.

An additional goal is to ensure that the repeated administration of these novel nanomedicines induces potent anticancer activity, while maintaining low or absent toxicity in normal tissues, associated with a lack of, or tolerable, side effects.

Finally, we also aim to develop a formulation of amyloid structured inclusion bodies whose capacity for the sustained release of therapeutic nanoparticles into the blood could be subcutaneously administered once a month. Their development will circumvent the need to administer the nanoparticles twice a week by intravenous injection, thus allowing patients to stay at home during treatment and avoid hospitalization.

## OUR CHALLENGES

Ninety percent of cancer patients die of metastases that do not respond to current treatments. Therefore, patients who develop metastases are considered incurable. Through our research, we aim to answer the following questions:

1

Is the selective elimination of cancer stem cells a relevant clinical target to improve therapy in different cancer types with acquired resistance and disseminated disease?

2

Will protein-based targeted nanoparticles that incorporate non-apoptotic and immunogenic cell death polypeptides increase cure, response and survival rates while reducing side effects once tested in patients?

3

What are the underlying mechanisms that dictate the highly selective accumulation of protein nanoparticles targeting the CXCR4 receptor we observe in cancer tissues?

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Biotechnology, nanomedicine, targeted drug delivery, oncotherapy, metastases





ABOUT US  
**RESEARCH  
GROUPS**

**35** | **CHRONIC LYMPHOCYTIC LEUKEMIA  
LED BY CAROLINA MORENO**

**GROUP  
MEMBERS**

**MORENO ATANASIO, CAROLINA**  
Group Leader

**MORA RAYA, ALBA**  
Postdoctoral Researcher

**CUELLAR GARCIA, CAROLINA**  
Postdoctoral Researcher

**JARA BUSTAMANTE, PAOLA**  
Lab Technician

## OVERVIEW

Chronic lymphocytic leukemia (CLL), the most common adult B-cell malignancy in Western countries. It is characterized by the accumulation of monoclonal CD5+ B cells with a characteristic immunophenotype in peripheral blood, bone marrow, and lymphoid tissues. The clinical course of the disease is extremely heterogeneous and, as a result, the individual life-expectancy ranges from a few years to a virtually normal lifespan. Despite the important progress in its therapy, CLL is still considered an incurable disease.

## OUR RESEARCH

Our group is constantly making hard efforts to gain further insights into the characteristics of the leukemic compartment and deepen on the impact of the tumor microenvironment. This knowledge can contribute to better understand the biologic events involved in the disease development and be used to develop new strategies for clinical management and therapy and eventually to improve the quality of life of patients with CLL.

Our group has been involved in several pivotal clinical trials that led to the approval of new drugs for CLL therapy. Also, we are involved in several projects focused on the characterization of residual leukemic cells, autoimmunity and mechanisms at work in BCR signaling.

## OUR GOALS

Our group is constantly making hard efforts to gain further insights into the characteristics of the leukemic compartment and deepen on the impact of the tumor microenvironment. This knowledge can contribute to better understand the biologic events involved in the disease development and be used to develop new strategies for clinical management and therapy and eventually to improve the quality of life of patients with CLL. Through our research, we aim to:

- 1.** Make available better diagnosis, prognosis and therapeutical strategies in CLL.
- 2.** Extend the knowledge of molecular and cellular events implicated in CLL development.
- 3.** Prevent clinical relapses in CLL.
- 4.** Design feasible approaches to monitor MRD in CLL.
- 5.** Find a cure to CLL patients.

## OUR CHALLENGES

Inherent to the disease, CLL patients have a deregulated immune system, which predispose them to have disease complications, including recurrent infections and autoimmune phenomena. The understanding of how the immune cells of tumor microenvironment contribute to maintain the survival of leukemic cells is essential to design future therapeutic strategies aimed, not only at eradicating leukemic cells, but also at restoring the immune system.

In the last years our group has started some projects aimed at describing the immune status and functional characteristics of B and T cell populations during the evolution of the disease and after therapy with targeted therapies.

## ABOUT US RESEARCH GROUPS

## KEYWORDS

Chronic Lymphocytic Leukemia, B-cell receptor signaling, minimal residual disease, liquid biopsy, tumor microenvironment



ABOUT US  
**RESEARCH  
GROUPS**

36

**HEMATOLOGY RESEARCH  
LED BY DAVID GALLARDO**

**GROUP  
MEMBERS**

**GALLARDO GIRALT, DAVID**  
Group Leader

**SANTOS CARVAJAL, NAZLY**  
Medical Doctor

**DÍAZ SANTA, JOHANA**  
Medical Doctor

**GONZÁLEZ MONTES, YOLANDA**  
Medical Doctor

**CRUZ GARCÍA, DAVID**  
Medical Doctor

**COLL JORDÀ, ROSA**  
Medical Doctor

**LLOVERAS GUELQUE, NATÀLIA**  
Medical Doctor

**RONCERO VIDAL, JOSEP M<sup>a</sup>**  
Medical Doctor

**TUSET ANDÚJAR, ESPERANZA**  
Medical Doctor

**KELLEHER, NICHOLLAS**  
Medical Doctor

**BUSTINS TARRATS, ANNA**  
Medical Doctor

**ANGONA FIGUERAS, ANNA**  
Medical Doctor

**VILA BOU, JORDI**  
Medical Doctor

**BLANCO BLANCO, ANTONIO**  
Medical Doctor

**MOSTACEDO MARASOVIC, SILVIA**  
Medical Doctor

**SITGES ARRIAGA, MARTA**  
Medical Doctor

**LLOPIS PUIGMARTÍ, XESCA**  
Nurse Case Manager

**GONZÁLEZ BÁRTULOS, MARTA**  
Lab Technician

**GONZÁLEZ BÁRTULOS, MARTA**  
Lab Technician, PhD Student



## OVERVIEW

The research group in hematology is devoted to clinical and translational trials in hematology, focused on diagnosis, prognosis and the development of new therapies to treat hematological malignancies such as leukemias and myelomas.

## OUR RESEARCH

The group uses a variety of approaches such as genetic studies of the immune response, pharmacogenomics in response to treatment, analysis of polymorphisms as disease predictors and the study of cell populations using flow cytometry for the characterization of residual disease.

## OUR GOALS

Through our research, we aim to:

- 1.** Investigate biological, clinical, and epidemiological aspects of hematological diseases.
- 2.** Carry out translational research projects focused on finding prognostic factors or treatment response predictors.
- 3.** Carry out clinical research, promoting participation in clinical trials for hematological diseases and participating in national and international cooperative groups.

## KEYWORDS

Hematology, myeloblastic leukemia, multiple myeloma, chronic lymphatic leukemia, residual disease



## ABOUT US **RESEARCH GROUPS**

# 37

**MYELOID NEOPLASMS (CAMPUS CLINIC)  
LED BY JORDI ESTEVE**

### **GROUP MEMBERS**

ESTEVE REYNER, JORDI  
Group Leader

### **OVERVIEW**

Myeloid neoplasms are a group of diseases in which the bone marrow produces an abnormal quantity of precursors for red blood cells, platelets or certain types of white blood cells. This leads to a variety of symptoms and from fatigue to bones fragility and, eventually, to a higher risk of developing Acute Myeloid Leukemia (AML).

Despite the advances produced during the last decades, not all those diagnosed benefit from efficient therapies.

Advancing in the knowledge of myeloid neoplasms is, therefore, paramount to increase both prognosis and survival of patients.

### **OUR RESEARCH**

Our research group is searching for key molecular features of myeloid neoplasms

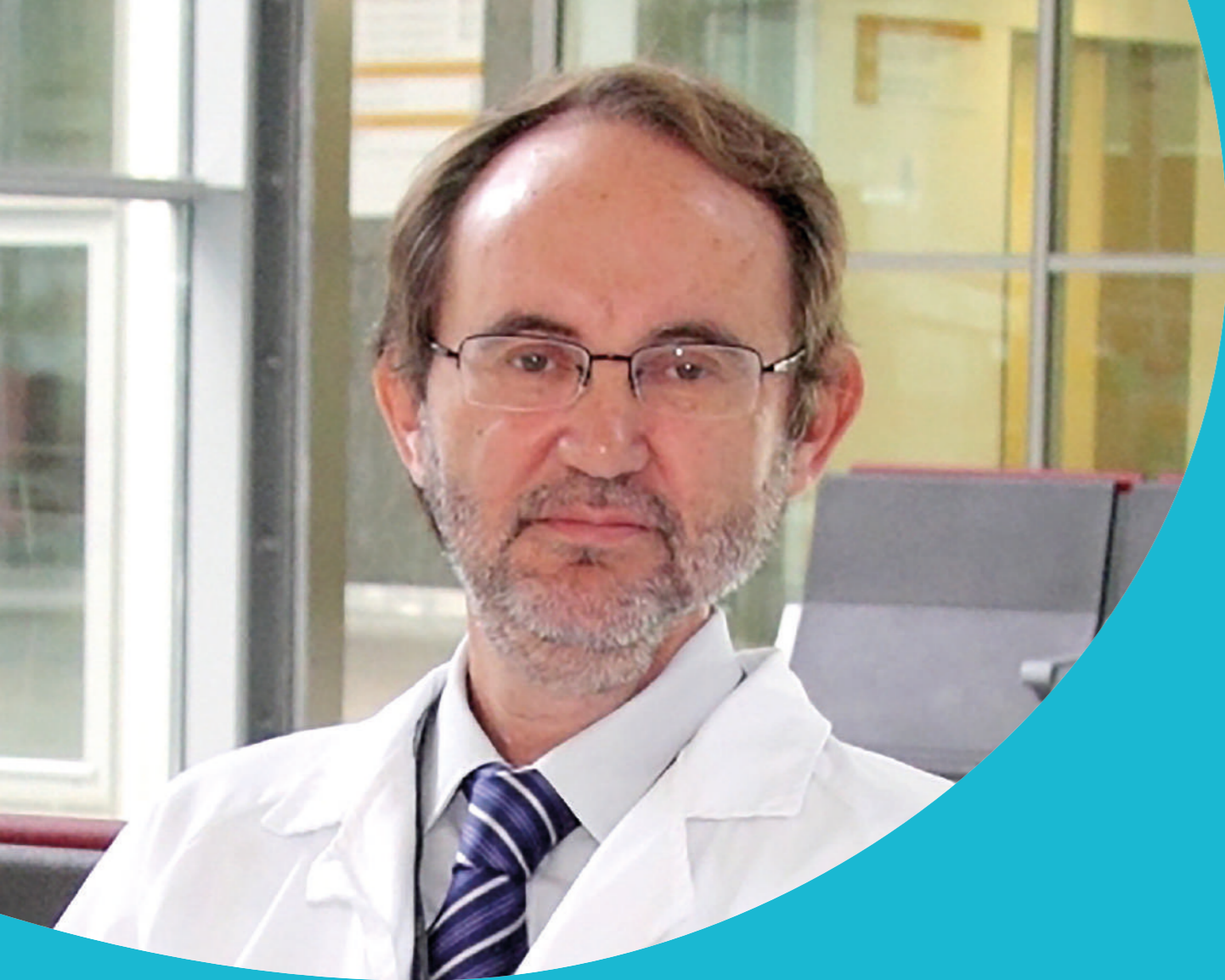
that could be used as therapeutic targets. In particular, we are focusing our efforts towards:

- Myeloma and other monoclonal gammopathies
- Mechanisms of progression in monoclonal gammopathies
- Myeloid neoplasms
- Lymphoid neoplasms

Also, we are seeking to improve the overall knowledge of the neoplasm microenvironment, the conditions where malignant cells live and proliferate, as well as how the body defenses respond to it.

### **KEYWORDS**

Pediatric leukemia, stem cells, immunotherapy, MLL rearrangements, PDX models



ABOUT US  
**RESEARCH  
GROUPS**

38

**HEMATOLOGICAL DISEASES, TRANSPLANT  
AND CELL THERAPY LED BY JORDI SIERRA**

**GROUP  
MEMBERS**

**SIERRA SIERRA, JORDI**  
Group Leader

**ALVAREZ FERNANDEZ, CARMEN**  
Postdoctoral Researcher

**ESCRIBA GARCIA, LAURA**  
Postdoctoral Researcher

**GRANELL GORROCHATEGUI, MIQUEL**  
Postdoctoral Researcher

**BRUNET MAURI, SALUT**  
Postdoctoral Researcher

**MIQUELEIZ ALAMOS, SARA**  
Postdoctoral Researcher

**GARRIDO DIAZ, ANA**  
PhD Student

**ESQUIROL SANFELIU, ALBERT**  
Lab Technician



## ABOUT US

# RESEARCH GROUPS

### OUR RESEARCH

Our research focuses on the molecular and cellular physiopathology of blood cancers, particularly on acute myeloid leukemia (LMA) and chronic lymphatic leukemia (CLL) where we seek to find new treatment options targeting molecular features. Also, we study the prognostic value

of clinical and biological features in malignant hemopathies, like LMA and CLL.

We study the transplant of hematopoietic progenitors and its complications and develop new academic CAR-T cells enriched in T-memory stem cells to treat T and B Hodgkin lymphomas.

### OUR GOALS

The main goals of our research are:

- 1.** Identify new prognostic parameters for risk and therapeutic stratifications.
- 2.** Molecularly characterize acute myeloid leukemia and determine the prognostic value of known genes and other genes of uncertain significance.
- 3.** Evaluate targeted therapy in cell lines and animal models (together with Dr. Mangues' group).
- 4.** Reduce toxicity and increase the availability of allogeneic transplants.
- 5.** Preclinical (mouse) and clinical studies on immunotherapy for lymphoproliferative diseases. Development of non-commercial CAR-T cell therapies.

Our overall goal is to contribute towards curing 100% of childhood leukemias or convert them into chronic conditions, without generating lifelong toxicities..

### OUR CHALLENGES

It is paramount that we improve the prognosis of hematological patients by using new more precise therapies, and less toxic. Therefore, through our research, we aim to:

- 1** Improve the genotypic and immunophenotypic characterization of AML and CLL, to identify new prognostic factors and administer targeted therapy.
- 2** Improve the safety and effectiveness of hematopoietic transplantation and expand the number of patients who can benefit from it.
- 3** Develop new CAR-T products that enhance the currently commercially available ones.

### KEYWORDS

Hematopoietic transplantation, CAR-T cells, Immunotherapy, Acute Myeloid Leukemia, Chronic Lymphocytic Leukemia



ABOUT US  
**RESEARCH  
GROUPS**

39

**HEMATOLOGICAL DIAGNOSIS  
LED BY JOSEP NOMDEDÉU**

**GROUP  
MEMBERS**

JOSEP NOMDEDÉU  
Group Leader

## ABOUT US

# RESEARCH GROUPS

### OUR RESEARCH

In our lab, we focus on both malignant and non-malignant hemopathologies to offer better diagnostics, understand its biological characteristics, and develop new treatments. We study acute leukemias multi-omics and platelet pathologies like thrombocytopenias, thrombocytopathies and thrombosis.

#### MALIGNANT HAEMOPATHOLOGIES

- Diagnostic activities: morphological, immunophenotyping, cytogenetic and molecular characterization of acute leukemias).
- Biological characteristics responding to therapy.
- New treatments using cell line models aimed at molecular targets.
- Genomic (microarrays) and proteomic platforms for diagnosis.
- Murine models development.

#### NONCANCEROUS HAEMOPATHOLOGIES

- Diagnosis and characterization of thrombocytopenia, thrombocytopathies and other platelet pathologies.
- In the framework of the GAIT-2 project (genetic analysis of idiopathic thrombophilia, phase 2), to seek new phenotypes that favor the development of thrombosis, specifically related to the structure and function of platelets and other blood cells.

### OUR CHALLENGES

Through our research, we aim to:

1

Consolidate the characterization of hematological tumors and complex, rare, and genetic noncancerous hematopathology's.

2

Include the results of mass-analysis genomic and proteomic platforms in diagnostic algorithms and establish prognostic factors for hematological disorders.

3

Develop functional cell culture and animal (murine) models.

4

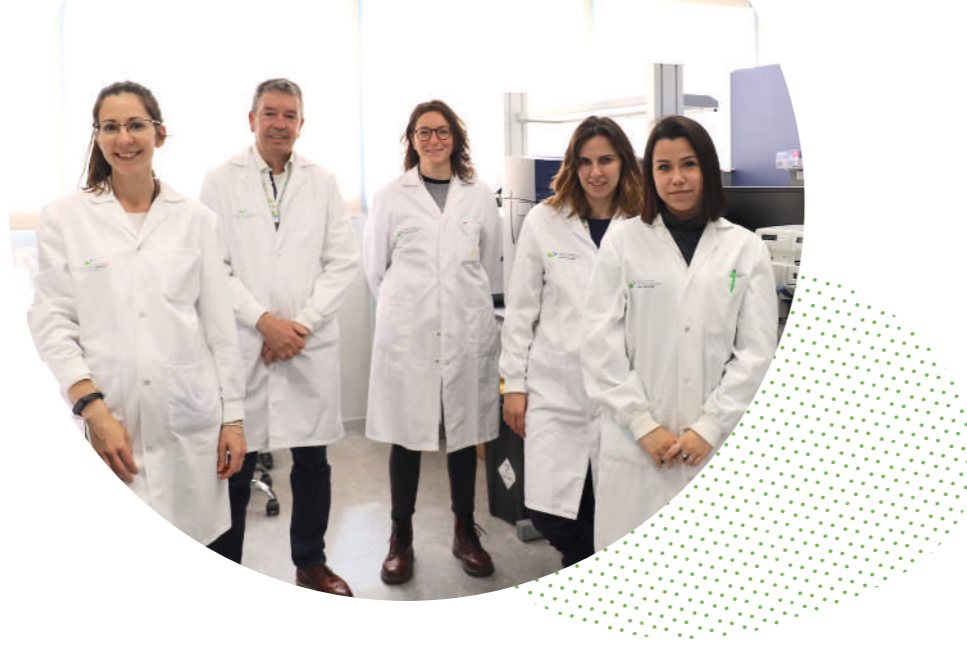
Consolidate cooperation with the GAIT-2 project, especially regarding platelet and other blood cell participation in thrombosis generation.

### KEYWORDS

Malignant hemopathologies, thrombocytopathies, thrombosis



ABOUT US  
**CORE  
FACILITIES**



MICROARRAYS  
UNIT

**UNIT MANAGER:**  
Francesc Solé

**STAFF:**  
Mar Mallo, Nuri de Haro,  
Jessica Tijero, Aida Silverio

The Microarrays Unit (UM) is a service focused on DNA and RNA microarray solutions towards a personalized medicine and participates in the Cytogenetic European Quality Assessment (CEQA).

**MOLECULAR CYTOGENETICS**

Microarray studies can offer various solutions for cytogenetic applications:

- Detection of whole genome gains and losses at a high resolution.
- Analysis of whole genome absences of heterozygosity.
- SNPs genotyping and genome-wide association studies.
- RNA Analysis Solution
- Gene expression profile studies on either human or mouse are suitable for:
- Detection of genes and pathways involved in diseases, treatment responses and biological processes.
- Predictive models based on gene expression profiles.
- Pharmacogenomics and toxicogenomics studies.
- Alternative splicing detection.
- Classification of samples on gene signatures.
- Analysis of miRNA.
- Microarray analysis on compromised samples with degraded and/or low quantity samples.
- Quality sample analysis

In addition, to the microarray procedure we also offer DNA and RNA quantification and quality control analysis.

**HIGH THROUGHPUT QPCR**

The Biomark HD system is a high throughput qPCR that runs IFCs in either real-time or end-point read modes, bringing PCR solutions to a range of applications. The 48x48 Dynamic Array combines up to 48 samples and 48 assays, generating 2304 different assays in one single run. The 96x96 Dynamic Array combines up to 96 samples and 96 assays, generating 9216 different assays. In addition, the FLEXsix IFC incorporates six 12x12 partitions that can be organized in any configuration, in up to six separate experimental runs.

**Applications**

- Genotyping
- Targeted Gene expression
- Digital PCR

**Equipment**

- Affymetrix Research Platform: GCS3000 with autoloader
- Agilent Bioanalyzer 2100
- NanoDrop 2000 Spectrophotometer

ABOUT US  
**CORE  
FACILITIES**



**CYTOGENETICS  
UNIT**

**UNIT MANAGER:**  
**Francesc Solé**

The Cytogenetics Unit is responsible for analytical tests belonging to Special Hematology from samples of whole blood, serum, plasma, urine, body fluids, bone marrow, lymph nodes, spleen, and tumor masses. The available analysis include: Special Hemostasis, Erythropathology, Special Cytology, Immunophenotype, Cytogenetics and Molecular Biology.

The Cytogenetics Unit at Josep Carreras Leukaemia Research Institute includes the Laboratory of Cytogenetics of the Institut Català d'Oncologia (ICO).

**Services**

- Conventional cytogenetics culture and karyotype performance
- Fluorescence in situ hybridization (FISH) with commercial probes
- QF-PCR



**SAMPLE  
HANDLING  
CIRCUIT UNIT**

**UNIT MANAGER:**  
**Francesc Solé**

**STAFF:**

Rocío Ruíz, Jessica Aranda,  
Aida Silverio

The Josep Carreras Leukaemia Research Institute (IJC) Campus ICO-GTP houses the Germans Trias i Pujol Hospital and Institute (IGTP-HUGTP) Sample Handling Circuit Unit, which manages the processing and storage of voluntarily donated samples of hematopoiesis neoplasms. The samples are stored in the collection entitled 'IJC Leukemia and other blood disease Sample Collection'. The IGTP-HUGTP Sample Handling Circuit Unit receives the bulk of its samples from the Catalan Institute of Oncology at the Germans Trias i Pujol Hospital (ICO-HUGTP). Samples received from other hospitals are processed in an identical way.

The technical staff of the IJC have created a database of patients, donors and samples received and processed according to required specifications for the tracking of each sample in the collection. The staff verify the quality, security and tracking of the data and samples throughout the process and starting at extraction.

ABOUT US  
**CORE  
FACILITIES**



PROTEOMICS  
UNIT

**UNIT MANAGER:**  
Carolina de la Torre

**STAFF:**  
Joan Josep Bech,  
Bernat Cucurull

The Proteomics Unit of the Josep Carreras Leukaemia Research Institute, part of the Carlos III Health Institute (ISCIII) and the Proteomics Network ProteoRed, offers mass spectrometry services to the academic and to the private sector under request.

The unit's main activity is to promote the incorporation of proteomics as a key tool for the development of clinical and basic projects at our institution. Our main work consists of offering innovative, high-quality proteomic and peptidomic services that allow the best therapeutic and human health solutions to be selected.

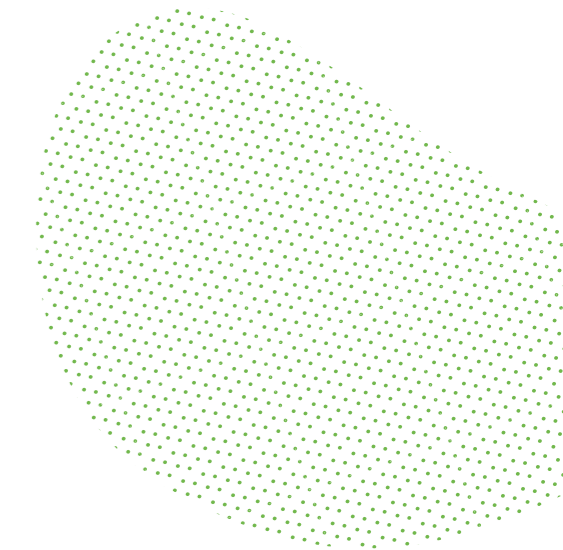
**HOW DO WE SUPPORT YOU?**

- By providing scientific and technological support to high-level research projects in the field of proteomics according to international standard procedures.
- By providing researchers with scientific advice, from the project's planning and experimental design stage to the execution phase, processing of samples and interpretation of results, and support during presentations and writing of results for publication.
- Through dissemination and training for researchers on the methodology and applications of the techniques offered.
- By contributing to the promotion of innovation in health technologies and the transfer of the knowledge generated to the public health service, and supporting genetic, epigenetic and pharmacogenetic diagnosis.

**OUR SERVICES:**

Proteomic analysis is a very powerful approach to addressing key challenges in clinical and health research. This approach can be used for different aspects of clinical and health sciences, such as biomarker discovery, drug target identification and food technology.

Biomarker discovery from a wide range of samples to improve precision medicine at different levels: a) early diagnostic, and b) prognosis to predict disease progression and guide treatment selection.





ABOUT US  
**CORE  
FACILITIES**



## GENOMICS UNIT

**UNIT MANAGER:**  
**Damiana Álvarez-Errico**

**STAFF:**  
Carles Arribas,  
Laila Aledón

The Genomics Unit aims to provide the IJC community and external researchers with scientific services. Several state-of-the-art technologies have been implemented in the unit to assess relevant genomic and epigenomic features and thus unravel basic molecular mechanisms of disease and contribute to the discovery of therapeutic targets and biomarkers.

The Unit is equipped with cutting-edge technology to offer solutions for both basic and translational epigenomic and genomic studies on many sample types (primary cells, cell lines, frozen and paraffin-embedded tissues). We have long-standing experience in array-based genome-wide DNA methylation

analysis and pyrosequencing. We use next-generation sequencing (NGS) technology to investigate subsets of genes or specific genome regions with Illumina's MiSeqDx System.

The applications provided by the Genomics Unit at IJC are:

- Infinium MethylationEPIC™ BeadChip technology (Illumina)
- MiSeqTMDx NGS sequencer (Illumina)
- PyroMark™Q48 (Qiagen)

## BIOINFORMATICS UNIT

**UNIT MANAGER:**  
**Angelika Merkel**

**STAFF:**  
Izar de Villasante

The Bioinformatics Unit at IJC provides both internal and external researchers with high-quality computational analysis services to cover all aspects of research projects related to clinical and biological data.

The Bioinformatics Unit offers a wide range of services, including experimental design for NGS or microarray experiments, statistical consulting, NGS and microarray data analysis and integration, scientific database management and software design and implementation. It also provides training on different bioinformatics topics and the use of HPC (high-performance computing) resources.



### OUR SERVICES:

- Bioinformatics data analysis, including consulting, experimental design advice, technology selection, final report generation and interpretation.
- Support for project proposal writing, data provider selection and budget planning.
- Custom bioinformatic analyses and tailored software development.
- Clinical software development compliant with ISO-13485.
- Large-volume data storage, management and submission to public repositories.
- Bioinformatics analysis training.

### BIOINFORMATICS ANALYSIS:

- Epigenomics and epitranscriptomics: 5mC and 5hmC DNA methylation arrays (infinium methylation 450K and EPIC), whole-genome bisulfite sequencing (WGBS), reduced-representation bisulfite sequencing (RRBS), bsRNA-Seq, m6A and m1A RNA methylomes by NGS and array.
- Genomics: whole-exome sequencing (WES), WGS, SNP and CNV arrays, ChIP-seq, Hi-C.
- Transcriptomics: RNA-Seq, Ribo-seq, RIP-seq, iCLIP, CLIP-seq, single-cell transcriptomics.
- Metagenomics: 16s rRNA.

ABOUT US  
**CORE FACILITIES**



The Cell immortalization Unit of the Josep Carreras Leukaemia Research Institute offers Infection of B-cells with Epstein–Barr virus (EBV) leads to more and subsequent immortalization. This is considered as the method of choice for generating lymphoblastoid cell lines (LCLs). Cell culture is an essential tool to study the fundamentals of genetic background variables. With the development of personalized medicine, this applies increasingly to the development and safety testing of drugs. Infection of B-cells with Epstein–Barr virus (EBV) leads to more and subsequent immortalization. This is considered as the method of choice for generating lymphoblastoid cell lines (LCLs). After successfully production of LCLs, different parameters including temperature, serum concentration, type of culture medium, and CO2 concentration have to be evaluated on

CELL IMMORTALIZATION UNIT

**UNIT MANAGER:**  
Carolina de la Torre

**STAFF:**  
Fernando Setien

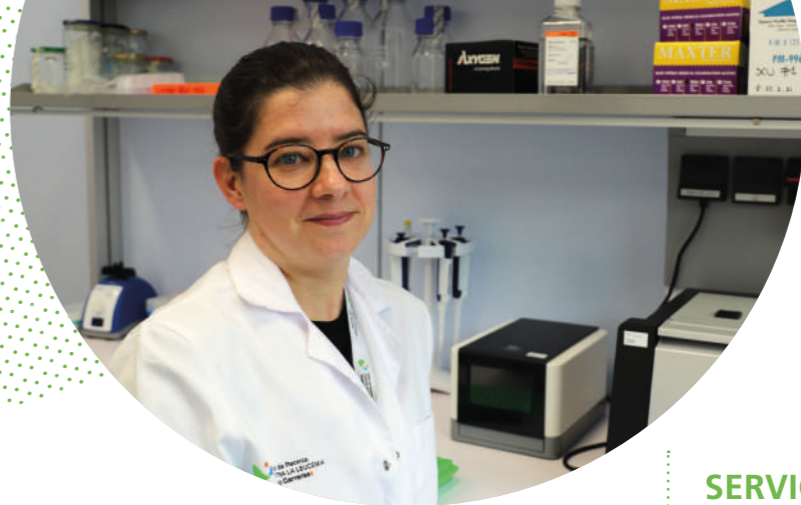
EBV-transformed B-cells. Our unit are able to produce LCLs and optimize condition.

**APPLICATIONS:**

- This immortalization technology enables rapid, efficient, and reliable production of unlimited numbers of personalized cells.
- To produce control material for rare genetic disorders.
- Lymphocyte immortalization technique let to preserve of DNA, RNA and proteins samples, that appears to be a valid strategy for further studies.
- To determine optimized condition for reliable and reproducible LCLs from different sources.
- Testing drugs analysis.
- Allows us to have enough biological sample without having to access the patient again

SINGLE CELL UNIT

**UNIT MANAGER:**  
Regina Antoni Alandes



The Single Cell Unit aims to provide scientific services to the Josep Carreras Institute community and is equipped with cutting-edge technology to apply single-cell technology to basic and translational genomic and transcriptomic studies. Single Cell approaches can reveal the cellular heterogeneity of tumors and help identify cells resistant to standard treatments or more prone to proliferate. The Single Cell Unit is equipped with a Chromium™ Controller (10x Genomics), a Mission Bio Tapestry Platform and the C1 system (Fluidigm). These technologies provide a comprehensive, scalable solution for cell characterization, gene expression profiling and DNA sequencing of up to tens of thousands of cells.

**SERVICES:**

**Chromium™ Controller (10x Genomics):**

- Single-cell RNA-seq (gene expression)
- Single-cell RNA-seq (gene expression) + Feature Barcoding
- Single-cell immune profiling (gene expression + TCR/BCR)
- Single-cell ATAC-seq

**Tapestry Platform (Mission Bio):**

- Single-cell targeted DNA-seq (mutation analysis)
- Single-cell targeted DNA-seq (mutation analysis + CNV analysis)
- Single-cell DNA-seq + cell-surface protein analysis

**C1 System (Fluidigm):**

- Single-cell mRNA sequencing
- Single-cell targeted gene expression
- Single-cell miRNA expression profiling





## ABOUT US MANAGEMENT UNITS



### STRATEGY DIRECTOR AND MANAGING DIRECTOR

**Garrido Anglada, Ana**

Together with Dr. Manel Esteller, Ana Garrido is part of the management team and contributes to the development of general policy and strategic planning, enabling and translating scientific vision and strategic objectives into a clearly articulated operational strategy.

#### The main objectives of the Strategic and Managing Director are:

- Strategic and operational organization of the Institute.
- Management, in accordance with the marked guidelines of the governing bodies, human resources, hiring staff, incidents, separation and termination of employment contracts, HR organizational policies, as well as the management of scholarships and grants, encouraging policies of Corporate Social Responsibility and attracting and retaining talent.
- Detection of the needs derived from the activity of the Institute Research Groups and the rest of the units from the economic and management point of view.
- Implementation of an agile model focused on researchers that favours their performance and the best concentration in scientific activity.



## ABOUT US

# MANAGEMENT UNITS

### Research Grants Unit

Its objective is the attraction of public and private competitive funding, both nationally and internationally, as well as the proactive management of research projects granted.

We support researchers throughout the life cycle of projects, from the detection of opportunities, the preparation of proposals and the training of research consortiums, to the management of projects in all areas beyond the economic.

#### Group members

**MORALES CUÑADO, CRISTINA**  
Project Manager

**PADIAL MELIÁN, VERÓNICA**  
Project Manager

**LAGUNAS VILA, LAIA**  
Project Manager

**EL BOUHALI EL MALLEM, SANAE**  
Project Manager

**GIL GUIÑON, ESTEL**  
Project Manager

**MAURI SÀNCHEZ, SARA**  
Project Manager

**VILLANUEVA DELGADO, ANAÍ NOEMÍ**  
Project Manager

**DOLSET VILLALOBOS, SARAI**  
Projects Support

**MORAL ROS, MARC**  
Projects Support

**ALLUÉ SANCHEZ, MÓNICA**  
Projects Support

**BEN FATINA, OMAR**  
Projects Assistant

**GARCIA MONTERO, ADRIÀ**  
Projects Assistant

**FERNANDEZ CAMBRAS, DANIEL**  
Projects Assistant

**EL-GHAUCHE EL-HALLAK, RANIA**  
International Business Development Manager

### Innovation Unit

The aim is to promote and boost innovation to improve the quality of life of society from the transfer of knowledge and developed technologies.

#### Group members

**RIERA GUERRA, ANNA**  
Innovation Manager

**FARRÉS ÀLVAREZ, CLARA**  
Innovation Officer

### Economic Management Unit

The Economic Development Manager manages and coordinates the financial control of the Institute. Its main functions are linked to the control and supervision of finances and, ultimately, it acts as a link between the Institute's management and accounting.

#### Group members

**BOIX MONTEMAYOR, HEURA**  
Economic Development Manager

### Human Resources Unit

Its mission is to plan, organize and execute all processes relative to the professional development of the staff and their commitment with the organization, within the framework of current regulations, including training opportunities and occupational risk prevention.

#### Group members

**JANÉ MORALES, NÚRIA**  
HR Manager

**PEREZ RODRIGUEZ, FERNANDO ROBERTO**  
HR Manager

**CARMONA FERNÁNDEZ, CAROLINA**  
HR Officer

**BODAS LOZANO, ANTHONY**  
HR Support

**ESCOBAR LÓPEZ, MAJDEL KAROLINA**  
HR Assistant

## ABOUT US

# MANAGEMENT UNITS

### Finance Unit

Its mission is to keep track of the actual finance situation of the Institute, rigorously and transparently, to support data driven strategic decision making in the short, medium and long run.

#### Group members

**CALONGE CORTÉS, MARIA CRISTINA**  
Financial Manager

**FINESTRES MARTINEZ, XAVIER**  
Finance Officer

**VILANOVA CUADRA, YAIZA**  
Finance Officer

**MURE FERNANDEZ, MIREIA**  
Finance Officer

**MATOS SILVA, AWILDA**  
Finance Support

**LAMATA SALVA, MARC**  
Finance Support

### Purchasing Unit

The unit's aim is to optimize purchasing at the institutional level according to the legal framework for public research bodies, to be more efficient, fast and agile, avoid unnecessary costs and save resources for research.

#### Group members

**JIMENEZ FERRER, NEUS**  
Purchasing Manager

**VERGÉS COLOMINAS, ANNA**  
Purchasing Officer

**REYES IBORRA, LAIA**  
Purchasing Officer

**GARCIA ESCODA, ALBERT**  
Purchasing Officer

**GERBOLÉS LÓPEZ, ANNA**  
Purchasing Officer

**MONTserrat SANCHEZ, QUIQUE**  
Purchasing Officer

**NIUBÓ BALCELLS, NURIA**  
Purchasing Support

**DESPRETZ SANZ, YANN**  
Purchasing Assistant

### IT Unit

The unit's objective is to support the Institute's staff in the use and purchasing of IT components -hardware, software and systems- and keep the institutional IT systems online and secure while ensuring its efficiency.

#### Group members

**JUBANY LÓPEZ, MARC**  
Manager

**GALLIGANI, DAVID**  
IT Officer

**CONTRERAS PEÑA, FRANCISCO**  
IT Officer

**SALA MOLAS, ELISENDA**  
IT Officer

**GALLARDO PEREZ, DIANA**  
IT Support

**DIAZ LOPEZ, SERGI**  
IT Assistant

### Data Management Unit

Aimed at delivering and maintaining all necessary infrastructure to efficiently keep institutional data available at all levels: strategic, technical, administrative and for transparent accountability in front of local or international management bodies.

#### Group members

**CARRIO REIG, MARTA**  
Data Manager

**DE HIGES ALBERICH, PAU TADAYUKI**  
Data Technician

### Communication Unit

Passionate about spreading the latest discoveries of our scientists and bring their research efforts closer to society, in any form. The unit strives to keep our partners closer and updated, and to foster the staff's sense of belonging.

#### Group members

**DÍAZ LÓPEZ, HELENA**  
Communication Manager

**BADAL SOLER, MARTI**  
Communication Officer

**BERZOSA FERNÁNDEZ, BEA**  
Communication Assistant

## ABOUT US

# MANAGEMENT UNITS

### Lab Management Unit

The aim is to support researchers in their daily work in laboratories so that their research can be carried out with the best equipment, in the best state and with maximum safety.

#### Group members

**PEREZ LADAGA, ALBERT**

Lab Manager

**RODRIGUEZ DE GUZMÁN GALLEGO, NOELIA**

Lab Management Assistant

**SAEZ GONZALEZ, CARLOS**

Lab Management Technician

**GARCIA FERRAN, ALBA**

Lab Management Technician

### Legal Services

Its mission is to support the Institute in all legal matters, particularly in establishing the collaboration framework with partner institutions, ongoing clinical trials and innovation aspects in biomedical research.

#### Group members

**LAFARGA TRAVER, JOSEP LLUIS**

Legal Services

### Facilities Unit

The aim is to support as a facilitator, manager, and organizer for the proper functioning of all the sites of the Institute in terms of their resources and associated with the development of our activity.

#### Group members

**ARQUÉ COMAS, CARLES**

Facilities Manager

### Support Unit

The Support Unit supports the governing bodies of the Institute in daily administrative tasks, agendas as well as in travel procedures.

#### Group members

**MARIN MANZANERA, ESPERANZA**

Management Assistant

**IZQUIERDO SÁNCHEZ, IRMA**

Management Administrative

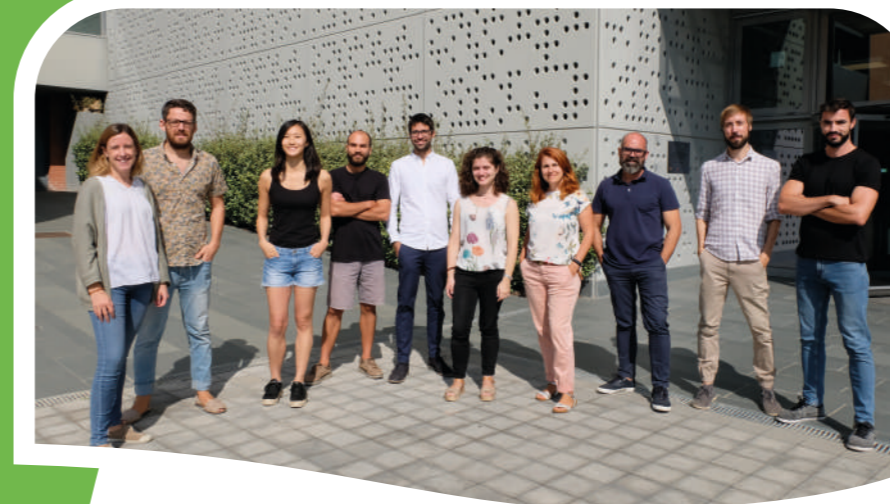
**AMADO BALLANO, ERIKA**

Management Assistant





COMMUNICATION  
**SELECTED PRESS  
RELEASES**



JANUARY:

**New epigenetic targets identified for the potential treatment of multiple myeloma.** Researchers at the Josep Carreras Leukaemia Research Institute have identified epigenetic alterations in multiple myeloma and have tested a compound that reverse those alterations, opening the door to a new therapeutic drug in the treatment of multiple myeloma.



FEBRUARY

**Myelodysplastic Syndrome heterogeneity revealed using single cell technology.** Researchers from the Myelodysplastic Syndrome group at Josep Carreras Leukaemia Research Institute describe the diversity of cancer cell populations coexisting in patients of the disease, using single cell resolution for the first time.



COMMUNICATION  
**SELECTED PRESS RELEASES**

**#100tífiques**  
 STEAMem la ciència  
 11 de febrer de 2021  
 Trenquem la bretxa de gènere a l'escola en les matèries STEAM!  
 www.100tífiques.cat

FEBRUARY

**Researchers from Josep Carreras Institute participate in the program 100tífiques aimed at fostering science careers among school girls.** Four young women researchers will talk about their research at 6th course in primary school or 1st ESO to bring the science research world closer to kids and offer relevant female roles in the science and technology field.



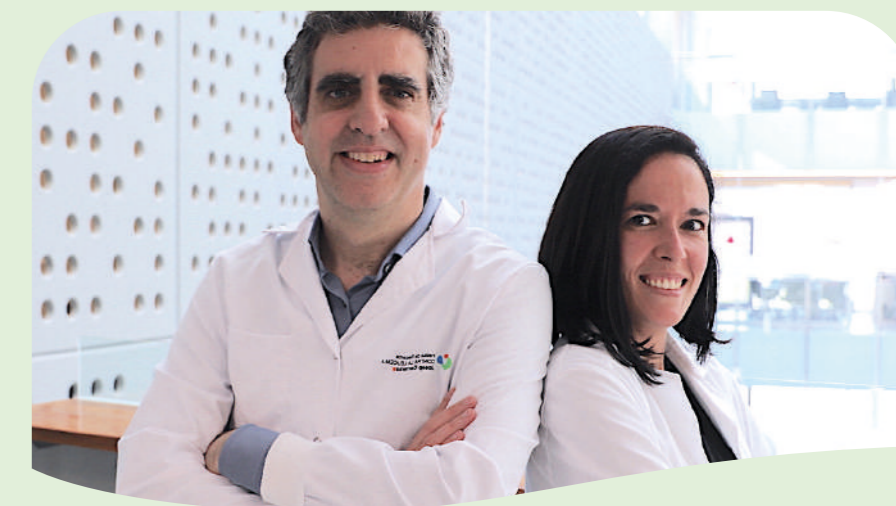
MARCH

**Sant Pau administers the first CAR-T of its own production for Hodgkin and non-Hodgkin lymphoma T in Europe.** The first CAR-T immunotherapy drug produced entirely in Sant Pau (academic) and administered to the first patient is part of a pioneering clinical trial in Europe of CAR-T phase I / II immunotherapy for the autologous treatment of classical Hodgkin lymphoma and relapsed / refractory CD30 + non-Hodgkin T CD30 + lymphoma, funded by the Carlos III Health Institute and the Josep Carreras Leukaemia Foundation.



MARCH

**The Josep Carreras Leukaemia Research Institute, new Member Institution at the International Cancer Proteogenome Consortium of the National Cancer Institute.** The Josep Carreras Leukaemia Research Institute signs a memorandum of understanding with the National Cancer Institute of the National Institutes of Health (NIH).



MAY

**Manel Esteller and Laura Belver elected as new International Members of the American Society of Hematology.** Dr. Manel Esteller and Dr. Laura Belver, Director and Group Leader of the Josep Carreras Leukaemia Research Institute, have been elected as International Members of the American Society of Hematology (ASH) by vote of the Executive Committee, the governing body of the Society. This membership is granted to researchers outside North America that have made a contribution to the field of hematology.

## COMMUNICATION SELECTED PRESS RELEASES



### JUNE

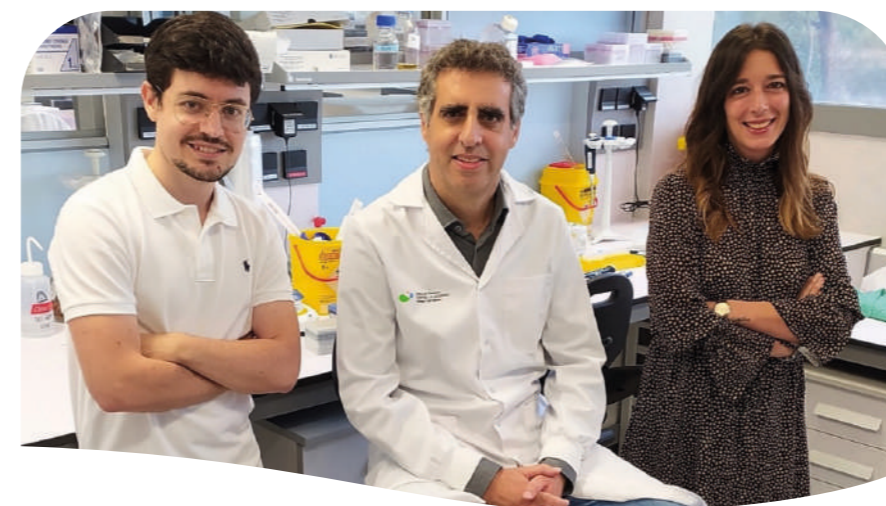
**Dr. Josep Maria Ribera coordinates the 4th ALL PETHEMA Workshop to update knowledge about Acute Lymphoblastic Leukemia.** Most of the Spanish experts in acute lymphoblastic leukemia (ALL) met on May 26 and 27 at the 4th Workshop dedicated to this disease organized by the PETHEMA group, belonging to the Spanish Society of Hematology and Hemotherapy (SEHH).

### Incidence and trends of haematological malignancies in Spain, 2002-2013



### SEPTEMBER

**REDECAN publishes the latest data on the incidence of hematological neoplasms in Spain.** Dr. Rafael Marcos-Gragera, member of the Josep Carreras Leukaemia Research Institute and leader of the Girona Cancer Registry group of the Institut Català d'Oncologia / Pla director d'oncologia, has coordinated the report published by the Spanish Network of Cancer Registries (REDECAN) on the incidence of hematological neoplasms that predicts the diagnosis of 26,000 new cases in 2021.



### OCTOBER

**Epigenetics predicts CAR-T efficacy. Epigenetics predicts the efficacy of T-lymphocyte treatments in hematological malignancies.** One in five cancers affects blood cells and lymph nodes, causing leukemias and lymphomas, respectively. Although their treatment with drugs has led to a great advance in their cure, there are cases where there is no clinical response or resistance to them is generated.



### NOVEMBER

**Manel Esteller receives the Narcís Monturiol Medal for scientific and technological merit from the Generalitat de Catalunya.** Manel Esteller, Director of the Josep Carreras Leukaemia Research Institute, ICREA Researcher and Professor of Genetics at the University of Barcelona, has received the Narcís Monturiol Medal at the scientific and technological merit from the Generalitat de Catalunya.



## COMMUNICATION SCIENTIFIC DISSEMINATION



Excellence is the basis of our Institute and encourages us to seek new initiatives to disseminate the results and impacts of our research to society. It is precisely for this reason that bringing the research of the Josep Carreras Institute closer to the public is one of the fundamental objectives of the strategic plan.

During the year 2021, and still under Covid-19 restrictions, the Institute deploys a whole range of dissemination actions. These actions are designed both to bring research closer to the public, and to position the Josep Research Institute as a pole of scientific excellence among the public.

In this spirit, we created a new informative video content called "Young researcher, what's up?", published on our YouTube channel, in which our young scientists explain their research projects in Catalan, Spanish and English.

In addition, we participate in the main scientific dissemination events of Barcelona and Catalonia through talks and workshops. These actions are indicated in the list below:

■ **Barcelona Science Festival**, organized by the City Council through the Barcelona Institute of Culture (ICUB): talk by Dr. Gerardo Ferrer and Epigenetics workshop by members of the Cancer Epigenetics group.

■ **UB Science Festival** (online), organized by the University of Barcelona, where Dr. Esteller is a professor: we presented a video about the different levels of organization of

information in a cell, done by several researchers.

■ **European Research Night**, organized by the Catalan Association for Scientific Communication with funding from the European program MSCA: talks in schools by various researchers and Workshop on Epigenetics at Cosmocaixa by the Dissemination Specialist of the Institute.

■ **Science Week 2021**, organized by the Catalan Foundation for Research and Innovation (FCRi): round table on new strategies in the fight against leukemia at the headquarters of the FCRi, with researchers from the Institute.

■ **#100tifiques**, organized by the FCRi and the BIST: networking and talks in schools by researchers from the center on the day of Women and Girls in Science, February 11th.

In addition, all the Conferences and Seminars organized by the Institute have been virtualized and streamed online, either for free on our channels or under registration. These talks are delivered by national and international speakers and have the aim to facilitate access to the latest developments in leukemia research.

The Josep Carreras Leukaemia Research Institute has maintained its collaboration with the Josep Carreras Foundation, with whom we have close ties. Every year, the Josep Carreras Foundation celebrates the Unstoppable Day and the Week against Leukemia and they organize many different activities for patients, relatives and civil society. Due to the COVID-19, those events were delivered fully online as TV shows and IJC researchers participated with a rich program of presentations, experiences,

testimonials and other online activities through their social networks.

This year, the Institute could not open its doors in the Can Ruti Open Day, organized annually at the same time as the National Science Week. We have instead participated in the online Can Ruti Open Day with micro-talks on Cancer Epigenetics, Aging and Leukemia, and also delivered an online class for High-School students on cancer research.

Finally, in the light of the new single cell analysis technologies emergence, the Institute has decided to open a specific channel on Twitter to disseminate the actions of the professionals of the center who exploit these new tools in the fight against leukemia. In this channel we also look at the benefits of the single cell analysis technologies and how they can complement classical methodologies.



# FACTS & FIGURES

## INNOVATION

The **Josep Carreras Institute** is committed to bringing the cure of leukemia and other hemotological malignances to patients. To do so and contribute to society and country's progress, it fosters the generation of new knowledge together with the development of new therapies and technologies.

In 2021, the results of our research in terms of innovation were as follows:

## FACTS & FIGURES INNOVATION

### INDICATORS 2021



**68**

Active Innovation Projects



**2**

Active Spin-Offs:  
Leukos Biotech and One Chain Immunotherapeutics



**68**

Patent families Portfolio  
(40% Licensed)



**7**

Proof of Concept Competitive calls granted:  
CaixaImpulse Validate; Líneas estratégicas (MICIN); IP BOOSTER (EC), Gínjol (i-CERCA), IESE-Tech Transfer Group



**10**

349k€ in granted PoC calls

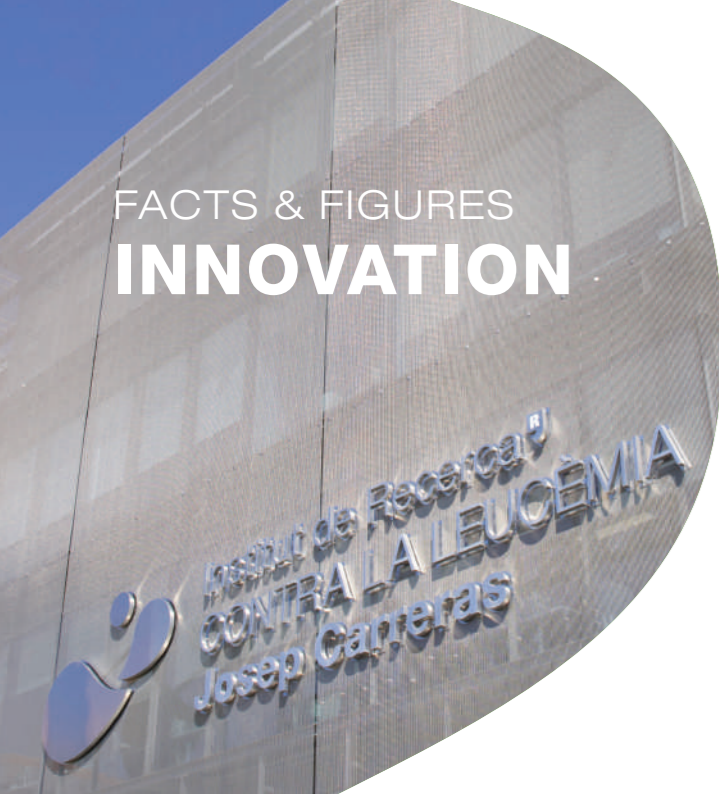


**106**

New Agreements signed in 2021



FACTS & FIGURES  
**INNOVATION**



The Josep Carreras Institute is betting on an innovative sponsorship tool

In 2021 The Josep Carreras Institute has made a bold proposal of Industrial sponsorship through a very uncommon tool in public research institutions of tax lease. Thanks to the collaboration with the company "Andamur" and the firms "AdastraVentures" and "LKS Next", the Institute has raised more than 300.000€ in 2021 (in a research project valued for more than 1M€) with the idea to consolidate and reinforce this sponsorship structure through the next years. The Institute has generate a new stable way of fundraising in order to reinforce their own research.

Regarding innovation, the great goal to be completed during 2021 is the implementation of the anointed Agrupacions d'Interès Econòmic (AIE), with the patronage of the company Andamur and the collaboration between AdastraVentures and LKS Next. In this way, the Institute has allocated more than €300,000 for the METILPIK21 project: "Study of the methylome in overgrowth syndromes caused by oncogenic mutations in PIK3CA", led by Dr. Mariona Graupera and Dr. Manel Esteller.

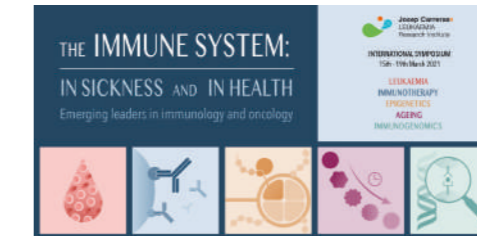
As far as technology transfer is concerned, the Institute has executed a license for the commercialization of a diagnostic kit.

During the year 2021, the Institute has managed 68 innovation projects, 10 families of patents and 2 spin-offs: Leukos Biotech, which is part of the research of the group of Dr. Ruth Muñoz Risueño and OneChain Immunotherapeutics, of the research of the group Dr. Pablo Menéndez. Overall, the set of actius innovation projects has resulted in the signing of 106 agreements worth 1.3 million euros. Additionally, during 2021, they have only tendered 50 competitive adjustments for the identification innovation projects, for a total value of 11 million euros, of which they face 14 pending resolutions and 9 of grants with a value of €600,000.

In 2021, specific financing has been obtained for valorization and concept proofs: CaixaImpulse Validate; R&D projects in strategic lines, in public-private collaboration; Intellectual Property Booster; I-Cerca, call for CERCA Gínjol patents fund, IESE-Tech Transfer Group.

## International congresses

In 2021, the Institute organized and participated in 4 international congresses, which were adapted to the global situation and virtualized.



March 15-19th, 2021  
**"The Immune System: in Sickness and in Health"** Congress.



April 7-9th April 2021  
**"Personalized and Precision Medicine"** Congress, (Participation of IJC).



Sept. 22-24th, 2021  
**"Physiology and Function of Histone Variants"** Congress.



Oct. 6-8th, 2021  
**"Vascular Malformations: from Fundamental Biology to Therapeutic opportunities"** Congress.



## FACTS & FIGURES

# TEACHING AND TRAINING

## Distinguished and Invited lectures

Twice a month the Institute has the pleasure to receive national and international well-renowned researchers in the cancer research-related field. They deliver a 1-hour lecture on their research, career and findings, which is open to all the Institute and the scientific community

### LECTURES at the Josep Carreras Leukaemia Research Institute

1st SEMESTER 2021

- JAN 29,** IGNACIO VARELA  
**Invited Lecture**  
**Instituto de Biomedicina y Biotecnología de Cantabria (IBBT), Spain**  
"Molecular characterization of the role of SWI-SNF alterations in lung cancer development"
- FEB 26,** KAREN VOUSDEN  
**Distinguished Lecture**  
**The Francis Crick Institute, United Kingdom**  
"The manifold functions of p53 in tumor suppression and beyond"
- APRIL 16,** DAN LANDAU  
**Distinguished Lecture**  
**Cancer Genomics & Evolutionary Dynamics at Weill Cornell Medicine and the New York Genome Center, USA**  
"Charting normal and malignant differentiation topologies via single cell multi-omics"

- FEB 12,** SÒNIA GUEDAN  
**Invited Lecture**  
**Clinic Hospital and IDIBAPS, Catalonia, Spain**  
"CAR-T cell therapy: engineering immune cells to fight cancer"
- FEB 19,** HANS CLEVERS  
**Distinguished Lecture**  
**Hubrecht Institute for Developmental Biology and Stem Cell Research and at the Princess Máxima Center for Pediatric Oncology, The Netherlands**  
"Organoids to model human disease"
- MARCH 5,** MARISOL SOENGAS  
**Invited Lecture**  
**Spanish National Cancer Research Centre (CNIO), Spain**  
"Immune suppressive pathways in cancer identified by live imaging of metastatic niches"
- MAY 28,** CAROLINE DIVE  
**Distinguished Lecture**  
**Cancer Research UK Manchester Institute, United Kingdom**  
"Liquid biopsies in lung cancer for improved patient outcomes"

### LECTURES at the Josep Carreras Leukaemia Research Institute

2nd SEMESTER 2021

- SEP 10,** LEE KRAUS  
**Distinguished Lecture**  
**UT Southwestern Medical Center, Texas, USA**  
"PARPs and ADP-ribosylation in Cancer: From Nucleosomes to Ribosomes"
- SEP 17,** NÚRIA MONTSERRAT  
**Invited Lecture**  
**Institute for Bioengineering of Catalonia (IBEC), Barcelona, Spain**  
"Engineering human pluripotent stem cells to understand human development and disease"
- OCT 1,** ANDRÉS HIDALGO  
**Invited Lecture**  
**Centro Nacional de Investigaciones Cardiovasculares (CNIC), Madrid, Spain**  
"Immune trafficking and the bone marrow"

- OCT 22,** SALVADOR MACIP  
**Invited Lecture**  
**Department of Molecular and Cell Biology, University of Leicester, UK**  
**Faculty of Health Sciences, Universitat Oberta de Catalunya, Barcelona**  
"Understanding resistance and relapse in B cell malignancies: a tale of mutation, oxidation and senescence"
- NOV 19,** MARK DAWSON  
**Distinguished Lecture**  
**Peter MacCallum Cancer Centre, Melbourne, Australia**  
"Cancer Epigenetics: Concepts, Challenges and Therapeutic Opportunities"
- NOV 19,** IANNIS AIFANTIS  
**Distinguished Lecture**  
**NYU School of Medicine, New York, USA**  
"Mitochondrial dynamics in leukemia drug response and the acquisition of therapy resistance"

- DEC 17,** IDO AMIT  
**Distinguished Lecture**  
**Weizmann Institute of Science, Rehovot, Israel**  
"The power of ONE: Immunology in the age of single cell genomics"



## TEACHING AND TRAINING

### YOUNG RESEARCHERS SEMINARS (39)

The Young Researchers Seminars are 20-min talks given by our PhD Students and young Postdoctoral Investigators, in which they explain an aspect of their research to their IJC fellows and respond to their questions. This is the perfect opportunity for them to practice an activity that they will have to face not only in their thesis defense, but also on numerous occasions throughout their research career.

### THESIS READ (12)

### CURRENT DOCTORAL THESIS (141)

### TRAINING COUSES

Researchers from the Institute periodically offer highly-specialized sciences courses.

#### FEBRUARY 24-25<sup>TH</sup>

- Cytogenetics and Molecular Genetics of Hematopoiesis Neoplasms

#### MAY 17-19<sup>TH</sup>

- Next Generation Diagnosis in Leukemia

#### NOVEMBER 19-20<sup>TH</sup>

- Preceptorship. Advances in Acute Lymphoblastic Leukemia

## FACTS & FIGURES

# TEACHING AND TRAINING

### COURSES AND SEMINARS

#### JANUARY

- **“Determinants of immunological evasion and immuno checkpoint inhibition response in cancer: the genetic front”** Montse Sanchez-Cespedes. Cancer Genetics Group, IJC
- **“Distinctive biomarker features in the endotheliopathy of covid-19 and septic síndromes”** Marta Palomo. Barcelona Endothelium Team (BET)
- **“Molecular characterization of the role of SWI-SNF alterations in lung cancer development”** Ignacio Varela, Instituto de Biomedicina y Biotecnología de Cantabria (IBBT), Spain

#### FEBRUARY

- **“The germline genome shapes the immune response and somatic landscape in human tumors.”** Eduard Porta. Cancer Immunogenomics Group, IJC
- **“The manifold functions of p53 in tumor suppression and beyond”** Karen Vousden, The Francis Crick Institute, United Kingdom
- **“Charting normal and malignant differentiation topologies via single cell multi-omics”** Dr. Dan Landau, Cancer Genomics & Evolutionary Dynamics at Weill Cornell Medicine and the New York Genome Center, USA
- **“Application of personalized medicine in patients with adult T-cell acute lymphoblastic leukemia (T-ALL).”** Eulalia Genesca. Acute lymphoblastic Leukemia (ALL) Group, IJC
- **“CAR-T cell therapy: engineering immune cells to fight cancer”** Sònia Guedan, Clinic Hospital and IDIBAPS, Catalonia, Spain

#### MARCH

- **“Organoids to model human disease”** Dr. Hans Clevers, Hubrecht Institute for Developmental Biology and Stem Cell Research and at the Princess Máxima Center for Pediatric Oncology, Nederland
- **“Effects of mutations, environmental exposures and drug interventions on gene expression”** Tanya Vavouri. Regulatory Genomics Group Leader, IJC
- **“Symposium: The Immune System: in sickness and in Health”** International emerging leaders in the fields of immunology and leukemia

#### APRIL

- **“New therapeutic approaches for acute leukemias”** Ruth Risueño, Leukemia stem cell Group, IJC
- **“PEMED 2020 Scientific Committee”** CHAIRMAN Prof. Manel Esteller, Director, Josep Carreras Leukaemia Research Institute

(IJC), Spain STEERING COMMITTEE Prof. Ann K. Daly, University Medical School, United Kingdom; Prof. Matthias Schwab, IKP and Dept of Clinical Pharmacology, Univ Hospital of Tuebingen, Germany; Prof. Anne-Marie Caminade, Coordination Chemistry Lab of Toulouse and University of Toulouse, France

- **“Immune suppressive pathways in cancer identified by live imaging of metastatic niches”.** Marisol Soengas, Spanish National Cancer Research Centre (CNIO), Spain
- **“The Genome in the Fourth Dimensional Context: Deciphering the role of Noncoding Mutations in Blood Diseases”** Biola M. Javierre, 3D Chromatin Organization Group, IJC
- **“A New Horizon in Precision Oncology – Proteogenomics”** Director, Office of Cancer Clinical Proteomics Research. National Cancer Institute, National Institutes of Health.

#### MAY

- **“ABO blood groups: from basics to molecular genetics/genomics, enzymology, evolution and disease association”** Fumiichiro Yamamoto, Immunohematology and Glycobiology Group, IJC
- **“Next Generation Diagnosis in Leukemia”** Online Platform
- **“Insights in endothelial cell biology”** Mariona Graupera. Endothelial Pathobiology and Microenvironment Group, IJC
- **“Mechanisms of disease progression in chronic lymphocytic leukemia”** Francesc Bosch. Vall d’Hebron Institut d’Oncologia (VHIO)
- **“Liquid biopsies in lung cancer for improved patient outcomes”** Caroline Dive, Cancer Research UK Manchester Institute, United Kingdom

#### JUNE

- **“Clinical translation of AML genomics”** Jordi Esteve, Myeloid Neoplasms Group. IJC
- **“Tissue-specific and time-dependent mechanisms of metastasis”** Roger Gomis, Growth Control and Cancer Metastasis, IRBBarcelona.

#### SEPTEMBER

- **“PARPs and ADP-ribosylation in Cancer: From Nucleosomes to Ribosomes”** Lee Kraus. UT Southwestern Medical Center.
- **“Engineering human pluripotent stem cells to understand human development and disease”** Prof. Nuria Montserrat. Institute for Bioengineering of Catalonia (IBEC)



## FACTS & FIGURES

# TEACHING AND TRAINING

### COURSES AND SEMINARS

#### OCTOBER

- **"Immune trafficking and the bone marrow"** Prof. Andrés Hidalgo, Centro Nacional de Investigaciones Cardiovasculares (CNIC)
- **"Understanding resistance and relapse in B cell malignancies: a tale of mutation, oxidation and senescence"** Prof. Salvador Macip. University of Leicester, UK and Universitat Oberta de Catalunya, Barcelona.
- **"Cytokines, Cell Crosstalk & Epigenetic Dysregulation in Immune Disease"** Prof. Esteban Ballestar, Epigenetics and Immune Disease Group, IJC.

#### NOVEMBER

- **"Different approaches to understand T-ALL chemotherapy resistance: a role for b-catenin"** Prof. Anna Bigas, Stem Cells and Cancer, Josep Carreras Institute
- **"Cancer Epigenetics: Concepts, Challenges and Therapeutic Opportunities"** Prof. Mark

Dawson. Peter MacCallum Cancer Centre, Melbourne, Australia

- **"Mitochondrial dynamics in leukemia drug response and the acquisition of therapy resistance"** Prof. Iannis Aifantis. NYU School of Medicine
- **"Novel markers and prognostic factors in HIV-related lymphomas"** Prof. Tomàs Navarro, Lymphoid Neoplasms group, IJC

#### DECEMBER

- **"Transcriptional control of B cell development and associated malignancies"** Prof. Maribel Parra, Lymphocyte Development and Disease Group, IJC.
- **"Metabolic heterogeneity as a driver of stem cell fate and tumorigenesis"** Prof. Carlos Sebastian, Metabolic Dynamics in Cancer, UB
- **"The power of ONE: Immunology in the age of single cell genomics"** Prof. Ido Amit. Weizmann Institute of Science.

## INSTITUTIONAL EVENTS

### MANAGEMENT RETREAT 25<sup>TH</sup> NOVEMBER 2021

The Josep Carreras Institute's Management Retreat served to create a shared vision among all staff categories within the organization. At this retreat, which was held at the Masia Cal Riera, our management staff collaborated with colleagues, engaged in meaningful leadership activities and built management team excellence.



### SCIENTIFIC RETREAT 27<sup>TH</sup> NOVEMBER 2021

The Institute's Scientific Retreat was held at the Montanyà Hotel & Lodgen, and participants talked about the past, present and future of the Josep Carreras Institute. It was the perfect opportunity for our researchers to share their insights about the work done during the year and gain a deeper knowledge of the projects of their closest colleagues.

## FINANCIAL DATA

The Institute's pioneering mixed-funding model is partially financed by the Josep Carreras Foundation. It also receives core funding from the Catalan government and is reliant on competitive funding for its research activities. In 2021, there was a **71,91%** increase in income from public funds and the provision of services. With respect to spending, this increased by **74,95%** compared to the previous year.

	2020	2021	
<b>INCOMES</b>	<b>11.716.754</b>	<b>16.292.754</b>	<b>71,91%</b>
CONTRIBUTIONS FROM THE GENERALITAT	3.644.424	3.587.060	
OTHER TRANSFERS (FIJC)	870.000	750.000	
SERVICES	1.037.549	3.033.238	
PROJECT	5.109.269	7.417.512	
OVERHEADS	1.055.512	1.504.945	
<b>OPERATIONAL EXPENSES</b>	<b>10.755.270</b>	<b>14.349.820</b>	<b>74,95%</b>
STAFFING COSTS	3.806.854	4.026.115	
INFORMATION TECHNOLOGIES SERVICES	155.850	161.536	
COMMUNICATION	9.211	31.938	
BUILDING MAINTENANCE	720.273	949.738	
LABORATORIES MAINTENANCE	114.049	214.312	
RESEARCH SUPPORT	277.880	26.273	
PROJECT	5.109.269	7.846.698	
SCIENTIFIC-TECHNICAL SERVICES (Platforms)	197.975	363.967	
BIOBANK	15.308	16.565	
MANAGEMENT SUPPORT SERVICES	225.595	263.925	
OTHER	81.482	67.841	
VAT PRORATA	26.812	80.454	
EXPENDITURE ON INVESTMENTS PENDING ACTIVATION			
HERITAGE		2.342	
REIMBURSEMENT OF SUBSIDIES AND OTHER MANAGEMENT LOSSES	14.711	298.116	
<b>RESULT OF THE ACTIVITY</b>	<b>961.484</b>	<b>1.942.934</b>	
<b>EXTRAORDINARY RESULT</b>	<b>7.494</b>	<b>0</b>	
<b>OPERATING INCOME</b>	<b>968.978</b>	<b>1.942.934</b>	
<b>FINANCIAL PERFORMANCE</b>	<b>-39.386</b>	<b>-1.107.638</b>	
<b>RESULT BEFORE AMORTIZATION</b>	<b>929.592</b>	<b>835.296</b>	
Amortization	-1.468.728	-1.527.277	
<b>RESULT</b>	<b>-539.136</b>	<b>-691.982</b>	



Dr. Josep Maria Ribera



Dr. Manel Esteller



Dr. Laura Belver

## Appointment

of **Dr. Josep Maria Ribera** as **Corresponding Academician of the Royal Academy of Medicine of Catalonia**. In the presentation, Dr. Ribera reviewed his academic and professional career in the field of acute lymphoblastic leukemia as a member of the Spanish Hematology Treatment Program (PETHEMA), and described the progress made in recent decades in the diagnosis and treatment of this disease, in both pediatric and adult patients.

## Appointment

of **Dr. Manel Esteller** and **Dr. Laura Belver** as **International Members of the American Society of Hematology (ASH) by the vote of the Executive Committee**, the governing body of the Society. This affiliation is awarded to researchers outside of North America who have made significant contributions to the field of hematology.

## Appointment

of **Dr. Manel Esteller** as a **member of the European Academy of Sciences**, for his important contributions to the Biomedical Sciences. Particularly due to his discoveries in the epigenetics of cancer, which have advanced the knowledge of this disease and opened the door to the application of new diagnostic tests and the development of new treatments for it.

## Award

of the **Fernández-Cruz Award for Excellence in Biomedical Research**, from the Fernández-Cruz Foundation, to **Dr. Esteller**.

## Recognition

of **Dr. Esteller** as **Highly Cited Researcher 2021 for Clarivate Web of Science**. Dr. Esteller received this honor for his work in Epigenetics, a field that studies all chemical modifications in biological material (DNA, RNA, and proteins) that, by regulating gene activity, provide cellular, tissue, and organic identity.

## Award

of the **Narcís Monturiol Medal for scientific and technological merit from the Generalitat de Catalunya to researcher Manel Esteller**, director of the Josep Carreras Leukaemia Research Institute, and leader of the Cancer Epigenetics group. The research of Dr. Esteller has made it possible to understand the contribution of epigenetic alterations to human disease, especially cancer, and to develop biomarkers of it as well as new treatments.





## COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

# CANCER AND LEUKEMIA EPIGENETICS AND BIOLOGY PROGRAM (PEBCL)

## 1. Cancer Epigenetics

### 2019 European Commission, MSCA-RISE-2019 Marie Skłodowska-Curie Actions – Innovative Training Networks

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 872391

**Title:** DevelOpmeNt of Cancer RNA TherapEutics

**Period:** 01/05/2020 - 30/04/2024

### 2019 Instituto de Salud Carlos III, Contratos Sara Borrell

**PI:** ESTELLER BADOSA, MANEL

**Reference:** CD19/00272

**Title:** Desarrollo de la isobutyl-deoxyxyboquinone en Glioma Pontino Intrínseco Difuso: una oportunidad terapéutica en oncología pediátrica

**Period:** 15/03/2020 - 14/03/2023

### 2019 Fundación Privada Olga Torres, Beques de recerca postdoctorals

**Researcher:** JOSHI, RICKY

**Title:** Studying the Clinical Impact of the Epitranscriptome for Colorectal Cancer Initiation, Progression and Treatment

**Period:** 01/01/2020 - 31/12/2021

### 2018 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la incorporació de personal investigador postdoctoral al sistema català de ciència i tecnologia

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 2018 BP 00250

**Title:** Senescence as a key factor in leukemogenesis and bone marrow homeostasis in aging

**Period:** 03/12/2019 - 31/08/2021

### 2019 European Commission, H2020 - MSCA - IF-2019 Marie Skłodowska-Curie Actions - Individual Fellowship

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 896403

**Title:** Senescence as a key factor in leukemogenesis and bone marrow homeostasis in aging

**Period:** 01/09/2021 - 31/08/2023

### 2016 Fundació La Marató de TV3, Marató 2016: Ictus i lesions medul·lars i cerebrals traumàtiques

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 201711.31

**Title:** Proyecto EPIGENESIS: Estudio epigenético y genético combinado con integrómica de datos y análisis funcional para encontrar genes asociados con el deterioro neurológico después de un ictus isquémico

## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

**Period:** 01/06/2019 - 31/12/2021

**2018 Ministerio de Ciencia, Innovación y Universidades , Acciones de dinamización "Redes Investigación"**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** RED2018-102471-T

**Title:** Metalofármacos Multifuncionales para Diagnóstico y Terapia

**Period:** 01/01/2020 - 31/12/2021

**2018 Fundación Científica de la Asociación Española Contra el Cáncer, Accelerator Award 2018**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** A29372/GEACC19003CED

**Title:** PREDICT-Meso: PRE-malignant Drivers Combined with Target-Drug validation in Mesothelioma

**Period:** 01/04/2020 - 30/03/2025

**2017 Fundación Científica de la Asociación Española Contra el Cáncer, Accelerator Award 2017**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** A26825/GEACC18004TAB

**Title:** ACRCELERATE: Colorectal Cancer Stratified Medicine Network

**Period:** 02/08/2019 - 31/10/2023

**2017 Fundación Científica de la Asociación Española Contra el Cáncer, Investigador AECC**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** INVES208DAVA

**Title:** EPINMUNE: Identificación de biomarcadores epigenéticos de predicción de respuesta a inmunoterapia

**Period:** 01/06/2019 - 30/11/2021

**2018 Ministerio de Educación, Cultura y Deporte, Ayudas para la formación de profesorado universitario (FPU)**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** FPU2017-02423

**Title:** Epigenética y Epi-transcriptómica del cáncer: el papel de las modificaciones del ARN en cáncer

**Period:** 01/07/2019 - 31/07/2022

**2018 MINISTERIO DE CIENCIA INNOVACIÓN Y UNIVERSIDADES, Retos Investigación**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** RTI2018-094049-B-I00

**Title:** Epigenetic and Genetic Disruption of RNA Modifications in Cancer (EPIRNA)

**Period:** 01/01/2019 - 31/12/2021

**2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 2017 SGR 1080

**Title:** Acreditación como "Grupo Consolidado Reconocido" al Grupo de Epigenética del Cáncer

**Period:** 01/01/2017 - 30/09/2021

**2019 Ministerio de Ciencia, Innovación y Universidades , Retos Colaboración**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** RTC2019-006951-1

**Title:** Immunotherapeutic Approach to the Treatment of Non-Hodgkin Lymphoma Through Preclinical Development and First Clinical Trials of a Selective HDAC6 Inhibitor

**Period:** 01/01/2020 - 31/12/2022

**2019 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** PRE2019-089958

**Title:** Disrupción epigenética y genética de las modificaciones del RNA en cáncer

**Period:** 01/08/2020 - 31/07/2024

**2020 Instituto de Salud Carlos III, Infraestructura de Medicina de Precisión asociada a la Ciencia y Tecnología (IMPACT)**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** IMP/00009

**Title:** Infraestructura de Medicina de Precisión asociada a la Ciencia y Tecnología

**Period:** 01/01/2021 - 31/12/2023

**2021 Fundación Uno Entre Cien Mil , VIII Beca "Fundación Unoentrecienmil" 2021 para la investigación en el área de la leucemia infantil**

**PI:** ESTELLER BADOSA, MANEL

**Title:** B-ALL troubling epigenetic dysregulation might represent a new opportunity for therapy

**Period:** 15/07/2021 - 14/07/2023

**2020 Ministerio de Universidades, Ayudas complementarias destinadas a beneficiarios de ayudas de Formación del Profesorado Universitario**

**Researcher:** BUENO COSTA, ALBERTO

**Reference:** EST21/00361

**Title:** Estancia en el Well Cornell Medicine Center, Department of Pathology and Laboratory Medicine (Dr Dan Landau), de Nueva York

**Period:** 09/11/2021 - 10/01/2022

**2021 Departament de Salut, Generalitat de Catalunya, Subvencions per a la contractació de Personal de Suport als Grups de recerca**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** SLT017/20/000218

**Title:** Contractació d'un tècnic especialista para realitzar protocols de seqüenciació de cèl·lula única

**Period:** 19/07/2021 - 31/12/2023

**2021 Fundació La Marató de TV3, Marató TV3: COVID-19**

**Researcher:** FERRER AGUILAR, GERARDO

**Reference:** 202131-32

**Title:** Multisystemic inflammatory syndrome associated with COVID-19 in children (MIS-C): genetic, epigenetic and immunopathogenic bases

**Period:** 23/09/2021 - 22/09/2024

**2020 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts a Doctorats Industrials**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** 2020 DI 20

**Title:** Novel algorithms for disruptive 3D microscopy data processing, visualization and analysis, using massive GPU parallelization and Artificial Intelligence (AI) applied to cancer epigenetics

**Period:** 01/11/2020 - 31/10/2023

**2019 Departament de Salut, Generalitat de Catalunya, Fons Europeus desenvolupament 2019**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** IU16-016723

**Title:** Construcció, habilitació i adquisició d'equipaments d'edificacions singulars de recerca centrades en la cura de les leucèmies i d'altres hemopaties malignes

**Period:** 01/01/2018 - 31/03/2023

**2021 IESE Business School, 2021 IESE Technology Transfer Group**

**Title:** Creation of an independent, non-profit biomedical research center dedicated to the study of the human genome through the immortalization of lymphocytes.

**Period:** 01/09/2021 - 31/12/2021

**2016 Instituto de Salud Carlos III, Incorporación de nuevas áreas temáticas y nuevos grupos CIBERR**

**PI:** ESTELLER BADOSA, MANEL

**Reference:** CB16/12/00312

**Title:** Center for the Biomedical Research Network in Oncology

**Period:** 01/01/2017 - 31/12/2022

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

## 2. Cancer Genetics

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** 2017 SGR 721

**Title:** Grup de Gens i Càncer, Grup de Recerca Consolidat

**Period:** 01/01/2017 - 30/09/2021

### 2019 Fundación Científica de la Asociación Española Contra el Cáncer, Investigador AECC

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** INVE19045ROME

**Title:** The CESAR Therapeutic Strategy (Cancer Epigenetic Short-circuit Adapted Response)

**Period:** 01/12/2019 - 30/11/2023

### 2017 Ministerio de Ciencia, Innovación y Universidades , Programa Estatal de I+D+i Orientada a los Retos de la Sociedad

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** SAF2017-82186-R

**Title:** Functional dissection of the MYC/MAX and SWI/SNF pathways regulation to boost the design of novel epigenetic cancer therapeutics

**Period:** 01/06/2019 - 30/09/2021

### 2018 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** PRE2018-084624

**Title:** Disección funcional de las vías moleculares MYC/MAX y SWI/SNF para potenciar el desarrollo de nuevas terapias epigenéticas en cancer

**Period:** 01/05/2020 - 30/06/2023

### 2020 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la contractació de personal investigador novell (FI-2021)

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** 2021 FI\_B 00315

**Title:** Inactivación genética de moléculas involucradas en represión transcripcional: análisis funcional y rol en el desarrollo del cáncer de pulmón

**Period:** 01/05/2021 - 30/04/2023

### 2020 Asociación Española de Afectados de Cáncer de Pulmón, Convocatorias Becas Dulce López – Fabrizio Facchini

**PI:** SANCHEZ CESPEDES, MONTSE

**Title:** Disección de los mecanismos genéticos y moleculares subyacentes a la recaída a los

tratamientos basados en inhibidores de RET en pacientes con cáncer de pulmón

**Period:** 01/01/2021 - 31/12/2021

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** SANCHEZ CESPEDES, MONTSE

**Reference:** PID2020-114541RB-I00

**Title:** Inactivation of molecules involved in transcriptional repression: functional analysis and role in lung cancer development. (TRARECAN)

**Period:** 01/09/2021 - 31/08/2024

### 2021 European Commission, Intellectual Property Booster 2021 - 1

**PI:** SANCHEZ CESPEDES, MONTSE

**Title:** KDM Subfamily 6 Protein Inhibitor for Use in the Treatment of Cancer

**Period:** 22/03/2021 - 22/06/2021

### 2021 Asociación para la Investigación del Cáncer de Pulmón en Mujeres, becas ICAPEM 2021 Pros Simón, Eva

**Title:** Disección de los mecanismos genéticos y moleculares de resistencia a los inhibidores tirosina quinasa en adenocarcinomas de pulmón de mujeres no fumadoras portadores de reordenamientos de ret

**Period:** 16/11/2021 - 31/12/2023

## 3. Chromatin Biology Laboratory

### 2019 European Commission, H2020 - MSCA - IF-2019 Marie Skłodowska-Curie Actions - Individual Fellowship

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** 895979

**Title:** Mechanisms of Sirtuin-dependent regulation of immunity and leukemogenesis

**Period:** 01/12/2020 - 30/11/2022

### 2017 Worldwide Cancer Research, April 2017 grant round

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** 18-0404

**Title:** Dissecting the specific contribution of Sirtuin ADP-ribosyltransferase and deacetylase activities in tumorigenesis

**Period:** 01/08/2019 - 31/10/2021

### 2017 Ministerio de Ciencia, Innovación y Universidades , Programa Estatal de I+D+i Orientada a los Retos de la Sociedad

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** SAF2017-88975-R

**Title:** Protección de la estabilidad del genoma por sirtuinas en condiciones de estrés y sus implicaciones en cáncer y envejecimiento

**Period:** 01/06/2019 - 30/06/2021

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** 2017 SGR 00148

**Title:** Grupo de Biología de la Cromatina, Grup de Recerca Consolidat

**Period:** 01/01/2017 - 30/09/2021

### 2020 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la contractació de personal investigador novell per a l'any 2020. FI-DGR 2020

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** 2020 FI\_B 00293

**Title:** "Sirtuin-dependent protection of genome stability under stress and its implications in cancer and aging"

**Period:** 01/04/2020 - 31/03/2023

### 2020 Fundación Científica de la Asociación Española Contra el Cáncer, Proyectos Estratégicos AECC 2020

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** PROYE20042VAQU

**Title:** Deciphering the role of SIRT7 in B-cell development and Leukemia formation

**Period:** 01/12/2020 - 30/11/2023

### 2018 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** PRE2018-084435

**Title:** Protección de la estabilidad del genoma por sirtuinas en condiciones de estrés y sus implicaciones en cáncer y envejecimiento

**Period:** 01/05/2020 - 31/08/2023

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** PID2020-117284RB-I00

**Title:** Role of Sirtuins in Epigenetic Regulation and Genome Integrity in Stress Response and their implication in Cancer and Aging (Sirepinome)

**Period:** 01/09/2021 - 31/08/2024

### 2020 European Commission, European Proteomics Infrastructure Consortium providing Access

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Title:** Study of Sirtuins' enzymatic duality in cancer



## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

**Period:** 01/03/2021 - 28/02/2022

### 2021 Deutsche Forschungsgemeinschaft, Walter Benjamin Abroad Fellowship 2021

**PI:** VAQUERO GARCÍA, ALEJANDRO

**Reference:** 493080688

**Title:** Role of the SIRT7/NPM pathway in lung cancer progression

**Period:** 01/11/2021 - 31/10/2023

## 4. Chromatin Metabolism and Cell Fate

### 2016 Instituto de Salud Carlos III, Proyectos integrados de excelencia

**PI:** BUSCHBECK, MARCUS

**Reference:** PIE16/00011

**Title:** Biomarkers and combinatorial drug targets for a personalized therapy for three major cancers.

**Period:** 01/01/2017 - 30/06/2021

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** BUSCHBECK, MARCUS

**Reference:** 2017 SGR 00305

**Title:** Modalitat GRPRE (Pre-Consolidats). Chromatin, Metabolism and Cell Fate.

**Period:** 01/01/2017 - 30/09/2021

### 2018 MINISTERIO DE CIENCIA INNOVACIÓN Y UNIVERSIDADES, Retos Investigación

**PI:** BUSCHBECK, MARCUS

**Reference:** RTI2018-094005-B-I00

**Title:** Regulation of 3D chromatin architecture by histone variants and their metabolite binding capacity.

**Period:** 01/01/2019 - 30/09/2022

### 2019 Fundació La Marató de TV3, Marató 2019: Càncer

**PI:** BUSCHBECK, MARCUS

**Reference:** 201907

**Title:** Exploring and exploiting histone variants as drug targets in acute myeloid leukemia

**Period:** 20/01/2021 - 19/01/2024

### 2018 European Commission, COST Actions 2018

**PI:** BUSCHBECK, MARCUS

**Reference:** CA18127

**Title:** International Nucleome Consortium

**Period:** 13/05/2019 - 12/05/2023

### 2016 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)

**PI:** BUSCHBECK, MARCUS

**Reference:** BES-2016-077251

**Title:** MacroH2A histone variants link genome architecture to metabolism

**Period:** 18/04/2017 - 17/04/2021

### 2018 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 18) 2018

**PI:** BUSCHBECK, MARCUS

**Reference:** DJCLS 14R/18

**Title:** Understanding and targeting epigenetic alterations in the hematopoietic stem cell niche for MDS therapy

**Period:** 01/05/2019 - 30/04/2022

### 2020 European Commission, H2020\_MSCA\_ITN-2020 Marie Skłodowska-Curie Actions – Innovative Training Networks

**PI:** BUSCHBECK, MARCUS

**Reference:** 953407

**Title:** Exploring cell-to-cell heterogeneity and exploiting epigenetic regulation for the interception of myeloid disease cells.

**Period:** 01/01/2021 - 31/12/2024

### 2019 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)

**PI:** BUSCHBECK, MARCUS

**Reference:** PRE2019-088529

**Title:** Regulación de la arquitectura tridimensional de la cromatina por parte de las variantes de histona macroh2a y su capacidad de unir metabolitos

**Period:** 01/10/2020 - 30/09/2024

### 2019 European Molecular Biology Organization, EMBO Workshop 2019

**PI:** BUSCHBECK, MARCUS

**Reference:** w20/08

**Title:** Physiology and functions of histone variants

**Period:** 22/09/2021 - 24/09/2021

### 2020 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 20) 2020

**PI:** BUSCHBECK, MARCUS

## 5. 3D Chromatin Organization

### 2018 Fundació "La Caixa", "La Caixa" postdoctoral Fellowships Junior Leader - Incoming

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** LCF/BQ/PI19/11690001

**Title:** Dynamic 3D Chromatin Organization in Human Hematopoiesis: revealing the contributions of non-coding genetic variants and mutations in blood malignancies with the aim of identifying novel disease-associated genes.

**Period:** 30/09/2019 - 29/09/2022

### 2018 MINISTERIO DE CIENCIA INNOVACIÓN Y UNIVERSIDADES, Retos Investigación

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** RTI2018-094788-A-I00

**Title:** Dynamic 3D Chromatin Organization in Human Hematopoiesis: description of novel genes associated with hematopoiesis disorders.

**Period:** 30/09/2019 - 29/09/2022

### 2019 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la contractació de personal investigador novell per a l'any 2019. FI-DGR 2019

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** 2019 FI\_B 00017

**Title:** Desxifrant noves dianes moleculars per a teràpies contra la leucèmia Limfoblàstica aguda infantil

**Period:** 01/04/2019 - 31/03/2022

### 2019 European Hematology Association, EHA Advanced Research Grant

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** 4823998

**Title:** Dissecting the Role of Non-coding Genome in B-precursor Acute Lymphoblastic Leukemia under the THREE-DIMENSIONAL Genome Architecture Point of View

**Period:** 01/09/2021 - 31/08/2023

### 2019 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 19) 2019

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** DJCLS 08R/2019

**Title:** Deciphering the Oncogenic Role of the PRC1 Complexes through Integration of Functional and Spatial Genomics in Diffuse Large B-cell Lymphoma

**Period:** 01/09/2020 - 28/02/2023

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

## 2019 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos doctorales para la formación de doctores (FPI)

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** PRE2019-088005

**Title:** Organización dinámica 3D de la cromatina en la hematopoyesis humana: descripción de nuevos genes asociados a enfermedades hematológicas

**Period:** 01/08/2020 - 31/07/2024

## 2020 Wellcome LEAP Inc, Human Organs, Physiology, and Engineering (HOPE)

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** HOPE-2021-2754490174

**Title:** Engineering human B cells: accelerating modelling of disease, drug screenings and translation of cancer immunotherapy

**Period:** 01/04/2021 - 31/03/2024

## 2021 Fundación Científica de la Asociación Española Contra el Cáncer, LAB AECC 2021

**PI:** JAVIERRE MARTINEZ, BIOLA M

**Reference:** LABAE21981JAVI

**Title:** Multi-omics approach to improve therapeutic management of T-cell acute lymphoblastic leukemia

**Period:** 01/04/2021 - 31/03/2024

## 6. Epigenetics and Immune Disease

### 2017 Ministerio de Ciencia, Innovación y Universidades , Programa Estatal de I+D+i Orientada a los Retos de la Sociedad

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** SAF2017-88086-R

**Title:** Myeloid cells and Epigenetic Plasticity: Mechanisms and Implications in Autoimmune and other Inflammatory Processes

**Period:** 01/06/2019 - 30/09/2021

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** 2017 SGR 00720

**Title:** Grup de Cromatina i Malaltia Grup de Recerca Consolidat

**Period:** 01/01/2017 - 30/09/2021

### 2019 Ministerio de Ciencia, Innovación y Universidades , Ayudas Juan de la Cierva - Formación 2019

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** FJC2019-040868-I

**Title:** Unravelling the molecular mechanisms of CAR-T cell activation in vivo on B-ALL patients

**Period:** 01/01/2021 - 18/04/2022

### 2018 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos doctorales para la formación de doctores (FPI)

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** PRE2018-083544

**Title:** Células mieloides y plasticidad epigenética: Mecanismos e implicaciones en procesos autoinmunes e inflamatorios

**Period:** 01/05/2020 - 30/06/2023

### 2018 Instituto de Salud Carlos III, Acciones complementarias de programación conjunta internacional

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** AC18/00057

**Title:** Integrative Multi-Omics Analysis of Primary Antibody Deficiency (PAD) Patients for Stratification According to Cellular Pathways (E-RARE)

**Period:** 01/04/2019 - 31/12/2021

### 2017 Instituto de Salud Carlos III, Contratos doctorales de formación en investigación en salud. Modalidad Contratos i-PFIS: doctorados IIS-empresa en ciencias y tecnologías de la salud

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** IFI17/00034

**Title:** Papel de los glucocorticoides, y otros fármacos relacionados, en la remodelación epigenética asociada a la adquisición telerogénesis en células mieloides: Implicaciones en el tratamiento de enfermedades autoinmunes.

**Period:** 15/01/2018 - 14/01/2022

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** BALLESTAR TARIN, ESTEBAN

**Reference:** PID2020-117212RB-I00

**Title:** Understanding the Role of Immune cell-cell Crosstalk in Epigenetic Dysregulation in Inflammation (InflaEpiTalk)

**Period:** 01/09/2021 - 31/08/2024

## 7. Lymphocyte Development and Disease

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** PARRA BOLA, MARIA ISABEL

**Reference:** 2017 SGR 00149

**Title:** Grup de Diferenciació Cel·lular, Grup de Recerca Pre-Consolidat

**Period:** 01/01/2017 - 30/09/2021

### 2019 Ministerio de Ciencia e Innovación, Acciones de dinamización «Europa Investigación»

**PI:** PARRA BOLA, MARIA ISABEL

**Reference:** EUR2019-103835

**Title:** B cell differentiation; unraveling gene silencing mechanisms

### 2017 Ministerio de Ciencia, Innovación y Universidades , Programa Estatal de I+D+i Orientada a los Retos de la Sociedad

**PI:** PARRA BOLA, MARIA ISABEL

**Reference:** SAF2017-87990-R

**Title:** Transcriptional repressive mechanisms in early and terminal B cell differentiation

**Period:** 01/06/2019 - 30/09/2021

### 2020 Fundación Científica de la Asociación Española Contra el Cáncer, Postdoctorales AECC 2020

**PI:** PARRA BOLA, MARIA ISABEL

**Reference:** POSTD20024DEBA

**Title:** Targeted Modulation of the Transcriptional Repressor Hdac7; towards Precision Medicine in Mll-Af4-Rearranged Infant Pro-B-All

**Period:** 01/12/2020 - 30/11/2022

### 2020 Federation of European Biochemical Societies, Beques de curta durada de la Federation of European Biochemical Societies (FEBS)

**Researcher:** AZAGRA RODRIGUEZ, ALBA

**Title:** Towards personalized medicine in Diffuse Large B Cell Lymphoma (DLBCL) using 3D organoids

**Period:** 05/04/2021 - 04/06/2021

### 2018 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos doctorales para la formación de doctores (FPI)

**PI:** PARRA BOLA, MARIA ISABEL

## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

**Reference:** PRE2018-083183

**Title:** Mecanismos de represión transcripcional en la diferenciación temprana y terminal de linfocitos B

**Period:** 01/05/2020 - 30/06/2023

### 2021 European Commission, Intellectual Property Booster 2021 - 2

**PI:** PARRA BOLA, MARIA ISABEL

**Title:** Mechanisms of transcriptional repression in early and terminal B cell differentiation (HDAC7-BLYM)

**Period:** 07/06/2021 - 22/09/2021

## 8. Regulatory Genomics

### 2019 Ministerio de Ciencia, Innovación y Universidades , Generación del Conocimiento

**PI:** VAVOURI, TANYA SOULTANA

**Reference:** PID2019-111676GB-I00

**Title:** Evolution of new PIWI-interacting RNAs in a mammal (ASliceOfPi)

**Period:** 01/06/2020 - 31/05/2023

## 9. Regulatory RNA and Chromatin

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** GUIL DOMÈNECH, SÒNIA

**Reference:** 2017 SGR 00722

**Title:** RNA regulador i cromatina, Grup de Recerca Consolidat

**Period:** 01/01/2017 - 30/09/2021

### 2019 Ministerio de Ciencia, Innovación y Universidades , Retos Investigación

**PI:** GUIL DOMÈNECH, SÒNIA

**Reference:** PID2019-111658RB-I00

**Title:** Pseudogenes as long noncoding oncofetal RNAs: functional characterization and implications for anticancer therapy.

**Period:** 01/06/2020 - 31/05/2023

### 2020 Sarcoma Foundation of America, 2020 Funding Opportunity Announcement

**PI:** GUIL DOMÈNECH, SÒNIA

**Reference:** SFA20-06

**Title:** The Role of Pseudogenes as Regulators of the Igf2bp2/Igf1r/Ras Axis in Rhabdomyosarcoma: Functional Characterization in Myogenesis and

Implications for Anticancer Therapy

**Period:** 01/07/2020 - 31/12/2021

### 2017 Instituto de Salud Carlos III, Contratos predoctorales de formación en investigación en salud. Modalidad Contratos i-PFIS: doctorados IIS-empresa en ciencias y tecnologías de la salud

**PI:** GUIL DOMÈNECH, SÒNIA

**Reference:** IFI17/00006

**Title:** Development of new personalized treatments against non Hodgkin lymphoma based on the inhibition of the epigenetic target HDAC6 and the obtention of new biomarkers for this target. Regeneración hematopoyética a partir de células madre pluripotentes.

**Period:** 01/01/2018 - 31/12/2021

## 10. Epigenetic Control of Hematopoiesis

### 2019 Worldwide Cancer Research, April 2019 grant round

**PI:** SARDINA ORTEGA, JOSÉ LUIS

**Reference:** 20-0269

**Title:** Uncovering the regulation of chromatin

structure by TET2 during leukemic cell fate decisions

**Period:** 01/03/2020 - 28/02/2023

### 2019 Instituto de Salud Carlos III, Contratos Miguel Servet - Tipo 1

**PI:** SARDINA ORTEGA, JOSÉ LUIS

**Reference:** MS19/00176

**Title:** Role of TET2 in chromatin structure regulation during the onset and development of myeloid malignancies

**Period:** 01/01/2020 - 31/12/2024

### 2019 Ministerio de Ciencia, Innovación y Universidades , Retos Investigación

**PI:** SARDINA ORTEGA, JOSÉ LUIS

**Reference:** PID2019-111243RA-I00

**Title:** Unveiling TET2 impact on chromatin structure at the leukemic onset

**Period:** 01/06/2020 - 31/05/2023

### 2020 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales en el marco del Plan Estatal de I+D+i (FPI)

**PI:** SARDINA ORTEGA, JOSÉ LUIS

**Reference:** PRE2020-093881

**Title:** Descifrando el impacto de tet2 sobre la estructura de la cromatina en el inicio leucémico

**Period:** 01/08/2021 - 31/07/2025

## 11. Transcriptional Dynamics in Leukemia

### 2019 Jerome Lejeune Foundation, Cycle 2019b-Down syndrome research

**PI:** CUARTERO BETRIU, SERGI

**Reference:** #1902

**Title:** Myeloid leukemia in Down syndrome: exploring the interplay between transcriptional regulation and immune signalling

**Period:** 01/04/2020 - 06/01/2023

### 2020 Leukemia Research Foundation, Hollis Brownstein Research Grants Program

**PI:** CUARTERO BETRIU, SERGI

**Reference:** 02/2020

**Title:** Identifying novel transcriptional vulnerabilities in MDS with cohesin mutations

**Period:** 01/09/2020 - 31/12/2021

### 2019 Fundació ""La Caixa"", Junior Leader 2020

**PI:** CUARTERO BETRIU, SERGI

**Reference:** LCF/BQ/PI20/11760002

**Title:** The role of inflammation in myeloid malignancies

**Period:** 01/09/2020 - 31/08/2023

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** CUARTERO BETRIU, SERGI

**Reference:** PID2020-117950RA-I00

**Title:** Deciphering the role of cohesin mutations and the 3D genome structure in myeloid leukemia (MYELO-3D)

**Period:** 01/09/2021 - 31/08/2024

## 12. Cancer Immunogenomics

### 2020 Fundación Científica de la Asociación Española Contra el Cáncer, LAB AECC 2020

**PI:** PORTA PARDO, EDUARD

**Reference:** LABAE20038PORT

**Title:** A cellular and molecular map of bladder cancer to guide neoadjuvant treatment

**Period:** 01/12/2020 - 30/11/2023



## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

### 2019 Ministerio de Ciencia, Innovación y Universidades , Subprograma de ayudas para contratos Ramon y Cajal

**PI:** PORTA PARDO, EDUARD

**Reference:** RYC2019-026415-I

**Title:** RYC Eduard Porta

**Period:** 01/09/2021 - 31/08/2026

## 13. Cancer Heterogeneity and Hierarchies

### 2018 Ministerio de Economía y Competitividad, Ramón y Cajal

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** RYC2018-024099-I

**Title:** In vivo models to study cellular hierarchies and cancer

**Period:** 01/05/2020 - 30/04/2025

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** PID2020-114647RA-I00

**Title:** Lineage tracing studies to unravel mammary epithelial hierarchies and cellular plasticity in breast

cancer

**Period:** 01/09/2021 - 31/08/2024

### 2020 Asociación Española de investigación sobre el Cáncer , ASEICA

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** AJT-PRJ2100397

**Title:** Monitoring senescence upon chemotherapy (MonSen)

**Period:** 01/12/2020 - 31/05/2022

### 2021 Departament de Salut, Generalitat de Catalunya, Subvencions per a la contractació de personal investigador en formació (PIF-SALUT)

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** SLT017/20/000140

**Title:** Estudis clonals per entendre les jerarquies i plasticitat c

el-lular en el càncer de mama

**Period:** 15/07/2021 - 31/12/2024

### 2021 Fundación de Investigación Oncológica, III PROYECTO FERO-GHD en Cáncer de Mama 2021

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** PFERO2021.01

**Title:** Tumor heterogeneity comprehension for an

improved diagnosis and treatment choice for breast cancer

**Period:** 01/06/2021 - 31/05/2023

### 2021 Fundación Científica de la Asociación Española Contra el Cáncer, LAB AECC 2021

**PI:** RODILLA BENITO, VERÓNICA

**Reference:** LABAE211626RODI

**Title:** Tumor heterogeneity comprehension for an improved diagnosis and treatment choice for breast cancer.

**Period:** 01/12/2021 - 30/11/2024

## 14. Leukemia and Immuno-Oncology

### 2020 Fundación Fero, BECA FERO EN INVESTIGACIÓN ONCOLÓGICA TRASLACIONAL 2020

**PI:** BELVER MIGUEL, LAURA

**Reference:** BFERO2020.03

**Title:** Molecular pathways and targeted therapies in Juvenile Myelomonocytic Leukemia

**Period:** 01/12/2020 - 30/11/2022

### 2020 Govern d'Andorra, Ajuts de tercer cicle de l'any 2020 - Modalitat 1 - Nous ajuts

**PI:** BELVER MIGUEL, LAURA

**Reference:** AJT-PRS2000176

**Title:** Oncogenic mechanisms and target therapies in JMML

**Period:** 02/01/2021 - 01/01/2024

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** BELVER MIGUEL, LAURA

**Reference:** PID2020-117645RA-I00

**Title:** Functional impact of enhancer-associated noncoding mutations in Juvenile Myelomonocytic Leukemia (JMML\_ENH)

**Period:** 01/09/2021 - 31/08/2024

## 15. Cellular Systems Genomics

### 2019 Ministerio de Ciencia, Innovación y Universidades , Ayudas Juan de la Cierva - Incorporación 2019

**PI:** MEREU, ELISABETTA

**Reference:** IJC2019-041346-I

**Title:** Juan de la Cierva Inc\_Elisabetta Mereu

**Period:** 01/05/2021 - 30/04/2024

## 16. Endothelial Pathobiology and Microenvironment

### 2020 European Commission, H2020\_MSCA\_ITN-2020 Marie Skłodowska-Curie Actions – Innovative Training Networks

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** 955951

**Title:** Deconstructing the evolution of metastasis

**Period:** 01/03/2021 - 28/02/2025

### 2020 Worldwide Cancer Research, March 2020 grant round

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** 21-0159

**Title:** Identifying properties of tumor suppressive pericytes for cancer therapy

**Period:** 01/03/2021 - 31/08/2024

### 2021 Fundació ""La Caixa"", Immunoepigenetics Research Program

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Title:** Estudi de l'activació de la via de la PI3K en malformacions vasculares oncogèniques

**Period:** 01/03/2021 - 30/06/2022

### 2019 Fundació ""La Caixa"", Junior Leader 2020

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** LCF/BQ/PR20/1177002

**Title:** Pathogenic and pharmacologic study of cutaneous capillary malformations and Sturge-Weber syndrome

**Period:** 01/03/2021 - 29/12/2023

### 2019 European Molecular Biology Organization, EMBO Workshop 2019

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** w20/44

**Title:** Vascular malformations: From fundamental biology to therapeutic opportunities

**Period:** 06/10/2021 - 08/10/2021

### 2021 Cloves Syndrome Community, 2021 Research Grant Program

**Researcher:** ANGULO URARTE, ANA

**Title:** Identifying the molecular impact of PIK3CA variants in PROS towards stratification of patients and personalized medicine.

**Period:** 01/09/2021 - 31/08/2022

## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

### 2020 European Commission, H2020 MSCA-ITN-2020 Marie Skłodowska-Curie Actions – Innovative Training Networks

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** 955534

**Title:** IPI3K/PTEN-related monogenic disease to understand cancer

**Period:** 01/07/2021 - 30/06/2025

### 2019 PTEN Research Foundation, 2020 call

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** IJC-21-001

**Title:** Preclinical investigation of vascular malformations in PTEN hamartoma tumor syndrome

**Period:** 01/05/2021 - 01/08/2024

### 2019 Fundación BBVA, Ayudas Fundación BBVA Equipos de Investigación Científica 2019

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** PR(19)BIO\_MET\_0061

**Title:** Endothelial molecular alterations induced by excessive energy intake - a new concept in obesity and metabolic disorders

**Period:** 01/02/2021 - 30/04/2023

### 2018 Fundación Científica de la Asociación Española Contra el Cáncer, GRUPOS TRASLACIONALES AECC

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** GCTRA18006CARR

**Title:** Vulnerabilities of Tumor and Stroma Interactions in Castration-Naïve Metastatic Prostate Cancer

**Period:** 01/04/2021 - 30/09/2023

### 2019 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la contractació de personal investigador novell per a l'any 2019. FI-DGR 2019

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** 2021 FI\_B1 00219

**Title:** Estudi patogènic i farmacològic de les malformacions capil·lars cutànies i del síndrome Struge-Weber

**Period:** 01/05/2021 - 30/04/2023

### 2020 Fundació ""La Caixa"", Health Research 2021

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** HR21-00046

**Title:** Decoding the paracrine control of metabolic fitness by endothelial nutrient signaling

**Period:** 01/12/2021 - 30/11/2024

### 2021 Fundación Científica de la Asociación Española Contra el Cáncer, Investigador AECC

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** INVES211084VILL

**Title:** Mapping stromal molecular programs that govern tumor aggressiveness in Prostate Cancer

**Period:** 01/07/2021 - 30/06/2023

### 2018 Fundació ""La Caixa"", Health Research 2018

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** HR18-00120

**Title:** Mapping the pathogenesis of vascular malformations

**Period:** 01/03/2021 - 01/11/2022

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** PID2020-116184RB-I00

**Title:** PIK3CA variants in PROS: cracking the Code of pathogenesis

**Period:** 01/09/2021 - 31/08/2024

### 2018 Ministerio de Ciencia, Innovación y Universidades , Ayudas para contratos predoctorales para la formación de doctores (FPI)

**PI:** GRAUPERA GARCIA - MILA, MARIONA

**Reference:** PRE2018-084283

**Period:** 01/11/2021 - 30/06/2023

## 17. T-Cell Lymphoma

### 2020 Ministerio de Ciencia e Innovación, Retos Investigación 2021

**PI:** MONDRAGÓN MARTÍNEZ, LAURA

**Reference:** PID2020-116049RA-I00

**Title:** Role of Apaf-1 IN tcr- thymoCytes maturatiOn and lymphomA developmenT - RAINCOAT

**Period:** 01/09/2021 - 31/08/2024

### 2019 Ministerio de Ciencia, Innovación y Universidades , Subprograma de ayudas para contratos Ramon y Cajal

**PI:** MONDRAGÓN MARTÍNEZ, LAURA

**Reference:** RYC2019-026522-I

**Period:** 01/04/2021 - 31/03/2026

## 18. Acute Lymphoblastic Leukemia (ALL)

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts per a la contractació de personal investigador novell per a l'any 2018. FI-DGR 2018

**Researcher:** GENESCÀ FERRER, EULÀLIA

**Reference:** 2018 FI\_B 00970, 2019 FI\_B100224, 2020 FI\_B2 00210

**Title:** Estudio de la resistencia al tratamiento en la leucemia linfoblástica aguda de subtipo T (LAL-T) del adulto. Búsqueda de nuevas alternativas terapéuticas

**Period:** 01/06/2018 - 30/10/2021

### 2019 Instituto de Salud Carlos III, Proyectos de investigación en Salud

Researcher: GENESCÀ FERRER, EULÀLIA

**Reference:** PI19/01828

**Title:** Use of Next Generation Sequencing (NGS) as a unique genomic technique to apply to improve diagnosis, prognosis and treatment of adult T-cell Acute Lymphoblastic Leukemia patients

**Period:** 01/01/2020 - 31/12/2022

### 2016 Fundación Científica de la Asociación Española Contra el Cáncer, Grupos Coordinados Estables de Investigación

**PI:** RIBERA SANTASUSANA, JOSEP MARIA

**Reference:** GC16173697BIGA

**Title:** Exploring Mechanisms of Resistance in Adult and Pediatric T-Acute Lymphoblastic Leukemia

**Period:** 01/11/2016 - 31/10/2022

### 2015 European Commission, H2020 JTI-IMI2 2015-06

**PI:** RIBERA SANTASUSANA, JOSEP MARIA

**Reference:** 116026

**Title:** Healthcare Alliance for Resourceful Medicines Offensive against Neoplasms in hematology

**Period:** 01/01/2017 - 30/06/2023

### 2019 European Commission, IMI2-2019-19-01

**PI:** RIBERA SANTASUSANA, JOSEP MARIA

**Reference:** 945406

**Title:** Healthcare Alliance for Resourceful Medicines Offensive against Neoplasms in Hematology - PLUS

**Period:** 01/10/2020 - 30/09/2023

## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

## EXPERIMENTAL AND CLINICAL HEMATOLOGY PROGRAM (PHEC)

### 19. Barcelona Endothelium Team (BET)

#### 2019 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 19) 2019

**PI:** CARRERAS PONS, ENRIC

**Reference:** DJCLS 03R/2019

**Title:** Enhancement of endothelial regeneration and endothelial function during GVHD

**Period:** 01/01/2020 - 30/04/2022

#### 2020 Fundació La Marató de TV3, Malalties Minoritàries

**Researcher:** PALOMO DE UDAETA, MARTA

**Reference:** 202026-10

**Title:** Analysis of the complement system as a therapeutic target in severe pre-eclampsia andHELLP syndrome

**Period:** 19/03/2021 - 18/03/2024

### 20. Myeloid Neoplasms

#### 2016 Instituto de Salud Carlos III, Proyectos de investigación en Salud

**PI:** ZAMORA PLANA, LURDES

**Reference:** PI16/01200

**Title:** Biological predictive factors for achieving deep molecular response and relapse after tyrosine kinase inhibitor discontinuation in patients with Chronic Myeloid Leukemia

**Period:** 01/01/2017 - 30/09/2021

### 21. Leukemia Stem Cell Group

#### 2019 Fundació La Marató de TV3, Marató 2019: Càncer

**PI:** MUÑOZ RISUEÑO, RUTH

**Reference:** 201930-30

**Title:** Lysosomal impairment-based therapeutic approach for leukemia

**Period:** 30/07/2020 - 29/07/2023

#### 2019 Agència de Gestió d'Ajuts Universitaris i de Recerca, Ajuts destinats a l'obtenció de prototipus i a la valorització i transferència dels resultats d'investigació generada per equips de recerca de Catalunya (PRODUCTE)

**PI:** MUÑOZ RISUEÑO, RUTH

**Reference:** 2019PROD00031

**Title:** Teràpies basades en CADs com a nou mecanisme autofàgic per a leucèmia

**Period:** 23/07/2020 - 22/04/2022

#### 2016 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 17) 2017

**PI:** MUÑOZ RISUEÑO, RUTH

**Reference:** DJCLS08/R2017

**Title:** CARs for Treatment of Pediatric T-Cell Acute Lymphoblastic Leukemia

**Period:** 08/08/2017 - 31/10/2021

#### 2019 Ministerio de Ciencia, Innovación y Universidades, Retos Investigación

**PI:** MUÑOZ RISUEÑO, RUTH

**Reference:** PID2019-111348RB-I00

**Title:** Chemosensitizers as a new therapeutic approach for acute myeloid leukemia (ChemoLeuko)

**Period:** 01/06/2020 - 31/05/2023

### 22. Lymphoid Neoplasms

#### 2019 Instituto de Salud Carlos III, Proyectos de investigación en Salud

**PI:** NAVARRO FERRANDO, JOSE TOMAS

**Reference:** PI19/01588

**Title:** Comprehensive epigenomic analysis of plasmablastic lymphoma: Identifying epigenetic features to improve diagnosis and patient outcomes

**Period:** 01/01/2020 - 31/12/2022

#### 2019 Gilead Sciences, SL, 7ª Convocatoria de Becas Gilead a la investigación Biomédica en VIH, Enfermedades hepáticas y Hemato-oncología

**PI:** NAVARRO FERRANDO, JOSE TOMAS

**Reference:** GLD19\_00121

**Title:** Disclosing the transcriptome of HIV-DLBCL for targeted therapy

**Period:** 01/01/2020 - 31/12/2021

### 23. Myelodysplastic Syndromes

#### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** SOLE RISTOL, FRANCESC

**Reference:** 2017 SGR 00288

**Title:** Modalitat GRC (Reconegut i Consolidat).

**Period:** 01/01/2017 - 30/09/2021

#### 2018 Instituto de Salud Carlos III, Acciones

#### complementarias de programación conjunta internacional

**PI:** SOLE RISTOL, FRANCESC

**Reference:** AC18/00002

**Title:** An integrated European platform to conduct translational studies in myelodysplastic syndromes based on the EuroBloodNet infrastructure

**Period:** 01/01/2019 - 31/12/2021

#### 2017 Instituto de Salud Carlos III, Proyectos de investigación en Salud

**PI:** SOLE RISTOL, FRANCESC

**Reference:** PI17/00575

**Title:** Aplicación de la secuenciación masiva (NGS) en el diagnóstico y pronóstico de síndromes mielodisplásicos/neoplasias mieloproliferativas.

**Period:** 01/01/2018 - 30/06/2022

#### 2018 Fundación Científica de la Asociación Española Contra el Cáncer, Ayudas a proyectos de investigación en cáncer TRANSCAN (Translational Research on Rare Cancers)

**PI:** SOLE RISTOL, FRANCESC

**Reference:** TRNSC18003SOL

**Title:** An integrated European platform to conduct translational studies in myelodysplastic syndromes based on the EuroBloodNet infrastructure

**Period:** 01/12/2018 - 31/03/2022



## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

### 2020 Instituto de Salud Carlos III, Proyectos de investigación en Salud (FIS)

**PI:** SOLE RISTOL, FRANCESC

**Reference:** PI20/00531

**Title:** Caracterización genética de las neoplasias mieloides asociadas a tratamiento (Therapy related myeloid neoplasms, TRMN)

**Period:** 01/01/2021 - 31/12/2023

### 2020 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 20) 2020

**PI:** SOLE RISTOL, FRANCESC

**Reference:** DJCLS 01R/2021

**Title:** Dissecting the mechanisms of clonal expansion in del(5q) myelodysplastic syndrome to selectively target the disease-initiating hematopoietic stem cells

**Period:** 01/10/2021 - 30/09/2024

## 22. Stem Cell Biology, developmental leukemia and immunotherapy

### 2017 Agència de Gestió d'Ajuts Universitaris i de Recerca, Grups de Recerca de Catalunya (SGR)

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 2017 SGR 00221

**Title:** Modalitat GRC (Consolidat). Concedit a IJC

**Period:** 01/01/2017 - 30/09/2021

### 2017 Ministerio de Economía y Competitividad, Retos-Colaboracion

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** RTC-2017-6367-1

**Title:** Obtención de hematíes in vitro a partir de iPSCs de donantes con fenotipos eritrocitarios seleccionados y optimizados mediante edición genómica, como alternativa a los paneles de hematíes actuales

**Period:** 01/09/2018 - 31/03/2022

### 2018 European Commission, H2020-SC1-BHC-2018-2020 (Topics 2018)

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 825749

**Title:** Childhood Leukemia: Overcoming distance between South America and Europe Regions

**Period:** 01/01/2019 - 31/12/2023

### 2017 Ministerio de Economía y Competitividad, Ayudas Juan de la Cierva - Incorporación 2017

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** IJCI-2017-33172

**Title:** Recreación de traslocaciones cromosómicas asociadas a leucemia pediátrica en progenitores humanos

**Period:** 01/02/2019 - 16/01/2022

### 2019 Fundación Científica de la Asociación Española Contra el Cáncer, Ideas Semilla AECC

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** IDEAS19005MENE

**Title:** Redirecting CAR T-Cells to the Bone Marrow: Improved CAR T-Cell Persistence and Anti-Leukemia Effects while Alleviating related Toxicity

**Period:** 01/10/2019 - 30/09/2021

### 2018 Fundació "La Caixa", Health Research 2018

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** HR18-00069

**Title:** Next-generation CAR-DOT cells for allogeneic adoptive cancer immunotherapy

**Period:** 15/09/2019 - 31/12/2022

### 2019 Instituto de Salud Carlos III, Contratos Sara Borrell

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** CD19/00013

**Title:** Towards a clinical Translation of the CD1A-directed car for relapse/refractory cortical T-cell acute Leukemia and Langerhans cell histiocytosis:

Feasibility, efficacy and Safety

**Period:** 01/01/2020 - 31/12/2022

### 2014 European Commission, ERC-2014-CoG

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 646903

**Title:** Genomic, cellular and developmental reconstruction of infant MLL-AF4+ Acute Lymphoblastic Leukemia

**Period:** 01/01/2016 - 30/06/2021

### 2017 European Cooperation in Science and Technology (COST), COST Actions 2017

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** CA16223

**Title:** Leukemia GENe Discovery by data sharing, mining and collaboration

**Period:** 26/10/2017 - 25/10/2021

### 2020 European Commission, Proof of Concept Grants 2020 (ERC-2020-PoC)

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 957466

**Title:** Clinical translation of a novel CD1a-directed CAR for relapse/refractory cortical T-cell Acute Lymphoblastic Leukemia: feasibility, efficacy and safety

**Period:** 01/01/2021 - 30/06/2022

### 2019 Deutsche José Carreras Leukämie Stiftung, Forschungsprojekte (R 19) 2019

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 19R2019

**Title:** Healthy Versus Non-Healthy Aging: Identifying Metabolic and Epigenetic Vulnerabilities of Single Leukemic Stem Cells in Acute Myeloid Leukemia Patients

**Period:** 01/01/2021 - 31/12/2022

### 2020 Instituto de Salud Carlos III, Proyectos de investigación en Salud (FIS)

**Researcher:** BUENO UROZ, CLARA

**Reference:** PI20/00822

**Title:** TIM3, una nueva y prometedora diana inmunoterapéutica en Ila-b de novo y en recaída

**Period:** 01/01/2021 - 31/12/2023

### 2019 Fundación Uno Entre Cien Mil , VI Beca Unoentrecienmil. Fundación para la investigación en el área de la leucemia aguda infantil del 2019

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** 1entre100mil

**Title:** Towards a clinical translation of the CD1a-directed CAR for relapse/refractory cortical T-cell Acute Lymphoblastic Leukemia and Langerhans Cell Histiocytosis: feasibility, efficacy and safety

**Period:** 01/06/2019 - 31/05/2021

### 2020 Fundación Científica de la Asociación Española Contra el Cáncer, Investigador AECC 2020

**Researcher:** BUENO UROZ, CLARA

**Reference:** INVES20011LÓPE

**Title:** Innovative Therapeutic Strategies for Infant Mixed Lineage Leukemiarearranged B-cell Acute Lymphoblastic Leukemia: NG2 and CD22 as therapeutic targets

**Period:** 01/09/2020 - 31/08/2022

### 2019 Ministerio de Ciencia, Innovación y Universidades , Retos Investigación

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** PID2019-108160RB-I00

**Title:** IM3, a promising novel immunotherapeutic target for de novo and relapsed B-cell acute lymphoblastic leukemia

**Period:** 01/06/2020 - 31/05/2023

### 2019 Ministerio de Universidades, Ayudas para la formación de profesorado universitario (FPU)

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** FPU19/00039

**Title:** Explorando células efectoras alogénicas para inmunoterapia en leucemia aguda

**Period:** 01/11/2020 - 31/03/2024

### 2021 Fundación Científica de la Asociación Española Contra el Cáncer, Investigador AECC

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** INVES211226MOLI

**Title:** Contribution of chromosome instability to the outcome of aneuploid childhood B-cell acute

## FACTS & FIGURES

# COMPETITIVE GRANTS AWARDED AND ACTIVE PROJECTS

lymphoblastic leukemia.

**Period:** 01/12/2021 - 30/11/2023

### 2021 Fundación Científica de la Asociación Española Contra el Cáncer, Proyectos generales AECC

**Researcher:** BUENO UROZ, CLARA

**Reference:** PRYGN211192BUEN

**Title:** Next Generation Universal T-cell Redirected ImmunoTherapy for Acute Leukemia

**Period:** 01/12/2021 - 30/11/2024

### 2021 Fundación Científica de la Asociación Española Contra el Cáncer, PRÁCTICAS VERANO AECC

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** PPLAB211825BURG

**Title:** Prácticas Laboratorio Verano AECC 2021 FRANCISCO JAVIER BURGOS RETAMAR

**Period:** 01/07/2021 - 31/08/2021

### 2021 Fundació "La Caixa", CaixaImpulse Validate 2021

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** LCF/PR/HR18/00069

**Title:** Tailored adoptive CAR T-cell Immunotherapy for Ewing Sarcoma

**Period:** 16/06/2021 - 16/06/2023

### 2021 Ministerio de Ciencia e Innovación, Proyectos de I+D+i en líneas estratégicas, en colaboración público-privada 2021

**Researcher:** BUENO UROZ, CLARA

**Reference:** PLEC2021-007518

**Title:** Recreating the Embryonic Niche for Hematopoietic Stem Cell Production and Derivatives in Human Gastruloids

**Period:** 01/12/2021 - 30/11/2024

### 2020 Ministerio de Ciencia, Innovación y Universidades, Ayudas para contratos predoctorales en el marco del Plan Estatal de I+D+i (FPI)

**PI:** MENÉNDEZ BUJÁN, PABLO

**Reference:** PRE2020-092778

**Title:** TIM3, una nueva y prometedora diana inmunoterapéutica en leucemia linfoblástica aguda bde novo y en recaída

**Period:** 01/08/2021 - 31/07/2025

## 23. Lymphoma Translational Group

### 2019 INTERREG POCTEFA, 3ª convocatoria de proyectos POCTEFA 2014-2020

**PI:** ROUÉ, GAËL

**Reference:** EFA360/19

**Title:** Red cooperativa franco-española para el análisis de proteinopatías y el desarrollo de terapias individualizadas en cánceres hematológicos

**Period:** 01/06/2019 - 31/05/2022

### 019 Gilead Sciences, SL, 7ª Convocatoria de Becas Gilead a la investigación Biomédica en VIH, Enfermedades hepáticas y Hemato-oncología

**PI:** ROUÉ, GAËL

**Reference:** GLD19\_00058

**Title:** Modulation of lymphoid microenvironment by intrinsic protein homeostasis in aggressive B-cell lymphoma

**Period:** 01/01/2020 - 31/12/2021

### 2018 Instituto de Salud Carlos III, Proyectos de Investigación en Salud (AES 2018). Modalidad proyectos en salud

**PI:** ROUÉ, GAËL

**Reference:** PI18/01383

**Title:** Development of a patient-derived xenografts platform for the evaluation of new targeted therapies in aggressive B-cell lymphomas

**Period:** 01/02/2020 - 31/12/2021

## 24. Chronic Lymphocytic Leukemia Group

**PI:** MORENO ATANASIO, CAROLINA

**Reference:** DJCLS 04 R/2021

**Title:** Definition of cellular components of the natural immune response in CLL

**Period:** 15/10/2021 - 14/10/2024

## 25. Proteomics Unit

### 2016 Fundació La Marató de TV3, Marató 2016: Ictus i lesions medul·lars i cerebrals traumàtiques

**Manager:** DE LA TORRE GÓMEZ, CAROLINA

**Reference:** 201719.32

**Title:** ITACAT: Impact of Thrombus Analysis in stroke patients in Catalonia

**Period:** 01/10/2019 - 31/01/2022

## 26. Communication Unit

### 021 Ajuntament de Barcelona, Convocatoria general de subvencions del Ajuntament de Barcelona para el año 2021

**Manager:** DÍAZ LÓPEZ, HELENA

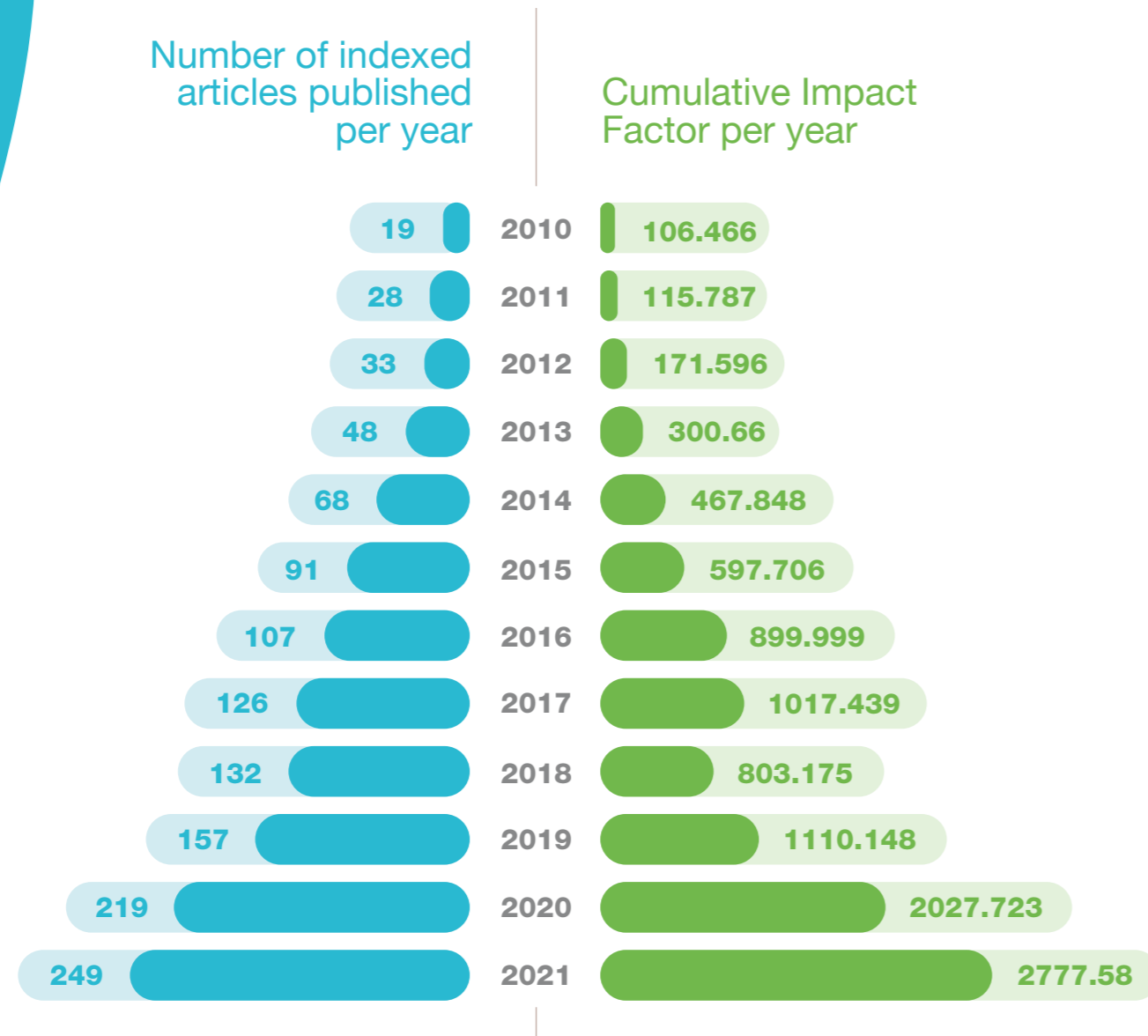
**Reference:** 21S00478-006

**Title:** InstiCIENCIA – Programa de divulgació i foment de vocacions científiques

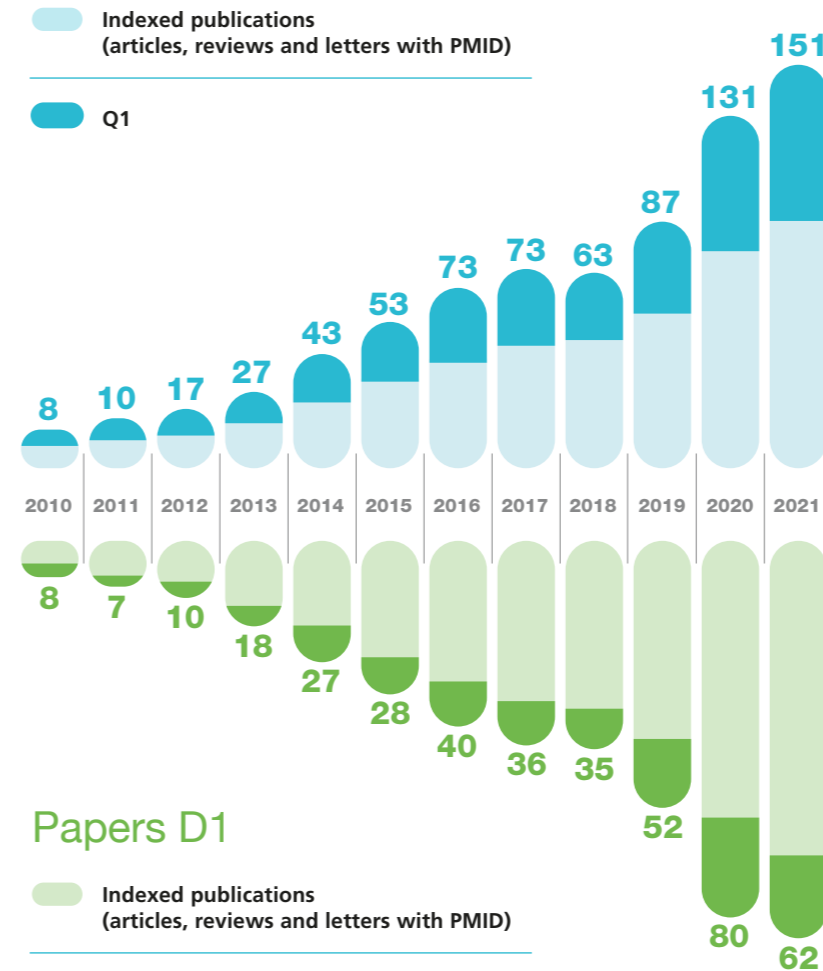
**Period:** 01/04/2021 - 31/12/2021

FACTS & FIGURES  
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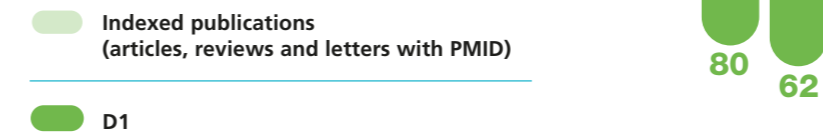
**INDICATORS**



**Papers Q1**

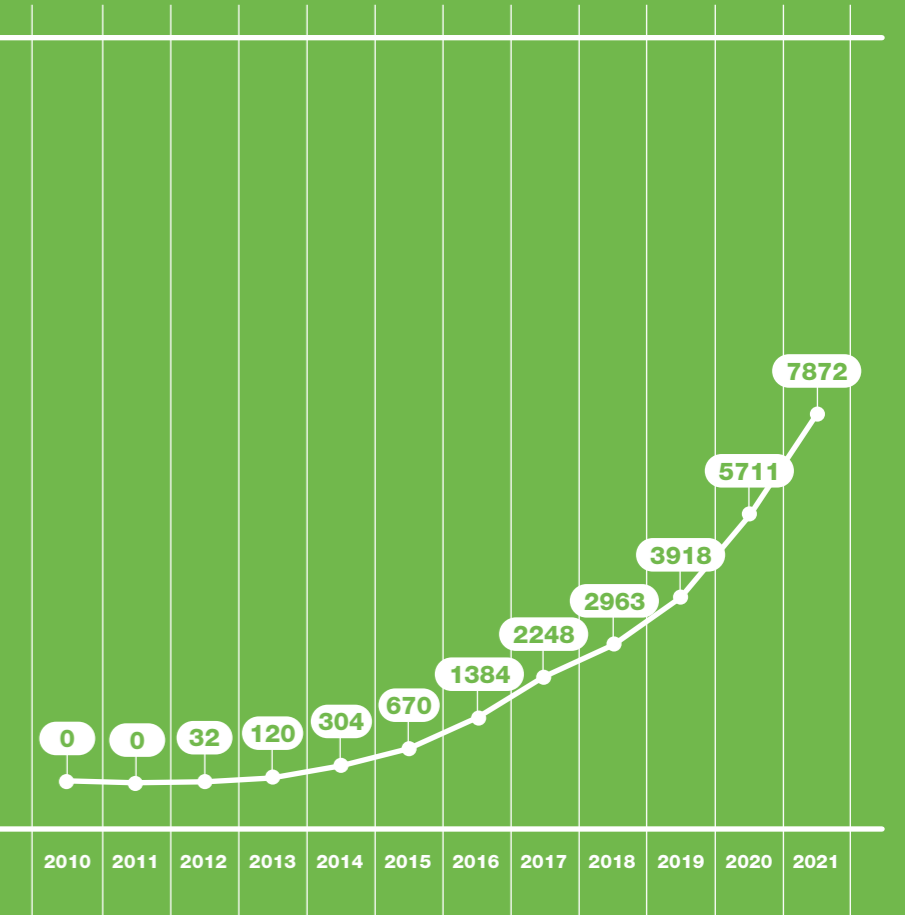


**Papers D1**



**Citations from Web of Science**

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### Cancer and Leukemia Epigenetics and Biology Program (PEBCL)

#### 1. Cancer Epigenetics

Alvarez-Palomo, AB; Requena-Osete, J; Delgado-Morales, R; Moreno-Manzano, V; Grau-Bove, C; Tejera, AM; Otero, MJ; Barrot, C; Santos-Barriopedro, I; **Vaquero, A**; Mezquita-Pla, J; Moran, S; Naya, CH; Garcia-Martinez, I; Perez, FV; Blasco, MA; **Esteller, M**; Edel, MJ

A synthetic mRNA cell reprogramming method using CYCLIN D1 promotes DNA repair generating improved genetically stable human induced pluripotent stem cells Stem Cells. 2021 Jul;39(7):866-881. doi: 10.1002/stem.3358. Epub 2021 Mar 3. Impact Factor: 5,8458 - Q1  
PMID: 33621399  
Citations: 3 [27/07/2022]

Caballero-Camino, FJ; Rivilla, I; Herraiez, E; Briz, O; Santos-Laso, A; Izquierdo-Sanchez, L; Lee-Law, PY; Rodrigues, PM; Munoz-Garrido, P; Jin, S; Peixoto, E; Richard, S; Gradilone, SA; Perugorria, MJ; **Esteller, M**; Bujanda, L; Marin, JGG; Banales, JM; Cossio, FP  
Synthetic Conjugates of Ursodeoxycholic Acid Inhibit Cystogenesis in Experimental Models of Polycystic Liver Disease Hepatology. 2021 Jan;73(1):186-203. doi: 10.1002/hep.31216. Epub 2020 Sep 22. Impact Factor: 17,298 - Q1  
PMID: 32145077  
Citations: 3 [27/07/2022]

**Campillo-Marcos I**; Monte-Serrano E; Navarro-Carrasco E; García-González R; Lazo PA.  
Lysine Methyltransferase Inhibitors Impair H4K20me2 and 53BP1 Foci in Response to DNA Damage in Sarcomas, a Synthetic Lethality Strategy Front Cell Dev Biol. 2021 Sep 3;9:715126. doi: 10.3389/fcell.2021.715126. eCollection 2021. Impact Factor: 6,081 - Q1  
PMID: 34540832  
Citations: 1 [27/07/2022]

**Campillo-Marcos, I; Alvarez-Errico, D; Alandes, RA; Mereu, E; Esteller, M**  
Single-cell technologies and analyses in hematopoiesis and hematological malignancies Exp Hematol. 2021 Jun;98:1-13. doi: 10.1016/j.exphem.2021.05.001. Epub 2021 May 9. Impact Factor: 3,249 - Q3  
PMID: 33979683  
Citations: 3 [27/07/2022]

Cappetta, M; Fernandez, L; Brignoni, L; Artagaveytia, N; Bonilla, C; **Lopez, M; Esteller, M**; Bertoni, B; **Berdasco, M**  
Discovery of novel DNA methylation biomarkers for non-invasive sporadic breast cancer detection in the Latino population Mol Oncol. 2021 Feb;15(2):473-486. doi: 10.1002/1878-0261.12842. Epub 2020 Nov 19. Impact Factor: 7,4498 - Q1  
PMID: 33145876  
Citations: 4 [27/07/2022]

Carrato, C; Sanz, C; Munoz-Marmol, AM; Blanco, I; Pineda, M; Del Valle, J; Damaso, E; **Esteller, M; Musulen, E** The Challenge of Diagnosing Constitutional Mismatch Repair Deficiency Syndrome in Brain Malignancies from Young Individuals Int J Mol Sci. 2021 Apr 28;22(9):4629. doi: 10.3390/ijms22094629. Impact Factor: 6,208 - Q1  
PMID: 33924881  
Citations: 2 [27/07/2022]

Ciampa, I; Operto, G; Falcon, C; Minguillon, C; **De Moura, MC; Pineyro, D; Esteller, M**; Molinuevo, JL; Guigo, R; Navarro, A; Gispert, J; Vilor-Tejedor, N Genetic Predisposition to Alzheimer's Disease Is Associated with Enlargement of Perivascular Spaces in Centrum Semiovale Region Genes (Basel). 2021 May 27;12(6):825. doi: 10.3390/genes12060825. Impact Factor: 4,141 - Q2

PMID: 34072165  
Citations: 0 [27/07/2022]

**Coll-SanMartin, L; Davalos, V; Pineyro, D; Rossello-Tortella, M; Bueno-Costa, A; Setien, F; Villanueva, A; Granada, I; Ruiz-Xiviller, N**; Kotter, A; Helm, M; Yokota, J; Kawabata-Iwakawa, R; Kohno, T; Esteller, M Gene Amplification-Associated Overexpression of the Selenoprotein tRNA Enzyme TRIT1 Confers Sensitivity to Arsenic Trioxide in Small-Cell Lung Cancer Cancers (Basel). 2021 Apr 14;13(8):1869. doi: 10.3390/cancers13081869. Impact Factor: 6,575 - Q1  
PMID: 33919717  
Citations: 0 [27/07/2022]

Conley, BA; Staudt, L; Takebe, N; Wheeler, DA; Wang, LH; Cardenas, MF; Korchina, V; Zenklusen, JC; McShane, LM; Tricoli, JV; Williams, PM; Lubensky, I; O'Sullivan-Coyne, G; Kohn, E; Little, RF; White, J; Malik, S; Harris, LN; Mann, B; Weil, C; Tarnuzzer, R; Karlovich, C; Rodgers, B; Shankar, L; Jacobs, PM; Nolan, T; Berryman, SM; Gastier-Foster, J; Bowen, J; Leraas, K; Shen, H; Laird, PW; **Esteller, M**; Miller, V; Johnson, A; Edmondson, EF; Giordano, TJ; Kim, B; Ivy, SP The Exceptional Responders Initiative: Feasibility of a National Cancer Institute Pilot Study J Natl Cancer Inst. 2021 Jan 4;113(1):27-37. doi: 10.1093/jnci/djaa061. Impact Factor: 11,816 - Q1  
PMID: 32339229  
Citations: 9 [27/07/2022]

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**de Moura, MC; Davalos, V;** Planas-Serra, L; **Alvarez-Errico, D; Arribas, C;** Ruiz, M; Aguilera-Albesa, S; Troya, J; Valencia-Ramos, J; Velez-Santamaria, V; Rodriguez-Palmero, A; Villar-Garcia, J; Horcajada, JP; Albu, S; Casanovas, C; Rull, A; Reverte, L; Dietl, B; Dalmau, D; Arranz, MJ; Lluica-Carol, L; Planas, AM; Perez-Tur, J; Fernandez-Cadenas, I; Villares, P; Tenorio, J; Colobran, R; Martin-Nalda, A; Soler-Palacin, P; Vidal, F; Pujol, A; **Esteller, M**  
Epigenome-wide association study of COVID-19 severity with respiratory failure EBioMedicine. 2021 Apr;66:103339. doi: 10.1016/j.ebiom.2021.103339. Epub 2021 Apr 15.  
Impact Factor: 11,205 - Q1  
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**Esteve-Puig, R;** Climent, F; **Pineyro, D;** Domingo-Domenech, E; **Davalos, V;** Encuentra, M; Rea, A; Espejo- Herrera, N; **Soler, M; Lopez, M; Ortiz-Barahona, V;** Tapia, G; **Navarro, JT;** Cid, J; Farre, L; Villanueva, A; Casanova, I; Mangues, R; Santamarina-Ojeda, P; Fernandez, AF; Fraga, MF; Piris, MA; Kol, N; Avrahami, C; Moshitch-Moshkovitz, S; Rechavi, G; Sureda, A; **Esteller, M**  
Epigenetic loss of m(1)A RNA demethylase ALKBH3 in Hodgkin lymphoma targets collagen, conferring poor clinical outcome  
Blood. 2021 Feb 18;137(7):994-999. doi: 10.1182/blood.202005823.  
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Farre L; Sanz G; **Ruiz-Xivillé N; Moura MC;** Tejera JFM; Gonçalves-Ribeiro S; Martinez-Iniesta M; Calaf M; Mosquera JL; Martín-Subero JI; **Granada I; Esteller M;** Domingo-Domenech E; Climent F; Villanueva A; Sureda A. Extramedullary multiple myeloma patient-derived orthotopic xenograft with a highly altered genome: combined molecular and therapeutic studies  
Dis Model Mech. 2021 Jul 1;14(7):dmm048223. doi: 10.1242/dmm.048223. Epub 2021 Jul 15.  
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PMID: 33988237  
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Fernández-Figueras MT; Carrato C; Saenz-Sardà X; **Musulén E;** Fuente MJ; Puig L.  
MicroRNA31 and MMP-1 contribute to the differentiated pathway of invasion -with enhanced epithelial-to- mesenchymal transition- in squamous cell carcinoma of the skin  
Arch Dermatol Res. 2021 Oct 13. doi: 10.1007/s00403-021-02288-x. Online ahead of print.  
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PMID: 34647185  
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Fernandez-Sanles, A; Sayols-Baixeras, S; Subirana, I; Senti, M; Perez-Fernandez, S; **Moura, MD; Esteller, M;** Marrugat, J; Elosua, R  
DNA methylation biomarkers of myocardial infarction and cardiovascular disease  
Clin Epigenetics. 2021 Apr 21;13(1):86. doi: 10.1186/

s13148-021-01078-6.  
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Fontana, A; Barbano, R; Dama, E; Pasculli, B; Rendina, M; Morrilli, MG; Melocchi, V; Castelveter, M; Valori, VM; Ravaioli, S; Bravaccini, S; Ciuffreda, L; Graziano, P; Maiello, E; Copetti, M; Fazio, VM; **Esteller, M;** Bianchi, F; Parrella, P  
Combined analysis of miR-200 family and its significance for breast cancer  
Sci Rep. 2021 Feb 3;11(1):2980. doi: 10.1038/s41598-021-82286-1.  
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**Garcia-Gomez, A; Li, TL; de la Calle-Fabregat, C; Rodriguez-Ubreva, J; Ciudad, L; Catala-Moll, F; Godoy- Tena, G;** Martin-Sanchez, M; San-Segundo, L; Muntion, S; Morales, X; Ortiz-de-Solorzano, C; Oyarzabal, J; Jose- Eneriz, ES; **Esteller, M;** Agirre, X; Prosper, F; Garayoa, M; **Ballestar, E**  
Targeting aberrant DNA methylation in mesenchymal stromal cells as a treatment for myeloma bone disease  
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Impact Factor: 17,694 - Q1  
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Garcia-Ruiz, B; **de Moura, MC;** Muntane, G; Martorell,

L; Bosch, E; **Esteller, M;** Canales-Rodriguez, EJ; Pomarol- Clotet, E; Jimenez, E; Vieta, E; Vilella, E  
DDR1 methylation is associated with bipolar disorder and the isoform expression and methylation of myelin genes Epigenomics. 2021 Jun;13(11):845-858. doi: 10.2217/epi-2021-0006. Epub 2021 May 4.  
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Garcia, P; Fernandez-Hernandez, R; Cuadrado, A; Coca, I; Gomez, A; Maqueda, M; Latorre-Pellicer, A; Puisac, B; Ramos, FJ; Sandoval, J; **Esteller, M;** Mosquera, JL; Rodriguez, J; Pie, J; Losada, A; Queralt, E  
Disruption of NIPBL/Sccl in Cornelia de Lange Syndrome provokes cohesin genome-wide redistribution with an impact in the transcriptome  
Nat Commun. 2021 Jul 27;12(1):4551. doi: 10.1038/s41467-021-24808-z.  
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Serrated epithelial lesion with dysplasia as a precursor to small bowel carcinoma associated with Crohn's disease Gastroenterol Hepatol. 2021 Aug-Sep;44(7):489-490. doi: 10.1016/j.gastrohep.2020.11.015. Epub 2020 Dec 5.  
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Stem Cells  
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DNA methylome in visceral adipose tissue can discriminate patients with and without colorectal cancer  
Epigenetics. 2021 Jul 26;1-12. doi: 10.1080/15592294.2021.1950991. Online ahead of print.  
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