



Directors' Message

Presentation

At a Glance

2020 Organization	9
Scientific Output	10
Funding	
Technology Transfer	11
Training & Outreach Activity	11
Awards & Recognitions	12

Knowledge & Technology Transfer

Patents	34
Spin-off	35
Research Agreements	
with Companies	36

Scientific Output

Selected Articles	38
Highlights	39
Edited Books & Book Chapters	43
Cover Pages	- 44
Editorial Activity	46
Full Publication List	47

14 Organization

General Assembly	15
International Scientific	
Advisory Board	15
Research Groups	16
Directorate & Management Area	20
	General Assembly International Scientific Advisory Board Research Groups Directorate & Management Area

Facilities

Molecular & Functional	
Imaging Facility	24
Technological Platforms	26

Funding

List of Funded Projects	
Starting in 2020	29
Total Funding	32

Training

PhD Programs	59
PhD Theses	60
Master & Vocational	
Training Programs	61
Research Secondments &	
Summer Interships	62
Seminars	62
Organization of Scientific	
Workshops/Conferences	64

66 Outreach

Open Days & Visits	71
Media	71

Directors' Message

2020 has not been a regular year in any respect. The unexpected COVID-19 pandemic has altered our lives in many ways during 2020, and research is not an exception. During several periods of the year, we were instructed such unusual measures as complete lockdown, severe limitations in the occupancy of labs and offices, as well as strong restrictions against interpersonal relationships and inperson meetings. The Direction of CIC biomaGUNE often had to implement new regulations on a very short term and most likely not always in the most efficient manner, but always aiming to keep health and safety of our personnel above everything else. We are happy that none of us suffered any severe infection so far and we hope it will stay like this. We would especially like to thank those who volunteered to work during the worst, early period of the pandemic, to assist the local hospitals when there was insufficient supply from companies. We should also thank the whole CIC biomaGUNE community for strictly following the (changing) directions and constantly cooperating and adapting to new work conditions.

In spite of all restrictions, communication difficulties, lack of travel and even severe lockdown, we can be most proud about the overall performance of our researchers. We had excellent scientific results, with a large amount of high (some outstanding) quality publications and patents, unusually high success in EU funding, as well as a record number of contracts with companies, which was only possible thanks to the fabulous support by our administration, IT and maintenance teams, as well as a great team spirit leading to plenty of internal collaborations. Still, this report will clearly show that some of the activity was hindered by the pandemic. In particular, we could hardly host visiting students and researchers, our seminar program was completely stopped for months and then restarted in an online modality, outreach activities were also heavily affected by difficulties in bringing people together. We are hoping that coming years will bring back our normal way of doing things, even though we will probably see some changes related to the recent experience, such as increased ratio of remote work or reduced travel schedules, we'll see.

We are also proud to have approved in 2020 the 1st Equality Plan reinforcing our commitment toward equality.

We invite you to browse through this report to exciting publications that were possible thanks to the enthusiasm and excellent performance of staff at CIC biomaGUNE, as well as our collaborators, both locally and internationally.

Kind regards,

Luis M. Liz-Marzán, Scientific Director José M. Mato, Director General



José M. Mato, Director General (left) and Luis M. Liz-Marzán, Scientific Director (right)



Presentation

The Center for Cooperative Research in Biomaterials **CIC biomaGUNE**, member of the Basque Research and Technology Alliance (**BRTA**), is a non-profit research organization created to promote scientific research and technological innovation at the highest levels in the Basque Country, following the BioBasque policy in order to create a new business sector based on biosciences.



Located in the Science and Technology Park of Gipuzkoa (Donostia-San Sebastián), the activity of CIC biomaGUNE is conducted by a team of 12 international and dynamic research groups, which develop high-level research at the interface between chemistry, physics and biology, with particular emphasis on the properties and applications of nanostructures at the biomolecular level. The final aim of CIC biomaGUNE's research is to contribute to understanding the way in which biological systems interact with nanometer-sized materials, at the molecular level. The main research lines deal with the design, preparation and characterization of biofunctional nanostructures and their *in vitro* and *in vivo* biological evaluation, to be used in the study of biological processes and the development of biomedical tools such as theranostic, multitherapeutic or multimode imaging platforms.

To carry out this ambitious program of research, the Center counts with a unique research infrastructure, equipped with advanced nanoscience, biochemistry and molecular imaging facilities, including fully



equipped research laboratories, Technological Platforms and the Molecular Imaging Facility, one of the most complete preclinical imaging research infrastructures in Europe.

CIC biomaGUNE started its activity in December 2006. In the course of almost fourteen years, with an average critical mass of around 120 researchers, the Center has obtained national and international recognition as a scientific leader and knowledge builder in the field of biomaterials. Among other achievements, our activity has resulted in the generation of knowledge (publications, patents, PhD theses), the internationalization of research through collaborations with international bodies and institutions, as well as opening our facilities to the scientific community and the industrial sector.

In 2018 CIC biomaGUNE earned the accreditation as a "María de Maeztu Unit of Excellence", the highest recognition of scientific excellence in Spain awarded by the Spanish State Research Agency (AEI) - which further confirms this recognition.

At a Glance



2020 Organization

Research Groups

- 1 Glycotechnology
- 2 Biomolecular Nanotechnology
- 3 Soft Matter Nanotechnology
- 4 Bionanoplasmonics
- 5 Bioengineered Particles
- 6 Carbon Bionanotechnology
- 7 Heterogeneous Biocatalysis
- 8 Regenerative Medicine
- 9 Computational Biophysics
- 10 Radiochemistry & Nuclear Imaging
- 11 Magnetic Resonance Imaging
- 12 Molecular & Functional Biomarkers

2 Molecular & Functional Imaging Facility

- 1 Radiochemistry
- 2 Nuclear Imaging (PET/SPECT/CT)
- 3 Magnetic Resonance
- Imaging (MRI) 4 Image Analytics
- E Animal Llause
- 5 Animal House

Technological Platforms

- 1 Nuclear Magnetic Resonance
- 2 Electron Microscopy
- **3** Mass Spectrometry
- 4 Surface Analysis & Fabrication
- 5 Colloidal Nanofabrication
- 6 Optical Spectroscopy

Support

- 1 Direction
- 2 General Management
- 3 Administration, Financial & HR Management
- 4 Project Management
- 5 Tech Transfer & Exploitation
- 6 Biosafety & Radioprotection
- 7 Information Technology
- 8 Maintenance



Scientific Output





Publications by gender



Funding



Public



Technology Transfer



Training and Outreach Activiy



11

Private

AWARDS & RECOGNITIO

Accreditations

AAALAC Accreditation to CIC biomaGUNE's animal facilities

CIC biomaGUNE received the AAALAC accreditation in 2015 for the first time and the accreditation was renewed in 2018.

Maria de Maeztu Unit of

Excellence accredited by the Spanish State Research Agency.

The Molecular & Functional Imaging Facility remains accredited as **Singular Scientific and Technical Infrastructure** (ICTS in Spanish), as a node of the ReDIB network.

Certification of CIC biomaGU-NE RD&I management system according to **UNE 166002:2014** standard.

Awards

Maurizio Prato,

Luis M. Liz-Marzán,

"Miguel Catalán-Paul

awarded with the prestigious **ERC-AdG** for his project e-DOTS. 31/03/2020

2020 SCF French-Spanish

Sabatier" Prize in recognition

of his eminent works in the

application of colloid chemistry to the field of nanoplasmonics. It is

also in recognition of the strong

links that he established with the



30/06/2020

Pedro Ramos-Cabrer,

French chemists.

awarded with the XVIII Research Prize of the Medical College of Córdoba, as one of the 3 best research publications during the year 2019, co-authored by at least one doctor of Spanish nationality.

18/11/2020

Wolfgang J. Parak, Highly Cited researcher (Clarivate Analytics) in the

Cross-field category.



20/11/2020

Luis M. Liz-Marzán, Highly Cited researcher (Clarivate Analytics) in the Cross-field category.





20/11/2020



CIC biomaGUNE





Best Poster

Pedro Ramos-Cabrer

The 15th European Molecular Imaging Meeting (vEMIM) has awarded the "**Best v-EMIM Poster-Pitches**" prize to the work "Longitudinal rs-fMRI study of functional alterations during demyelination and remyelination in the cuprizone murine model" from The Magnetic Resonance Imaging Lab.

24/08/2020



Rossana Passannante

The work entitled "Pharmacokinetics evaluation of new drugs with potential application in Duchenne muscular dystrophy", presented by Rossana Passannante, PhD student of the Radiochemistry and Nuclear Imaging Group received the award for the **Best Imaging Probes & Applications Poster-Pitch** within the 15th European Molecular Imaging Meeting-EMIM2020. Best Imaging Probes & Applications Poster-Pitch.

01/09/2020

Academy Membership



Luis M. Liz-Marzán, joins the Royal Galician Academy of Sciences (RAGC - Real Academia Gallega de Ciencias) as a Corresponding Member.

21/01/2020



Fernando López-Gallego, Ikerbasque Professor at CIC biomaGUNE joins the Young Academy of Spain.

04/06/2020



Organization



General Assembly



International Scientific Advisory Board



Prof. Peter Morris

Sir Peter Mansfield Magnetic Resonance Chemistry Centre University of Nottingham, UK



Prof. Monica Olvera de la Cruz

Department of Northwestern University, USA



Prof. Itamar Willner

Institute of Chemistry The Hebrew University of Jerusalem, Israel



Prof. Peter Seeberger

Max-Planck Institute of Colloids and Interfaces, Germany



Prof. Aránzazu del Campo

INM - Leibniz Institute for New Materials, Germany



Prof. Samuel I. Stupp

Institute for BioNanotechnology in Medicine Northwestern University, USA



Prof. Patrick Couvreur Université Paris-Sud. France

Research Groups

The main research lines revolve around the design, preparation and characterization of biofunctional nanostructures, to be used in the study of biological processes and the development of biomedical tools.



Sergio Moya - Principal Investigator

Glycotechnology

Niels Reichardt - Principal Investigator Sonia Serna - Research Associate

The Glycotechnology Laboratory carries out projects in basic and applied glycoscience, the science and technology of carbohydrates with a large untapped potential to provide innovative solutions to important social challenges such as personalized medicine, pharmaceutical products, food and biomaterials.

The group applies carbohydrate synthesis, materials science and molecular biology to the design of tools, probes and devices to elucidate the role and exploit the potential of sugars for cancer immune therapy, as biomarkers and biomaterials for biomedical applications.

Biomolecular Nanotechnology

Aitziber L. Cortajarena - Principal Investigator (Ikerbasque Professor) Ivan R. Sasselli - Research Associate (Fellow Gipuzkoa)

erc

The group focuses on protein engineering toward the generation of functional nanostructures and bioinspired materials for applications in nanobiotechnology and nanomedicine. The research focuses mainly on protein engineering in order to develop versatile platforms for the bottom-up fabrication of proteinbased hybrid functional biomaterials. We are also interested in the tailored biofunctionalization of nanomaterials for biomedical applications, from disease treatment to diagnosis.

The group also carries out research aimed at the development of new analytical and bioanalytical techniques, using metal and semiconductor nanoparticles and has also focused on the fabrication of novel biosensors employing different read-out methods, such as UV-visible and fluorescence spectroscopy, quartz crystal microbalance, electrochemistry and photo-electrochemistry. The Soft Matter Nanotechnology Laboratory makes use of elements of soft matter, mainly polyelectrolytes, in nanofabrication and in the development of hybrid materials for biomedical applications.

The group has expertise in several physico-chemical characterization techniques, especially in fluorescence spectroscopy. The group interests include the synthesis of polyelectrolytes, self-assembly, physical characterization, the development of drug delivery systems, and hybrid materials for tissue engineering.



Bionanoplasmonics

erc

Luis M. Liz-Marzán - Principal Investigator (Ikerbasque Professor) Isabel García-Martín - Research Associate (CIBER-BBN)

Dorleta Jiménez de Aberasturi - *Research Associate (Ikerbasque Fellow)* Oscar F. Silvestre - *Research Associate (Fellow Gipuzkoa)*

Malou Henriksen-Lacey - Research Associate The activity of the Bionanoplasmonics Laboratory focuses on the biomedical applications of plasmonic nanomaterials, including new chemical methods for the synthesis of colloidal metal nanoparticles with tailored size, shape and surface chemistry, their directed self-assembly and applications in biosensing, diagnostics and therapy, mainly based on plasmonic effects. One of the current central topics of the

group is the development of platforms that can be used for ultrasensitive detection based on SERS. The group is interested in the incorporation of such nanostructured substrates within devices for implementation of real detection techniques.



Wolfgang J. Parak - Principal Investigator Carlos Sánchez-Cano - Research Associate (Fellow Gipuzkoa)

The Bioengineered Particles Laboratory focuses on understanding the interaction of colloidal nanomaterials with the biological matter, such as proteins and cells.

Our work is dedicated to the synthesis of highly defined nanoparticle libraries, their physicochemical characterization, and correlation of biological effects to their physicochemical properties. We also develop new methods toward novel methodologies for physicochemical characterization *in situ* and in complex environments.

Carbon Bionanotechnology



Maurizio Prato - Principal Investigator (Ikerbasque Professor & AXA Chair)

The mission of the Carbon Bionanotechnology Laboratory is the design and synthesis of tailored carbon nanostructures for bio-nanotechnology applications and solar energy conversion through biomimetic approaches.

The group explores new synthetic protocols and new analytical methods, enabling innovative, controlled and reproducible ways toward the designer functionalization of carbon nanostructures, such as fullerenes, carbon nanotubes, graphene and carbon nanodots.

Regenerative Medicine

Ander Abarrategi - Junior Group Leader (Ikerbasque and Ramón y Cajal Fellow) The Regenerative Medicine Laboratory uses biomaterial-based approaches to

boost knowledge in stem-cell biology,

both in physiological and in pathological contexts. For this aim, we

generate bioactive and cell-laden 3D

structures potentially useful for

Heterogeneus Biocatalysis



Fernando López-Gallego- Principal Investigator (Ikerbasque Professor)

The Laboratory of Heterogeneous Biocatalysis is applying multi-enzyme systems to synthetic, environmental, medical and analytical chemistries by harnessing the exquisite selectivity of enzymes (biological catalysts) for the development of more sustainable and effective chemical processes. We are mimicking the spatial organization found inside the living organisms, but using ex-vivo systems supported on solid materials. To address such goal, we are interfacing chemistry and biology utilizing multidisciplinary tools that involve molecular biology, enzymology and materials chemistry.

regenerative medicine and disease modelling studies. The understanding of bone tissue is the core of our research. Briefly, we design, characterize and test different kinds of implantable devices to gain insight into specific tissue formation processes. From this information we define and modulate relevant mechanisms in the context of tissue regeneration and tissue pathology.

Computational Biophysics

Ivan Coluzza - *Principal Investigator* (Ikerbasque Professor)

Ivan R. Sasselli – *Research Associate* (*Fellow Gipuzkoa*)

The Computational Biophysics Laboratory is composed by an interdisciplinary team of scientists from different backgrounds but all with experience in computational modeling of biological systems and statistical mechanics.

Our research focuses on the application of statistical mechanics to soft-matter and complex biological systems.

Our goal is to build simple models of natural complex systems, such as proteins, and in doing so learn their fundamental function and copy it into artificial systems.

Radiochemistry & Nuclear Imaging

Jordi Llop - Principal Investigator

The activity of the Radiochemistry and Nuclear Imaging Laboratory focuses on the development of innovative radiochemistry and the application of positron emission tomography (PET) and single photon emission computed tomography (SPECT) tracers toward the investigation of biological, physiological and pathological processes in the fields of oncology, neurology, pneumology, infection and cardiovascular diseases.



Magnetic Resonance Imaging

Pedro Ramos-Cabrer - Principal

Investigator (Ikerbasque Professor) The Magnetic Resonance Imaging (MRI) Laboratory makes use of nanomaterials and magnetic resonance imaging techniques on animal models.

On one hand, we intend to characterize the onset and evolution of diseases of the central nervous system (CNS), from development of early markers to imaging methods that quantify the progression of the pathological processes and their consequences at anatomical and functional levels.

We additionally develop new therapeutic approaches to treat such diseases, with special emphasis on the penetration through the blood- brainbarrier and monitoring the effective release of drugs in the brain parenchyma.

Molecular & Functional Biomarkers

Jesús Ruiz-Cabello - Principal Investigator (Ikerbasque Professor) Susana Carregal-Romero - Research Associate (CIBER-BBN)

The Molecular & Functional Biomarkers Laboratory studies cardio-pulmonary and vascular diseases through functional and molecular imaging and system biology approaches.

The group is particularly interested in the potential of new imaging techniques, including nanotechnologybased applications, in early diagnosis of pulmonary and cardiovascular remodelling, the assessment of metabolic changes associated with cell growth, the structure and function of the right ventricle and cardiovascular coupling signals.

Activity Report 2020

Directorate

R&D&I

Transfer Manager

particular,

Management Unit

Cristina Díez García - Project Manager

A dedicated office to support and

strengthen the capacities of the Center,

in terms of attracting funding from

various Research Programs, in

strengthening links between academic and industrial environments, and promoting the transfer of research results to society and industry, especially to the biotechnology sector.

international

ones,

Marcos Simón Soria - Technology

José M. Mato - Director General Luis M. Liz-Marzán - Scientific Director Design, define and coordinate the Center's scientific strategies and activities.

General Management

Anna Llanes Pallàs - General Manager The General Manager is responsible for supervising the management of the Administration Department, Project Management, as well as the IT, Maintenance and Biosafety units of

the Center.

Administration

Sheyla García Medel - Administration & HR Manager

Elizabeth Noguera Olaechea - Finance & Control Manager

This department is responsible for the management of finances, accounting, administration, and the human resources of the Center.

Biosafety & Radioprotection

Paola Ferreira Cabeza - Biosafety & Radioprotection Manager

Dedicated to establish safe working conditions of all CIC biomaGUNE's personnel by promoting good laboratory practices. The service is also in charge of the appropriate operation of the Center's Radioactive Facility.

Computing & Communications

Mikel Gonzalez Lacunza - *IT Manager* The IT service is in charge of supporting the staff of the Center with setting up and maintaining computer-related equipment, software, data storage, e-mail servers, as well as the website and other social media.

Maintenance

Álvaro Ruiz Fernández - Maintenance Manager

This department takes care of the preventive, predictive and corrective maintenance of all facilities at CIC biomaGUNE.

CIC biomaGUNE's Personnel during 2020









All research lines, framed within the strategic research program of CIC biomaGUNE, are strongly supported by the Molecular & Functional Imaging Facility and the Technological Platforms, which constitute a major strength of the Center.

CIC biomaGUNE Technological Platforms & Molecular Imaging Facility

CIC biomaGUNE's state-of-the art facilities are depicted in the images below.





Housed within 900 m², the Molecular and Functional Imaging Facility at CIC biomaGUNE is an integrated bioimaging structure that offers state-of-the-art preclinical imaging instrumentation in Positron Emission Tomography (PET), Single Photon Emission Computed Tomography (SPECT), Computerized Tomography (CT), Magnetic Resonance Imaging (MRI), Optical/Fluorescence and Ultrasound (US) Imaging. It includes a fully equipped radiochemistry laboratory with a biomedical cyclotron, advanced microscopy equipment, and dedicated animal housing facility for rodents which holds AAALAC accreditation. The Facility is currently integrated in the "Distributed Biomedical Imaging Network" (ReDIB, www. redib.net), recognized by the Spanish Government as a Singular Scientific and Technical Infrastructure (ICTS). The infrastructure has been designed, built and equipped to tackle longitudinal and multimodal pre-clinical projects and to develop applications in the area of Preclinical Molecular and Functional Imaging and Nanomedicine.

Nuclear Imaging

Unai Cossío - *Platform Manager* Equipped with a hybrid PET-CT (eXplore Vista-CT) a full ring SPECT-CT (eXplore speCZT CT 120), the latter offering the possibility of multi-isotope studies with energy discrimination, and a trimodal PET-SPECT-CT.

Magnetic Resonance Imaging (MRI)

Daniel Padró - Platform Manager

Instrumentation to conduct advanced imaging and spectroscopic experiments applied to biological samples including small rodents, samples or tissue extracts and cell cultures is available. CIC biomaGUNE provides the instrumentation and the expertise to carry out a wide range of MRI and MRS experiments, and is equipped with surgery rooms for animal preparation and implementation of surgical models. Ancillary equipment is also available: anesthesia systems, MRI-compatible physiological monitoring systems, infusion pumps, and temperature regulation systems. Available equipment: MRI 7T/30 cm (70/30 USR), MRI 11.7T/16 cm (117/16 USR).



Radiochemistry Platform

Vanessa Gómez-Vallejo - Platform Manager

The platform is equipped with an IBA Cyclone 18/9 cyclotron able to accelerate protons (18 MeV) and deuterons (9 MeV) and is equipped with 7 targets for the routine production of [¹⁸F] F⁻, [¹⁸F] F⁻, [¹³N]-NH⁴⁺⁻, [¹⁵O]-O₂ [¹¹C]-CO₂ and [¹¹C]-CH₄. It also has a solid target for the production of ⁸⁹Zr and ⁶⁴Cu.

The radiochemistry laboratory equipped with 5 shielded hot cells housing versatile automatic synthesis, suitable for the production (synthesis, purification and quality control) of PET and SPECT radiotracers. The facility has specially designed modules for:

- Synthesis of [¹¹C] $CH_3I / [^{11}C] CH_3OTf$ from [¹¹C] $CO_2 / [^{11}C] CH_4$, and subsequent methylation reaction.
- ¹⁸F-fluorination by nucleophilic and electrophilic substitution.
- Radiotracer synthesis using microfluidics technology.
- Chelation reactions using radiometals (⁶⁸Ga, ⁶⁷Ga, ⁶⁴Cu, ⁸⁹Zr, etc.).

The quality control lab, sited into the production lab, is equipped with state of art equipment to perform complete quality control of synthesized radiotracers, including radio-HPLC, radio-GC, radio-TLC, and gamma spectrometry.

Pre-Clinical Analytics

Unai Cossío - Platform Manager

The Image Analytics service takes care of processing all the outcoming multimodal images obtained within the Molecular Imaging Unit (PET, SPECT, CT, and MRI). Working on different operating systems, we carry out coregistration, segmentation, and quantification of multimodal images. The combination of all multimodal imaging techniques confined in the Imaging Unit with a reliable image analysis process offers a complete and powerful Imaging facility to researchers. Four workstations and a data storage system in the Terabyte scale enable image reconstruction, processing, quantification and archiving.

Animal Facility

Ainhoa Cano - Platform Manager

The MRI Unit and the Nuclear Imaging Unit sandwich a dedicated animal housing area, which holds accreditation of the **Association for Assessment and Accreditation of Laboratory Animal Care International (AAALAC)** and is prepared to house up to 800 mice and 400 rats in individually ventilated cages, with an experimental area specifically devoted to perform longitudinal studies. The animal house is complemented with microsurgery areas for animal preparation.

Technological Platforms

Managed by specialized Platform Managers, they provide technical and scientific support to the research activities conducted at CIC biomaGUNE. They include the following stateof-the-art research infrastructures:

Mass Spectrometry

Javier Calvo - Platform Manager Equipped with **MALDI-TOF, ICP-MS**, it provides several high quality mass spectrometry techniques for the analysis of small molecules, complex biomolecules and nanomaterials. The platform is equipped with modern instrumentation and offers different ionization methods.

Electron Microscopy

Marco Möller - Platform Manager Equipped with SEM-EDX, TEM – 120 keV and 200 keV, offers techniques to study nanoparticles and biological or soft polymer materials (cryo-TEM) at the micrometer and nanometer scale to determine the materials dimensions, shape and composition.

Nuclear Magnetic Resonance (NMR)

Daniel Padró - Platform Manager Equipped with a **500 MHz NMR spectrometer**, it provides essential service for the characterization of molecules with biological activity, from complex glycans to molecules used in the design of nanostructures for biomedical applications.



Surface Analysis & Fabrication

Desiré Di Silvio - *Platform Manager* Equipped with *XPS*, *3 x AFM*, *sputtering system*, it focuses on the analysis of materials at the surface level, including spectroscopic and microscopic techniques especially suited to surfaces. The platform offers also services for the deposition of thin layers with controlled manufacture at the nanoscale.

Colloidal Nanofabrication

Ana Sánchez-Iglesias - Platform Manager

Within the priority area of biofunctional nanomaterials and nanomedicine, colloidal synthesis plays an essential role. This platform provides knowledge, training and service for the manufacture of nanoparticles with a wide variety of compositions and morphologies.

Optical Spectroscopy & Imaging

Irantzu Llarena - Platform Manager Judith Langer - Platform Manager Equipped with 2 x Confocal, and 2 x Confocal-Raman microscopes, UV-VIS-NIR, and FT-IR spectrometers, Fluorimeter, Flow Cytometer, Cell Observer, CD, DLS, DCS, ITC, TGA, SPR, this platform offers a wide variety of techniques for the spectroscopic characterization of biomaterials and biosurfaces. Also included are optical microscopy techniques (confocal, Raman, etc.), to investigate the interaction of nanomaterials with cellular systems of diverse complexity.







Funding



The total funding for 2020 has amounted 11,868,675€ 45 new projects (from competitive funding sources) have been launched in 2020 with a total contribution of 7,731,721€

	ZA			
CALL	PI (Fellow)	Full Title	Period	Amount (€)
Elkartek 2019 - 2020	Jesús Ruiz-Cabello	biomaGUNE 2019 - Biomarcadores para el diagnóstico precoz y seguimiento de tratamiento en hipertensión pulmonar	2020-2021	576,430.90
Elkartek 2020 - 2020	Sergio Moya	biomaGUNE 2020 –Desarrollo de vectores no virales para terapia génica con aplicaciones oncológicas y en inmunoterapia	2020-2021	1,020,336.80
Ayudas a proyectos de investigación y desarrollo en salud 2020	Aitziber L. Cortajarena	Glioblastoma – Medicina personalizada en tumores cerebrales: desarrollo y validación de estrategias terapéuticas basadas en la inhibición de la auto-renovación	2020	13,460.40
Ayudas a proyectos de investigación y desarrollo en salud 2020	Aitziber L. Cortajarena	NKTPH - Papel de las células NK en el trasplante de precursores hematopoyéticos autólogo en el tratamiento de cáncer y enfermedades raras	2020	22,336.10
Ayudas a proyectos de investigación y desarrollo en salud 2020	Ander Abarrategi	Fragilidad - Validación de un patrón molecular para el diagnóstico y estratificación de la fragilidad	2020	11,202.21
Ayudas a proyectos de investigación y desarrollo en salud 2020	Ander Abarrategi	EXA MINA - Dolor y pérdida de la capacidad funcional en las personas mayores: Desarrollo de procedimientos y tecnologías para su valoración integral	2020	12,223.16
BIKAINTEK2020 – Contratación de doctores	Aitziber L. Cortajarena/Nan Coluzza (Giulia Magi Meconi)	SwitchProt - Diseño de novo de andamios de proteínas con estructuras intercambiables	2020-2022	46,471.68
BIKAINTEK2020 – Contratación de doctores	Fernando López-Gallego (Alejandro Herrero Orrero)	KIMICoA - Fabricación de biocatalizadores heterogéneos multifuncionales para la síntesis sostenible de precursores de coenzima A con aplicaciones en química fina y médica	2020-2022	46,471.68
Ayudas urgentes I+D COVID-19_2020	Niels C. Reichardt	Biochip COVID-19 - Desarrollo de microarrays de anticuerpos y antígenos como plataforma para el diagnóstico y estudio epidemiológico de Covid-19	2020	104,123.38
Ayudas urgentes I+D COVID-19_2020	Aitziber L. Cortajarena	Nano-CoV - Plataformas nanotécnológicas para la detección rápida de anticuerpos anti-SARS-CoV-2: diagnostico y evaluación de desarrollo de	2020	90,000.00
Equipamiento Científico_EKIZIEN 2020	Maurizio Prato	inmunidad Espectrómetro de infrarrojo (IR) por Transformada de Fourier (FT)	2020	90,000.00



CALL	PI (Fellow)	Period	Amount (€)
Ayudas para contratos predoctorales 2019	Aitziber L. Cortajarena (Laura Pérez)	2020-2024	98,250.00
Ayudas para contratos predoctorales 2019	Jordi Llop (Óscar Moreno)	2020-2024	98,250.00
Ayudas para contratos predoctorales 2019	Fernando López-Gallego (Francisco Javier Santiago-Arcos)	2020-2023	98,250.00
Juan de la Cierva - Formación 2018	Luis M. Liz-Marzán (Xiaolu Zhuo)	2020-2022	50,000.00
Juan de la Cierva - Formación 2018	Maurizio Prato (Alessandro Silvestri)	2020-2022	50,000.00
Juan de la Cierva - Incorporación 2018	Maurizio Prato (Huilei Hou)	2020-2023	93,000.00
Ramón y Cajal 2018	Ander Abarrategi	2020-2024	208,600.00

CALL	PI	Full Title	Period	Amount (€)
Proyectos de I+D+i 2019 "Retos Investigación"	Aitziber L. Cortajarena	ProTOOLS - Protein engineering to generate advanced tools for biomedical and biotechnological applications	2020-2023	175,450.00
Proyectos de I+D+i 2019 "Retos Investigación"	Maurizio Prato	Multi-DOTS - Tailored Carbon Nanodots as Novel and Safe Multifuncional Materials for Nano- and Bio- Applications	2020-2023	181,500.00
Proyectos de I+D+i 2019 "Retos Investigación"	Dorleta Jiménez de Aberasturi	InOrINK-3D - Smart bioinks based on multifunctional inorganic-organic composites and application to a 3D- printed pulmonary artery model	2020-2023	96,800.00
"Redes de Investigación" 2018	Fernando López-Gallego	REDBIOCAT - Red temática de biocatálisis y tecnología enzimática; soluciones hacia una química sostenible	2020-2022	6,000.00
"Redes de Investigación" 2018	Luis M. Liz-Marzán	SOMMa+ - Alianza de Centros Severo Ochoa y Unidades Maria de Maeztu - SOMMA	2020-2021	
"Redes de Investigación" 2018	Jesús Ruiz-Cabello	nanoCARE - Red de nanomedicina para el diagnóstico y tratamiento de enfermedades de alto impacto social: cancer, aterosclerosis y enfermedades infecciosas	2020-2021	
"Redes de Investigación" 2018	Aitziber L. Cortajarena	BIONAPEP - Red de Péptidos en Biomedicina y Nanociencia	2020-2021	



European Commission				
CALL	PI	Full Title	Period	Amount (€)
H2020 H2020-NMBP- ST-IND-2018-2020	Maurizio Prato	GrapheneCore3 - Graphene Flagship Core 3	2020-2023	363,750.00
H2020 H2020-SGA-FET- GRAPHENE-2017	Maurizio Prato	DECADE - DistributEd Chemicals And fuels production from CO ₂ in photoelectrocatalytic Devices	2020-2024	320,746.25
H2020-FETOPEN-2018- 2020	Aitziber L. Cortajarena	ARTIBLED - Engineered Artificial Proteins for Biological Light-Emitting Diodes	2020-2023	484,636.25
H2020-FETOPEN-2018- 2020	Luis M. Liz-Marzán	POSEIDON - NanoPhOtonic devices applying SEIf-assembled colloIDs for novel ON-chip light sources	2020-2023	246,031.25
H2020-MSCA-ITN-2019	Jesús Ruiz-Cabello	NOVA-MRI - Novel Applications in 19F Magnetic Resonance Imaging	2020-2024	250,904.88
H2020-MSCA-ITN-2019	Fernando López-Gallego	INTERfaces - Heterogeneous biocatalytic reaction cascades training network	2020-2023	250,904.88

Gipuzkoako Foru Aldundia Diputación Foral de Gipuzkoa				
CALL	PI (Fellow)	Full Title	Period	Amount (€)
Programa RED – Investigación 2020	Pedro Ramos-Cabrer	BBB-trans - Biomateriales y protocolos efectivos para el transporte de fármacos al sistema nervioso central	2020-2021	90,804.00
Programa RED – Investigación 2020	Aitziber L. Cortajarena	Espectrofluorímetro - Adquisición de una Espectrofluorímetro	2020-2021	47,150.00
Programa Fellows Gipuzkoa 2020	Wolfgang Parak (Carlos Sánchez-Cano)	NanoBioMeSi - Nano-materiales metálicos en muestras Biológicas; explorando interacciones entre Metales usando radiación de Sincrotrón	2020-2021	35,000.00
Programa Fellows Gipuzkoa 2020	Aitziber L. Cortajarena/ Ivan Coluzza	EINPySB - Estudio de la Interfase entre Nanomateriales Peptídicos y Sistemas Biológicos	2020-2021	39,492.00
Programa Fellows Gipuzkoa 2020	Luis M. Liz-Marzán (Oscar Ferreira Silvestre)	NanoMito4Imaging - Nanoterapias dirigidas a la mitocondria y herramientas de imagen de precisión avanzadas para la administración de medicamentos	2020-2021	39,492.00
Ayudas para la investigación contra enfermedades severas 2020	Niels C. Reichardt	Robot pipeteo - Proyecto de Adquisición de un Robot de Pipeteo para Plataforma Diagnóstica COVID-19	2020-2021	29,323.00
Acompañamiento en el proceso de creación de empresas de base tecnológica y/o innovadoras 2020	Aitziber L. Cortajarena	Serket BioMed – Proyecto de acompañamiento en el proceso de creación de una empresa de base tecnológica	2020-2021	50,000.00
Promoción del talento y el aprendizaje de las personas en las empresas 2020	CIC biomaGUNE	bmG-HRS4R – Proyecto de Implantación del Sello de Excelencia en Recursos Humanos de Investigación	2020-2021	10,794.11

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CALL	PI	Full Title	Period	Amount (€)
Talento Innovador 2020	lvan Coluzz	DesFIA- Búsqueda de dianas terapéuticas guiada por inteligencia artificial	2020-2021	20,000.00
Talento Innovador 2020	Alejandro Criado	FLU-gFET - Detección de flujo multivalente de virus SARSCoV-2	2020-2021	20,000.00
 Obra Social Fundació "la Caixa	,,			
CALL	PI (Fellow	v) Full Title	Period	Amount (€)
Health Research Call 2020	Jesús Ruiz-Cabello	o ATHEROCONVERGENCE - Flow-driven inflammation and arterial wall remodeling in atherosclerosis: mechanisms and therapeutic potential	2020-2023 2020-2023	124,630.00
Doctoral Fellowship INPhINIT Incoming 2020	Aitziber L. Cortajarena (Gabriela Gue	er L. Protein-based functional materials jarena ela Guedes)		
b∔oef				
CALL	PI	Full Title	Period	Amount (€)
Ayudas 2019 a proyectos de investigación en Cáncer de Pulmón	Luis M. Liz-Marzán	INDICATE - INvitro Dlagnostics for CAncer TEsting: Desarrollo de un análisis de sangre económico, sensible y rápido para mejorar el diagnóstico y las decisiones de tratamiento en pacientes con cáncer de pulmón	2020-2023	7,920.00

Total Funding: 7,731,721€



Knowledge & Technology Transfer



In 2020 3 new **Priority Patent Applications** have been filed, 1 **Patent** was **Granted** and 17 **Collaboration Agreements** with industrial or clinical partners signed.

CIC biomaGUNE is making a substantial effort to promote the transfer to the market of research results produced at the center. In this respect, an initiative has been launched to identify and support the development of those technologies with a clear market potential, namely the development unit (DU) initiative, within the R&D&I management unit. The DU acts as an antenna and incubator of ideas, with the main objective of identifying and transferring research results from the center into the market, by protecting the results, encouraging their exploitation and

strengthening patent licensing. Projects to be matured within the DU are initially evaluated by the Scientific Direction of the center, on the basis of criteria that include the consistency of the initiative with the center's activity and the benefit that the initiative can obtain from the infrastructure/resources of the center for its maturation. In addition, the R&D&I management unit works on establishing new contracts with companies, promoting cross-sectorial research and maximizing the transfer of knowledge and technology.

Patents

The generation of patents is one of the instruments that indicate the transfer of knowledge from the Center to Society and to business development. The graph shows the evolution over the years.



2020 Patent Applications

Glycan mediated spleen targeting using extracellular vesicles

N. C. Reichardt, C. Williams, J. M. Falcón

> EP20382981.7 13.11.2020

Isoalloxazinebased catalytic systems and uses thereof.

F. López-Gallego, L. Salassa

> EP20383067.4 07.12.2020

Layered substrate and uses thereof.

L. M. Liz-Marzán, J. Plou, M. Charconnet, I. García

> EP20383131.8 21.12.2020

Pharmaceutical composition comprising fluorine-18 labelled gases J. Llop Roig, V. Gómez-Vallejo, T. Reese, A. Lecuona.

> EP17733822.5 US10849994B2





www.aspariaglycomics.com

In 2016, CIC biomaGUNE researchers and private investors joined efforts to set up **Asparia Glycomics**, a spin-off company specialized in the production and marketing of reagents, reference standards, kits and software, intended for glycan analysis in clinical diagnosis and for quality control of biopharmaceuticals.

The business initiative markets the leading technology developed by CIC biomaGUNE's Glycotechnology Laboratory, directed by Niels Reichardt, to quantify and identify glycans more accurately and faster than existing solutions on the market. Asparia Glycomics offers unique, stable, isotope labeled glycans as internal standards, custom made quantification software and reagents for glycan analysis by mass spectrometry.

Asparia Glycomics' line of work is based on the development of its own technology that can be useful for the diagnosis and prognosis of cancer, diabetes or autoimmune diseases such as rheumatoid arthritis, among others.

The products and technology developed by the Glycotechnology Laboratory, reviewed and validated by the CIC biomaGUNE Development Unit are aimed at researchers, academia and pharmaceutical companies around the world.

During 2020 and after a change of management, the company has been able to break even and has established a position as a reliable provider of cutting-edge glycan analysis and synthesis projects for leading biotech companies and academic researchers worldwide.

The company has been certified as Innovative SME by the Spanish government.

The collaboration between Asparia Glycomics and CIC biomaGUNE has continued during 2020 with several ongoing research projects and 3 co-directed PhD Theses.

Research Agreements with Companies



CIC biomaGUNE develops joint research activities in collaboration with different types of organizations and companies, including spin-offs, SMEs, large companies and research organizations.

This research activity has a sharp During 2020, besides several focus on innovation, to which CIC biomaGUNE contributes with industry and the initiation of scientific knowledge. Joint projects conversations under development include companies, 17 collaboration privately funded research activities as well as research contracts and consultancy services.

technical services provided to with many agreements with 11 industrial or clinical partners were signed.

1 / Collaboration Agreements with

Industrial or Clinical Partners

- Hospital Clínico Universitario de Valladolid
- Acuratio Europe S.L.
- AQUADAT Efficient Innovation, s.l.
- ASPARIA GLYCOMICS S.L
- IXTAL srl; Naturality Research & Development, S.L.
- Numio Tecnologías, S.L.
- Onena Medicines S.L.
- Out and Out Chemistry, Pr LLC
- Starget Pharma, Ltd.
- TiCom Sp. z o.o.
Scientific Output



In 2020, **188 research articles** have been published in international scientific journals with an **average impact factor** of **8.5** and received **8736 citations**.

Journal	Factor	N° of Articles
Chemical Society Reviews	42.85	1
Nature	42.78	1
Science	41.84	1
Nature Nanotechnology	31.54	1
Advanced Materials	27.40	2
Nature Immunology	20.48	1
Chem	19.73	1
Advanced Functional Materials	16.84	6
Advanced Science	15.84	1
Journal of the American Chemical Society	14.61	4
ACS Nano	14.59	8
Science Advances	13.12	2
Angewandte Chemie International Edition	12.96	5
ACS Catalysis	12.35	2
Materials Horizons	12.32	1
Nature Communications	12.12	2
Journal of Clinical Investigation	11.86	1
Small	11.46	3
Nano Letters	11.24	2
Biosensors and Bioelectronics	10.26	1

Highlights



White-emitting Protein-Metal Nanocluster Phosphors for Highly Performing Biohybrid Light-Emitting Diodes

Aires, A; Fernández-Luna, V; Fernández-Cestau, J; Costa, RD; Cortajarena, AL.

Nano Letters 2020, 20, 2710–2716 DOI: 10.1021/acs.nanolett.0c00324

This work presents a simple in situ synthesis and stabilization of fluorescent gold nanoclusters (AuNCs) with different sizes using engineered protein scaffolds in water. The protein-AuNC hybrids show a dual emission (450 and 700 nm) with a record photoluminescence quantum yield of 20%. These features impelled us to apply them to biohybrid light-emitting diodes as color downconverting filters or biophosphors. Efficient white emission (x/y CIE color coordinates of 0.31/0.29) and stabilities of more than 800 h were achieved. This represents a 2 orders of magnitude enhancement compared to the prior art. Besides the outstanding performance, the protein scaffold also infers a unique anisotropic emission character that is considered as a proof-of-concept of high interest for single-point lighting and display.



Micelle-directed chiral seeded growth on anisotropic gold nanocrystals

González-Rubio, G; Mosquera, J; Kumar, V; Pedrazo-Tardajos, A; Llombart, P; Solís, DM; Lobato, I; Noya, EG; Guerrero-Martínez, A; Taboada, JM; Obelleiro, F; MacDowell, LG; Bals, S; Liz-Marzán, LM.

Science, 2020, 368, 1472-1477 DOI: 10.1126/science.aba0980

This work reports a mechanism by which gold atoms are deposited by means of chemical reduction onto previously formed gold nanorods to produce a quasihelicoidal structure (the particles acquire chirality). This geometry enables these "nanoscrews" to interact with circularly polarized light much more efficiently than what is achieved with any other known object. These properties could lead to the detecting of biomolecules in a very selective and very sensitive way.

Surfactant-assisted seeded growth of metal nanoparticles (NPs) can be engineered to produce anisotropic gold nanocrystals with high chiroptical activity through the templating effect of chiral micelles formed in the presence of dissymmetric cosurfactants. Mixed micelles adsorb on gold nanorods, forming guasihelical patterns that direct seeded growth into NPs with pronounced morphological and optical handedness. Sharp chiral wrinkles lead to chiral plasmon modes with high dissymmetry factors (~0.20). Through variation of the dimensions of chiral wrinkles, the chiroptical properties can be tuned within the visible and near-infrared electromagnetic spectrum. The micelle-directed mechanism allows extension to other systems, such as the seeded growth of chiral platinum shells on gold nanorods. This approach provides a reproducible, simple, and scalable method toward the fabrication of NPs with high chiral optical activity.



Chemo-Enzymatic Synthesis of S. mansoni O-Glycans and Their Evaluation as Ligands for C-Type Lectin Receptors MGL, DC-SIGN, and DC-SIGNR

Pham, J.; Hernández, A.; Cioce, A.; Achilli, S.; Goti, g.; Vivès, C.; Thepaut, M.; Bernardi, A.; Fieschi, F. and Reichardt, NC.

Chem. Eur. J., 26, 12818-12830 DOI: 10.1002/chem.202000291

Due to their interactions with C-type lectin receptors (CLRs), glycans from the helminth Schistosoma mansoni represent promising leads for treatment of autoimmune diseases, allergies or cancer. We chemo-enzymatically synthesized nine O-glycans based on the two predominant O-glycan cores observed in the infectious stages of schistosomiasis, the mucin core 2 and the S. mansoni core. The O-glycans were fucosylated next to a selection of *N-glycans* directly on a microarray slide using a recombinant fucosyltransferase and GDP-fucose or GDP-6-azidofucose as donor. Binding assays with fluorescently labelled human CLRs DC-SIGN, DC-SIGNR and MGL revealed the novel O-glycan O8 as the best ligand for MGL from our panel. Significant binding to DC-SIGN was also found for azido-fucosylated glycans. Contrasting binding specificities were observed between the monovalent carbohydrate recognition domain (CRD) and the tetravalent extracellular domain (ECD) of DC-SIGNR.



Na⁺ controls hypoxic signalling by the mitochondrial respiratory chain

Hernansanz-Agustín, P; Choya-Foces, C; Carregal-Romero, S; Ramos, E; Oliva, T; Villa-Piña, T; Moreno, L; Izquierdo-Álvarez, A; Cabrera-García, JD; Cortés, A; Lechuga-Vieco, AV; Jadiya, P; Navarro, E; Parada, E; Palomino-Antolín, A; Tello, D; Acín-Pérez, R; Rodríguez-Aguilera, JC; Navas, P; Cogolludo, Á; López-Montero, I; Martínez-del-Pozo, Á; Egea, J; López, MG; Elrod, JW; Ruíz-Cabello, J; Bogdanova, A; Enríquez, JA; Martínez-Ruiz, A.

Nature 2020, 586, 287–291 DOI: 10.1038/s41586-020-2551-y

A multidisciplinary study published in Nature in which the Laboratory of Molecular and Functional Biomarkers of CIC biomaGUNE has participated, led by Ikerbasque Professor Jesús Ruiz-Cabello —specialist in the study of pulmonary vascular diseases—, have made a major advance toward deciphering the mechanism through which the production of reactive oxygen species (ROS) increases in the early phases of hypoxia (acute reduction in tissue oxygen). The finding represents an important advance in the understanding of cell physiology and could be exploited in the future to treat a variety of diseases in which hypoxia plays a role, such as stroke and heart attack.



I-124 Radiolabeling of a Au-III-NHC Complex for *In Vivo* Biodistribution Studies

Guarra, F; Terenzi, A; Pirker, C; Passannante, R; Baier, D; Zangrando, E; Gómez-Vallejo, V; Biver, T; Gabbiani, C; Berger, W; Llop, J; Salassa, L.

Angew. Chem. Int. Ed., 2020, 59, 17130-17136 DOI: 10.1002/anie.202008046

Au^{III} complexes with N-heterocyclic carbene (NHC) ligands have shown remarkable potential as anticancer agents, yet their fate in vivo has not been thoroughly examined and understood. Reported herein is the synthesis of new Au^{III}-NHC complexes by direct oxidation with radioactive [¹²⁴]]I₂ as a valuable strategy to monitor the *in vivo* biodistribution of this class of compounds using positron emission tomography (PET). While *in vitro* analyses provide direct evidence for the importance of Au^{III}-to-Au^{II} reduction to achieve full anticancer activity, *in vivo* studies reveal that a fraction of the Au^{III}-NHC prodrug is not immediately reduced after administration but able to reach the major organs before metabolic activation.

Functional rewiring across spinal injuries via biomimetic nanofiber scaffolds

Usmani S; Franceschi Biagioni A; Medelin M; Scaini D; Casani R; Aurand E; Padro D; Egimendia A; Ramos-Cabrer P; Scarselli M, De Crescenzi M, Prato M; Ballerini L.

PNAS 2020 117, 25212-25218 DOI: 10.1073/pnas.2005708117

The re-growth of severed axons is fundamental to reestablish motor control after spinal-cord injury (SCI). Ongoing efforts to promote axonal regeneration after SCI have involved multiple strategies only partially successful. Our study introduces an artificial carbonnanotube based scaffold that, once implanted in SCI rats, improves motor function recovery. Confocal microscopy analysis plus fiber tracking by magnetic resonance imaging and neurotracer labelling of longdistance cortico-spinal axons, suggest that recovery might be partly attributable to successful crossing of the lesion site by regenerating fibers. Since manipulating SCI microenvironment properties, such as mechanical and electrical ones, may promote biological responses, we propose this artificial scaffold as a prototype to exploit the physics governing spinal regenerative plasticity.





Synthesis and Characterization of Antibody-Protected Bimetallic Nanoclusters with Catalytic Properties

Mora-Sanz, V; Saa, L; Briz, N; Pavlov, V.

Chem. Mater. 2020, 32, 8286–8293, DOI: 10.1021/acs.chemmater.0c02096

In this work, a novel and facile method for the synthesis of gold-platinum bimetallic nanoclusters (NCs) embedded in the structure of an IgG (Au/Pt NCs-IgG) is presented. Proteins have been widely used as scaffolds for the synthesis of atomic clusters. The harsh conditions usually required during the synthesis imply the denaturation of different proteins and the loss of their biological activity. We propose a strategy for the synthesis of NCs employing IgG as a scaffold, performed under physiological conditions in order to keep the IgG structure unaltered, allowing the resulting material to bind to Protein G and the antigen. NCs composed of two different types of atoms exhibit higher catalytic activity than monometallic NCs due to the synergistic effect of two diverse atoms. This peroxidase-like activity and the maintained affinity for its antigen make Au/Pt NCs-IgG a suitable material for its use as a detection antibody in a direct sandwich enzyme-linked immunosorbent assay (ELISA). The use of Au/Pt NCs-IgG as an alternative to IgG tethered to horseradish peroxidase in ELISA boosts the limit of detection (LOD) by 56 times.

Selective oxidation of alkyl and aryl glyceryl monoethers catalysed by an engineered and immobilised glycerol dehydrogenasechain

Velasco-Lozano, S; Roca, M; Leal, A; Mayoral, JA; Pires, E; Moliner, V; López-Gallego, F.

Chem. Sci., 2020, 11, 12009-12020 DOI: 10.1039/D0SC04471G

Enzymes acting over glyceryl ethers are scarce in living cells, and consequently biocatalytic transformations of these molecules are rare despite their interest for industrial chemistry. In this work, we have engineered and immobilised a glycerol dehydrogenase from Bacillus stearothermophilus (BsGlyDH) to accept a battery of alkyl/aryl glyceryl monoethers and catalyse their enantioselective oxidation to yield the corresponding 3-alkoxy/aryloxy-1-hydroxyacetones. QM/MM computational studies decipher the key role of D123 in the oxidation catalytic mechanism, and reveal that this enzyme is highly enantioselective towards S-isomers (ee > 99%). Through structureguided site-selective mutagenesis, we find that the mutation L252A sculpts the active site to accommodate a productive configuration of 3monoalkyl glycerols. This mutation enhances the kcat 163-fold towards 3-ethoxypropan-1,2-diol, resulting in a specific activity similar to the one found for the wild-type towards glycerol. Furthermore, we immobilised the L252A variant to intensify the process, demonstrating the reusability and increasing the operational stability of the resulting heterogeneous biocatalyst. Finally, we manage to integrate this immobilised enzyme into a one-pot chemoenzymatic process to convert glycidol and ethanol into 3-ethoxy-1-hydroxyacetone and (R)-3-ethoxypropan-1,2-diol, without affecting the oxidation activity. These results thus expand the uses of engineered glycerol dehydrogenases in applied biocatalysis for the kinetic resolution of glycerol ethers and the manufacturing of substituted hydroxyacetones.

Edited Books and Book Chapters



Colloidal Synthesis of Plasmonic Nanometals

Liz-Marzán, L.M. 2020, Jenny Stanford Publishing; 1st edition ISBN: 9789814800679



Colloids for Nanobiotechnology: Synthesis, Characterization and Potential Applications

Feliu, N.; Parak, W. J. Frontiers of Nanoscience, Vol 16, 2020, Elsevier. ISBN: 978-0081028285



Trends in Quorum Sensing and Quorum Quenching: New Perspectives and Applications;

"Analysis of Quorum Sensing by Surface-Enhanced Raman Scattering Spectroscopy"

Bodelón, G.; De Marchi-Lourenco,S.; Pérez-Juste, J.; Pastoriza-Santos, I.; Liz-Marzán, L.M.

CRC Press, Ch. 5, 2020, pp. 59-77 ISBN: 9780367224288



Immobilization of Enzymes and Cells

Guisan, JM.; Bolivar, JM; López-Gallego, F; Rocha-Martin, J. Springer, 2020, Vol. 2100 ISBN: 978-1-0716-0215-7



Encapsulation of Enzymes in Protein Films

Sánchez-de-Alcázar, Liutkus M.; Cortajarena, A. L.

Immobilization of Enzymes and Cells, Springer, 2020, Vol. 2100, pp. 211-226

Encapsulation of Enzymes in Porous Capsules via Particle Templating

Zyzin, MV; Ramos-Cabrer, P; Carregal-Romero, S.

Immobilization of Enzymes and Cells, Springer, 2020, Vol. 2100, pp. 227-241

ISBN: 978-1-0716-0215-7



MRI to CTA Translation for Pulmonary Artery Evaluation Using CycleGANs Trained with Unpaired Data

Stephens, M.; San José Estépar, R.; Ruiz-Cabello, J.; Arganda-Carreras, I.; Macía, I.; López-Linares, K.

Thoracic Image Analysis; Springer, 2020; pp 118–129. Part of the *Lecture Notes in Computer Science* book series (LNCS, volume 12502) Also part of the *Image Processing*, *Computer Vision*, *Pattern Recognition*, *and Graphics* book sub series (LNIP, volume 12502)

ISBN: 0302-9743

Cover Pages



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Thrombo-tag, an *in vivo* formed nanotracer for the detection of thrombi in mice by fast pre-targeted molecular imaging

Adrover, JM; Pellico, J; Fernández-Barahona, I; Martín-Salamanca, S; Ruiz-Cabello, J; Hidalgo, A; Herranz, F.

Nanoscale 2020, 12, 2297



C ROWAL SOCIETY rsc.ll/chemical-science

Selective oxidation of alkyl and aryl glyceryl monoethers catalysed by an engineered and immobilised glycerol dehydrogenase

Velasco-Lozano, S; Roca, M; Leal, A; Mayoral, JA; Pires, E; Moliner, V; López-Gallego, F. Chem. Sci. 2020. 11. 12009



Discovering Biomolecules with Huisgenase Activity: Designed Repeat Proteins as Biocatalysts for (3+2) Cycloadditions

Rivilla, I; Odriozola-Gimeno, M; Aires, A; Gimeno, A; Jiménez-Barbero, J; Torrent-Sucarrat, M; Cortajarena, AL; Cossío, FP.

J. Am. Chem. Soc. 2020, 142, 762-776



Biogenic fluorescent protein-silk fibroin phosphors for high performing lightemitting diodes

Fernandez-Luna, V; Fernandez-Blazquez, JP; Monclus, MA; Rojo, FJ; Daza, R; Sánchez-de-Alcázar, D; Cortajarena, AL; Costa, RD

Mater. Horiz. 2020, 7, 1790-1800



Fluorinated carbohydrates as chemical probes for molecular recognition studies. Current status and perspectives

Linclau, B; Arda, A; Reichardt, NC; Sollogoub, M; Unione, L; Vincent, SP; Jiménez-Barbero, J. Chem. Soc. Rev. 2020, 49,

3863-3888



Templated-Assembly of CsPbBr(3)Perovskite Nanocrystals into 2D Photonic Supercrystals with Amplified Spontaneous Emission

Polavarapu, L; Vila-Liarte, D; Feil, MW; Manzi, A; García-Pomar, JL; Huang, H; Döblinger, M; Liz-Marzán, LM; Feldmann, J; Mihi, A

Angew. Chem. Int. Ed. 2020, 59, 17750-17756



ALS PUDICATIONS Mathumac Not Chat Not Net.

Dual Monitoring of Surface Reactions in Real Time by Combined Surface-Plasmon Resonance and Field-Effect Transistor Interrogation

Aspermair, P; Ramach, U; Reiner-Rozman, C; Fossati, S; Lechner, B; Moya, S; Azzaroni, O; Dostalek, J; Szunerits, S; Knoll, W; Bintinger, J.

J. Am. Chem. Soc. 2020, 142, 11709-1171



Colloidal Superstructures with Triangular Cores: Size Effects on SERS Efficiency

Höller, RPM; Kuttner, C; Mayer, M; Wang, R; Dulle, M; Contreras-Cáceres, R; Fery, A; Liz-Marzán, LM.

ACS Photonics 2020, 7, 1839-1848



In Silico Evidence That Protein Unfolding is a Precursor of Protein Aggregation

Bianco, V; Franzese, G; Coluzza, I. ChemPhysChem 2020, 21, 377-384



CONTRACTORY CONTRA

A study of the complex interaction between poly allylamine hydrochloride and negatively charged poly(N-isopropylacrylamideco-methacrylic acid) microgels

Giussi, JM; Martínez-Moro, M; Iborra, A; Cortez, ML; Di Silvio, D; Llarena, I; Longo, GS; Azzaroni, O; Moya, S.

Soft Matter 2020, 16, 881-890



ACS Publications

Tailored Nanoscale Plasmon-Enhanced Vibrational Electron Spectroscopy

Tizei, LH; Mkhitaryan, V; Lourenço-Martins, H; Scarabelli, L; Watanabe, K; Taniguchi, T; Tencé, M; Blazit, J; Li, X; Gloter, A; Zobelli, A; Schmidt, F; Liz-Marzán, LM; García de Abajo, FJ; Stephan, O; Kociak, M.

Nano Lett. 2020, 20, 2973–2979





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Surfactant-Assisted Symmetry Breaking in Colloidal Gold Nanocrystal Growth

González-Rubio, G; Scarabelli, L; Guerrero-Martínez, A; Liz-Marzán, LM

ChemNanoMat 2020, 6, 698-707



Protein-directed crystalline 2D fullerene assemblies

Liutkus, M; López-Andarias, A; Mejías, SH; López-Andarias, J; Gil-Carton, D; Feixas, F; Osuna, S; Matsuda, W; Sakurai, T; Seki, S; Atienza, C; Martín, N; Cortajarena, AL.

Nanoscale 2020, 12, 3614-3622



Water Soluble Cationic Poly(3,4-

Ethylenedioxythiophene) PEDOT-N as a Versatile Conducting Polymer for Bioelectronics

Minudri, D; Mantione, D; Domínguez-Alfaro, A; Moya, S; Maza, E; Bellacanzone, C; Antognazza, MR; Mecerreyes, D.

Adv. Electron. Mater. 2020, 6-10, 2000510



Growth of ZIF-8 MOF Films with Tunable Porosity by using Poly (1vinylimidazole) Brushes as 3D Primers

Allegretto, JA; Iborra, A; Giussi, JM; von Biderling, C; Ceolín, M; Moya, S; Azzaroni, O; Rafti, M.

Chem. Eur. J. 2020, 26, 12388-12396

Editorial Activity

ADVANCING SCIENCE, SERVING SOCI

Sience Reviewing Editor: *Luis M. Liz-Marzán*



ACS Nano Associate Editors: *Wolfgang Parak, Luis M. Liz-Marzán* Editorial Advisory Board: *Maurizio Prato*

ACS Applied Bio Materials Associate Editor: *Aitziber L. Cortajarena*

Accounts of Chemical Research Editorial Advisory Board: *Luis M. Liz-Marzán*

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Bioconjugate Chemistry Editorial Advisory Board: *Wolfgang Parak*



Faraday Discussions Advisory Board: *Luis M. Liz-Marzán*

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Materials Advances Advisory Board: *Luis M. Liz-Marzán*



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Nanotoxicology Editorial Board: Wolfgang Parak

MDPI

International Journal of Molecular Sciences Special Issue Editor: *Ander Abarrategi*

SPRINGER NATURE

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Colloid and Interface Science Communications Advisory Board: *Wolfgang Parak*

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Full Publications List

Kang, NX; Wang, Q; Ojeda, R; Wang, WJ; Fu, FY; Moro, MM; Ramirez, MD; Moya, S; Coy, E; Salmon, L; Pozzo, JL; Astruc, D. Visible-Light Acceleration of H₂ Evolution from Aqueous Solutions of Inorganic Hydrides Catalyzed by Gold-Transition-Metal Nanoalloys. ACS Appl. Mater. Interfaces 2020, 12, 53816-53826

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Training



The SARS-COV-2 pandemic hindered some of the training activity. In particular, we could hardly host visiting students and researchers, our seminar program was completely stopped for months and then restarted in an online modality.

Yet, in 2020 there have been 66 active **PhD tesis**, 11 of them successfully completed during the year. Our researchers performed 16 secondments at collaborating institutions, and we hosted 15 stays of visiting researchers. The Center organized 18 seminars, 5 workshops/conferences and our researchers have participated as co-organizers of 6 international conferences.

PhD Program

The objective of CIC biomaGUNE's PhD Program is to a provide PhD students with top quality multidisciplinary training at the interface between biology, chemistry, nanobiotechnology and materials science. The students benefit from international training and are exposed to different research areas. The training program offers weekly lectures by leading scientists, technical training courses on a broad range of scientific techniques and instrumentation, opportunities for

short stays at renowned international research institutions, as well as complementary training in soft skills to prepare doctoral candidates to become highly qualified, autonomous and skilled professionals. The joint training of PhD candidates fosters closer ties and cooperation between research groups and researchers of the institutions involved.

CIC biomaGUNE has agreements with several universities (which are the degree-grating bodies) to enroll PhD students in different PhD programs. Additionally, CIC biomaGUNE principal researchers are PhD Program Professors in the following Doctoral programs from the University of the Basque Country (UPV/EHU), CIC biomaGUNE participates in the following Doctoral programs.

- Synthetic & Industrial Chemistry
- Applied Chemistry & Polymeric Materials
- Molecular Biology & Biomedicine
- Medicine & Surgery
- Biomedical Research programs

Co-supervised PhD Scheme



Since 2016, CIC biomaGUNE is running a program of co-supervised (and co-funded) PhD theses, in collaboration with other research institutions, technology centers and enterprises in the Basque Country. In 2020, 25 PhD theses were co-directed with the following institutions:

CIC biomaGUNE has agreements with industrial partners, to conduct industrially-oriented PhD research projects.

- POLYMAT Basque Center for Macromolecular Design & Engineering
- DIPC
- CIC bioGUNE
- CIC nanoGUNE
- IIS BioCruces
- IIS BIODONOSTIA

- TECNALIA
- GAIKER
- CIDETEC
- ACHUCARRO Basque Center for Neuroscience
- UPV-EHU
- BIOFISIKA Basque Center for Biophysics



PhD Theses

During 2020, 11 students obtained their **PhD** at CIC biomaGUNE



Daniel Sánchez de Alcazar

Engineered proteins as molecular templates for functional materials

Supervisor: Aitziber L. Cortajarena Defense Date: 10/01/2020

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Rafaela Paz García Álvarez Protein corona and optoacustic properties of gold nanoparticles of different size and shape Supervisor: Prof. Luis M. Liz-

Marzán & Prof. Kostas Kostarelos

Defense Date: 14/02/2020



Beatriz Díez Buitrago Enzymatic modulation of nanomaterials and its application to biosensing Supervisor: Dr. Valery Pavlov Defense Date: 07/04/2020



Pablo Valverde Sánchez NMR Insights into DC-SIGN recognition of fucosylated antigens

Supervisor: Dr. Niels C. Reichardt & Prof. Jesús Jiménez Barbero

Defense Date: 27/04/2020



llaria Calloni NMR studies of the interactions of glycan antigens with monoclonal antibodies

Supervisor: Dr. Niels C. Reichardt & Prof. Jesús Jiménez Barbero Defense Date: 29/06/2020



Donostia-San Sebastián-2020

Krishna Reddy Pulagam

Synthesis, radiolabeling and preclinical evaluation of gold-nanomaterial based boron carriers as drug delivery systems: Potential application to boron neutron capture therapy Supervisor: *Dr. Jordi Llop* Defense Date: 01/07/2020



Irene V. J. Feiner

A pre-targeting approach to boron neutron capture therapy: towards multipurpose boronenriched therapeutic agents

Supervisor: *Dr. Jordi Llop* Defense Date: 06/07/2020





Juan Pedro Merino-Rusillo Modifications of Graphene Prepared by Chemical Vapor Deposition for Diagnostic Applications

Supervisor: Prof. Maurizio Prato & Dr. Alejandro Criado Defense Date: 02/09/2020



Charles Williams

Glycoengineered Exosomes as Vehicles for Gene and Drug Delivery

Supervisor: Dr. Niels C. Reichardt & Dr. Juan Falcón Defense Date: 07/10/2020



Marcos Navascuez Lominchar Nanoemulsions as drug carriers: preclinical evaluation of pulmonary delivered drug-loaded nanocapsules

Supervisor: Dr. Jordi Llop & Dr. Iraida Loinaz Defense Date: 30/10/2020



Marta Martínez Moro Biological Fate of Soft Nanomaterials: A Physico-Chemical and Translocation Study Supervisor: *Dr. Sergio Moya* Defense Date: 21/12/2020

Master & Vocational Training Programs

Master Students

CIC biomaGUNE has agreements with several universities (which are the degreegrating bodies) in different Master programs. Additionally, in partnership with the **University of the Basque Country (UPV/EHU**), CIC biomaGUNE participates in the following MSc courses by providing lectures and direction of master thesis:

- Molecular Biology and Biomedicine
- Nanoscience
- Chemistry and Polymers

Vocational Training Programs

CIC biomaGUNE has agreements with **CPES CESA BHIP, Don Bosco and Colegio Inmakulada Ikastetxea,** Centers for Intermediate and Superior level vocational training in the fields of Chemistry or Biosciences to host training internships of students. Several training placements take place every year at different laboratories of the Center. Selected students are mentored and supervised by postdoctoral researchers or PhD students and receive hands-on training.

Research Secondments & Summer Internships

In the framework of different national and international projects, CIC biomaGUNE has established collaborations with international institutions to **second** and **host research stays of PhD students and postdoctoral researchers**. During 2020, our researchers performed 16 secondments at collaborating partners and we hosted 15 stays of visiting researchers.

This activity was clearly hindered by the pandemic situation and the summer undergraduate program could not take place in 2020.

Research stays	2015	2016	2017	2018	2019	2020
From CIC biomaGUNE	30	30	19	11	23	16
To CIC biomaGUNE	59	105	92	84	98	15
Visiting Professors/Sabbaticals	1	4	1	2	2	
Erasmus placements	1	7	7	2	8	
Summer placements	2	5	6	6	15	
Vocational training	4	7	5	3	5	

Seminars

CIC biomaGUNE runs an annual program of scientific seminars, which includes:

i) International seminars delivered by internationally recognized researchers of varying scientific backgrounds and fields.

ii) **Postdoc seminars** delivered by selected postdoctoral fellows to share progress in each of our laboratories to the rest of the Center.

iii) PhD seminars delivered by PhD students of the Center.

iv) **Training seminars** delivered by CIC biomaGUNE's PIs or Platform Managers, aimed at strengthening the technical training program and soft skills training. Altogether, these programs contribute to the career development of our researchers. During 2020, 16 seminars have been delivered most of them in an online modality after a long break caused by the pandemic situation.



09/01/2020 14/10/2020 Neutralizing Dual-SMAD Inhibition Proteins as a Conjugated polymers for neural stimulation and their Therapeutic Mechanism of Action application for vision restoration in retinal dystrophies Dr. Neethan A. Lobo - Onena Medicines Prof. Fabio Benfenati - University of Genoa 30/01/2020 21/10/2020 Aberrant activation of developmental programs in How to keep women (and men) in science adult disease Prof. Dr. Petra Rudolf - Zernike Institute for Prof. Ángela Nieto - Instituto de Neurociencias Advanced Materials, University of Groningen (CSIC-UMH) 29/10/2020 06/02/2020 CIC biomaGUNE 2nd PhD Day Manufacturing of Recombinant Adeno-associated **CIC biomaGUNE** Viral Vector (rAAV) For Human Gene Therapy Treatments 04/11/2020 Dr. César Trigueros - Viralgen Nanotoxicology: does it matter how cells die? Prof. Bengt Fadeel, M.D., Ph.D. - Institute of 27/02/2020 Environmental Medicine, Karolinska Institutet Sperm-based microbots and their potential to improve ART success and reproductive health 11/11/2020 Dr. Mariana Medina - Institute for Integrative Inorganic-organic hybrid materials for biomedicine Nanosciences Dr. Dorleta Jiménez de Aberasturi -**CIC biomaGUNE** 04/03/2020 Looking for targets to prevent non-alcoholic fatty liver 18/11/2020 disease progression Heterochiral Peptide Assembly: Entry to Prof. Patricia Aspichueta - UPV/EHU Wonderland through the Mirror Dr. Silvia Marchesan - University of Trieste 23/09/2020 A pre-targeting approach to boron neutron 02/12/2020 capture therapy: towards multipurpose boron-Glycoengineering of Extracellular Vesicles enriched therapeutic agents Dr. Charles Williams - CIC biomaGUNE Dr. Irene Feiner - CIC biomaGUNE 09/12/2020 30/09/2020 Functional nanomaterials from diatoms microalgae Modifications of Graphene Prepared by Chemical Prof. Gianluca M. Farinola - Università degli Studi di Vapor Deposition for Diagnostic Applications Bari Aldo Moro Dr. Juan Pedro Merino - CIC biomaGUNE 17/12/2020 Our Glycans: Tales of non-self, self and superself

Dr. Niels C. Reichardt - CIC biomaGUNE

CIC biomaGUNE seeks to enhance its national and international visibility and reputation by organizing conferences, workshops and seminars that share knowledge about the latest research and advances in the field of biomaterials. The following events have been organized during 2020: Additionally, our researchers chaired or co-organized a number of international symposia:



Outreach

One of the strategic objectives of CIC biomaGUNE is the communication and dissemination of knowledge within and beyond the academic community. In 2020, despite the pandemic situation, our commitment toward outreach, promotion of STEAM careers in young girls and visibility of women in science has continued, as clearly reflected in the list of conducted activities:

Emakumeak Zientzian

International Day of Women & Girls in Science

The Basque Technology Park Network joins the campaign "Women and Girls in science" on the occasion of the celebration on February 11 of the "International Day of Women and Girls in science". This campaign aims to make visible the female talent working in companies and centers of the Technology Parks of Araba, Bizkaia and Gipuzkoa. For this campaign, they have counted with the participation of:

Susana Carregal-Romero, Research Associate, researches on the pulmonary administration of nanomedicines and the effect of their physicochemical properties on their fate and efficiency.

Dina Niculaes, Postdoctoral Researcher in the Biomolecular Nanotechnology group, investigating the feasibility of a test for antibody detection based on fluorescent metallic nanoclusters stabilized by designer proteins.

Clara García-Astrain, Postdoctoral Researcher in the BioNanoPlasmonics group, investigates the development of new 3D nanocomposite materials for tumor growth monitoring.

Irantzu Llarena, Head of the Optical Microscopy Platform, which involves teaching, technical work and research in the characterization of biomaterial and biological systems.

The image of yesterday, today, tomorrow

Jordi Llop

Opinion piece for El Correo.

Inspira 2020

The INSPIRA project is a pioneering project in Euskadi for the promotion of the scientific-technological vocation (STEAM: Science, Technology, Engineering, Arts and Maths) among girls.

The mentors accompany students during a period of two months with the following aim:

- Provide new references of nearby women technologists
- For boys and girls to discover the STEAM professions
- Raise awareness of the need for the development of the country to occur between men and women
- Sensitize and guide on the career in technology
- Make visible and value women technologists

The mentors from CIC biomaGUNE for 2020 have been *Elisa Bindini, Vanessa Gómez-Vallejo, Clara García-Astrain* and *Angie Ramírez.*

Instagram Live Talk between *Aitziber L. Cortajarena* and *Marián García (Boticaría García)*, who has a PhD in Pharmacy and has a popular outreach handle on Instagram where she talks about different topics.

They discussed about the response of the scientific community to the Covid-19.

Encounter of scientific lives 2020

CIC biomaGUNE will participate in the 10th "Encuentro de Vidas next Científicas 2020" organized by Eureka! Zientzia Museoa, located in the Parke of Gipuzkoa. This activity, took place on October 19 and November 2, and it is directed to 4° ESO and bachelorship students with the aim of boosting scientific vocations. The aim is to create a space where students can speak, ask questions and exchange opinions directly with various professionals from different scientific and technological fields. Thus, students who have in mind to study science- related studies will be able to find out about future employment opportunities, ask questions about what it means to take certain studies, detect possible difficulties to be encountered, etc.

Pedro Ramos-**Cabrer interview** in La Mecánica del Caracol

Pedro Ramos-Cabrer, Principal Investigator of the MRI group at CIC biomaGUNE, interviewed by Eva Caballero in "La Mecánica del Caracol" to talk about his work on reconnecting neurons from spinal cord injuries in vivo.

The results of this study by Ikerbasque Professor Pedro Ramos-Cabrer and Ikerbasque Professor and Axa Chair Maurizio Prato, both researchers at CIC biomaGUNE, have been published in the prestigious journal PNAS (Proceedings of the National Academy of Sciences of the U.S.A.) and open a way for a potential recovery of spinal cord injuries.

Rosalind Franklin, honoured at the Donostia WeekINN

As part of the WeekINN week organised by FOMENTO DSS, and within the framework of the event "Women scientists of yesterday and today", Clara García-Astrain, postdoctoral researcher in the BioNanoPlasmonics group, delivered a presentation about the life and the important role that Rosalind Franklin played in understanding the structure of DNA.

Jordi Llop, Head of the Radiochemistry and Nuclear Imaging group at CICbiomaGUNE, explains how noninvasive nuclear imaging techniques help in the diagnosis of diseases in the podcast "Hablando con Científicos".

con

II Aseica Joven **Video Competition**

Peio Azcoaga, pre-doctoral researcher in the Molecular and Functional Biomarkers group at CIC biomaGUNE and the breast cancer group at IIS Biodonostia, talks about oncostatin and breast cancer in his video for the "II Aseica Joven Video Competition".

Aitziber L. Cortajarena, interview on La Mecánica del Caracol on Radio Euskadi

Journalist Eva Caballero interviews CIC biomaGUNE Ikerbasque professor Aitziber L. Cortajarena about the e-Prot project that she coordinates and which has received a grant from the European Commission in the latest FET Open call. The researcher talks about the project, which aims to develop bioinspired, sustainable and efficient bioelectronic systems based on proteins as an alternative to the traditional technologies used in the electronics industry.

Niels C. Reichardt, interwiew in Cadena Ser

CIC biomaGUNE researcher Niels C. *Reichardt*, leader of the Glycotechnology Laboratory, interviewed by Aloña Velasco on Cadena SER's A vivir Euskadi programme. They talked about the research on COVID-19; Specifically, the project coordinated by Niels C. Reichardt, in which Neiker, Biodonostia, Asparia Glycomics and Vicomtech also participate, which aims to develop antibody and antigen microarrays as a platform for the diagnosis and epidemiological study of the virus, and which has received a grant from the Provincial Council of Gipuzkoa.

Open Days & Visits

CIC biomaGUNE regularly receives visits from high-school and university students who come to have a closer look at our activity. These visits typically include a lecture about biomaterials in the context of life sciences, an open discussion with CIC biomaGUNE researchers, and a guided facilities/laboratories tour to six technical platform, Confocal (Nanoparticle synthesis Microscopy, Radiochemistry platform, Molecular Imaging Facility, Scanning Electron Microscopy, Atomic Force Microscopy). The program of visits is run by Ana Sánchez-Iglesias, Marco Möller, Daniel Padró, Irantzu Llarena and Vanessa Gómez-Vallejo with the support from other Platform Managers as well as PhDs, Postdocs and Principal Investigators. Unfortunately, this activity was severly hindered by the pandemic situation.

Guided Visits at the Molecular Imaging Facility

Media

CIC biomaGUNE is committed to disseminating research results and other news, not only to the scientific community but also to the general public. Our presence at social media has remained with an increase of the number of followers/connections.

	2019	2020	1
Impacts in General Media	207	478	+271
Web Visits	36,715	60,447	+23,732
Twitter Followers	1827	2047	+220
Linkedin	1755	2643	+888
Facebook Likes	151	184	+33
Facebook Followers	165	208	+43

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