



Dirección General  
de Investigación  
e Innovación Tecnológica  
CONSEJERÍA DE EDUCACIÓN,  
CIENCIA Y UNIVERSIDADES

UNIÓN EUROPEA  
Fondos Estructurales  
*Invertimos en su futuro*



# PROGRAMAS DE I+D EN TECNOLOGÍAS 2018

ACRONIMO: **Tec4Bio** REF: **S2018/NMT-4443**

TITULO PROGRAMA: **Nuevas Tecnologías Aplicadas al Estudio de  
Nanomáquinas Biológicas**

PRESUPUESTO CONCEDIDO: **809.600€**



<https://tec4bio.es/>

Madrid, 17 y 18 de abril de 2024



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# ¿Quiénes participamos?



Fernando **MORENO-HERRERO**. Centro nacional de Biotecnología, CSIC (COORD)  
*Molecular Biophysics. DNA-protein interactions.*



Ricardo **GARCÍA**. Instituto de Ciencias de Materiales de Madrid, CSIC  
*Atomic Force Microscopy. Cellular mechano-biology.*



Gustavo **PLAZA**. Universidad Politécnica de Madrid  
*Engineering methods in micro-rheology.*



Miguel Angel **DEL POZO**. Spanish National Center for Cardiovascular Research. CNIC  
*Mechano-transduction and mechano-adaptation at the cellular level.*



Oscar **LLORCA**. Spanish National Center for Cancer Research. CNIO  
*Structural biology. Cryo-EM methods.*



Jorge **ALEGRE-CEBOLLADA**. Spanish National Center for Cardiovascular Research. CNIC  
*Molecular mechano-biology of proteins.*

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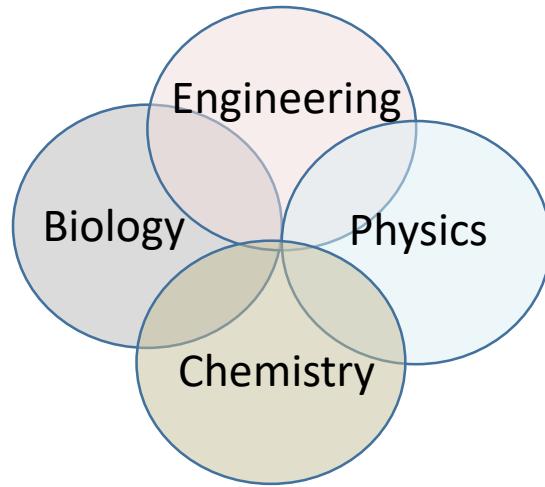
UNIVERSIDAD  
POLITÉCNICA  
DE MADRID





# ¿Qué objetivos planteamos?

**Tec4Bio** consolidates a team of engineers, physicists, and biologists to develop and apply **new biophysical technologies** to answer key questions in **cardiovascular disease** and **organization and repair of DNA**



How physical forces and mechanical properties of cells and macromolecular complexes contribute to cell development, physiology and disease

A. **Development** of instruments and new methodologies in molecular and cellular mechanobiology

B. **Application** of new technologies to the study of biological nanomachines at the multiscale level





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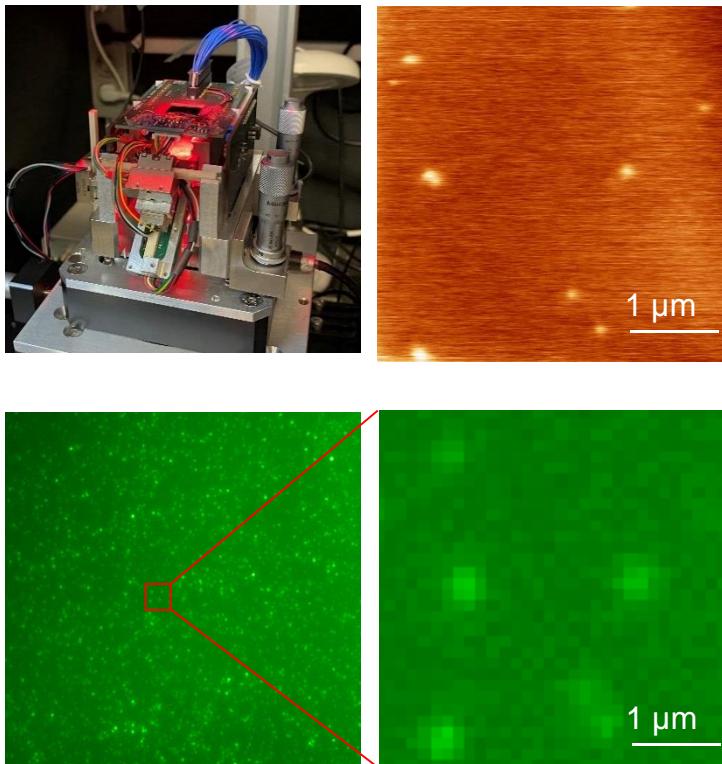


# ¿Qué resultados hemos obtenido?

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## A. Development of instruments and new methodologies in molecular and cellular mechanobiology

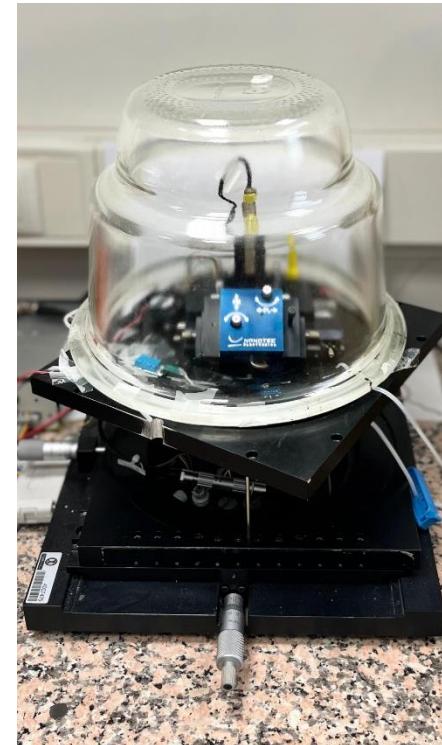
### A combined AFM-TIRF setup



...and other SM methodologies.

Labs involved: Moreno-Herrero, Plaza, García, Alegre-Cebollada

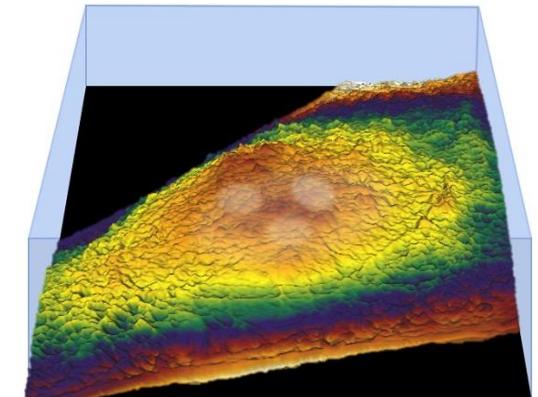
### A micro-incubator for cell mechanical studies



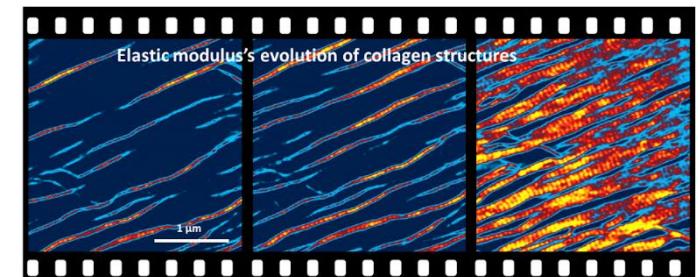
González-Bermudez et al Inmunología 2022

### High-resolution AFM force spectroscopy

Cells



Collagen structures



Gisbert et al ACS Nano 2021

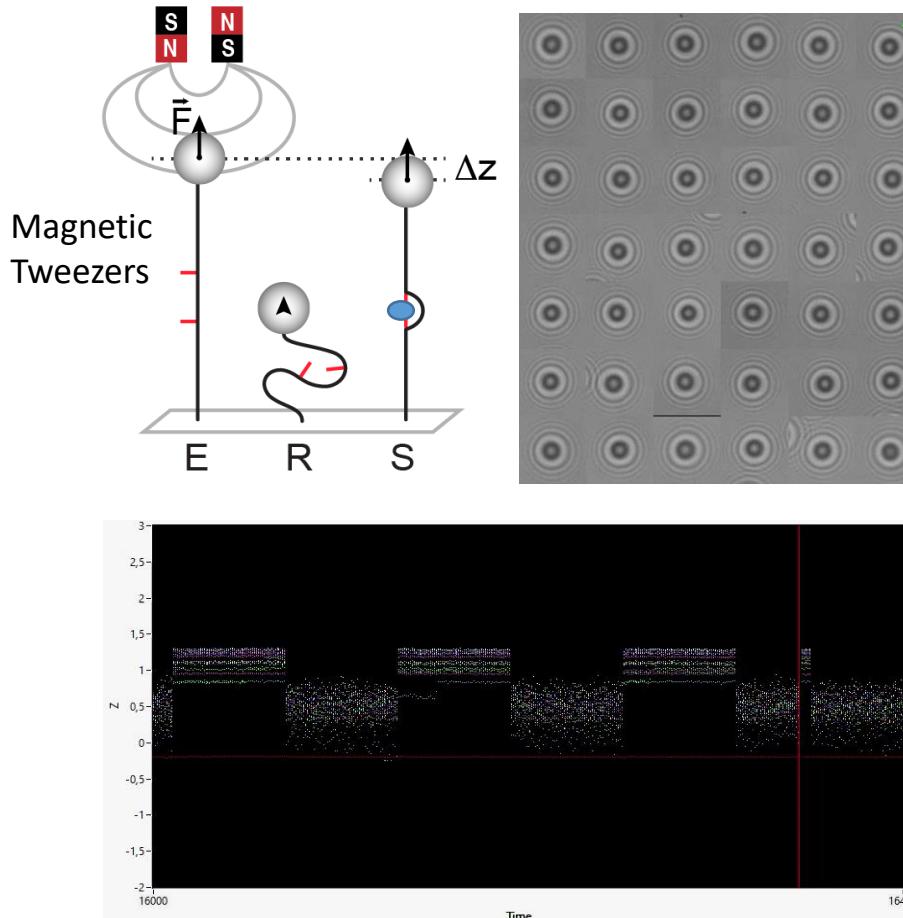
Gisbert et al Small 2023



## B. Application of new technologies to the study of biological nanomachines at the multiscale level

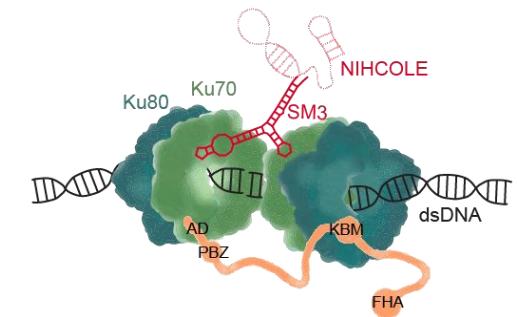
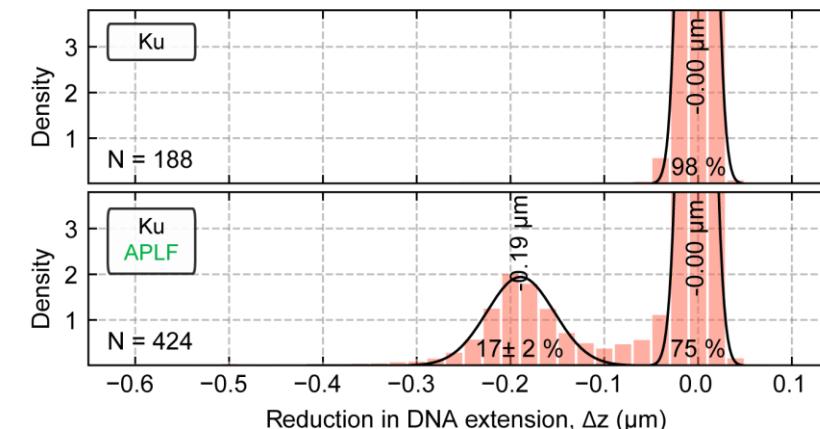
### Biophysical studies of NHEJ complexes

Mimicking a DSB to study the DNA repair pathways



### APLF, Ku and NIHCOLE in NHEJ

- In NHEJ, the Ku-APLF complex supports DNA end synapsis for several minutes under piconewton forces
- lncRNA NIHCOLE fortifies the DNA synapsis



Unfried et al. *Cancer Research* 2021

Coloma et al *Nucleic Acids. Res.* 2023

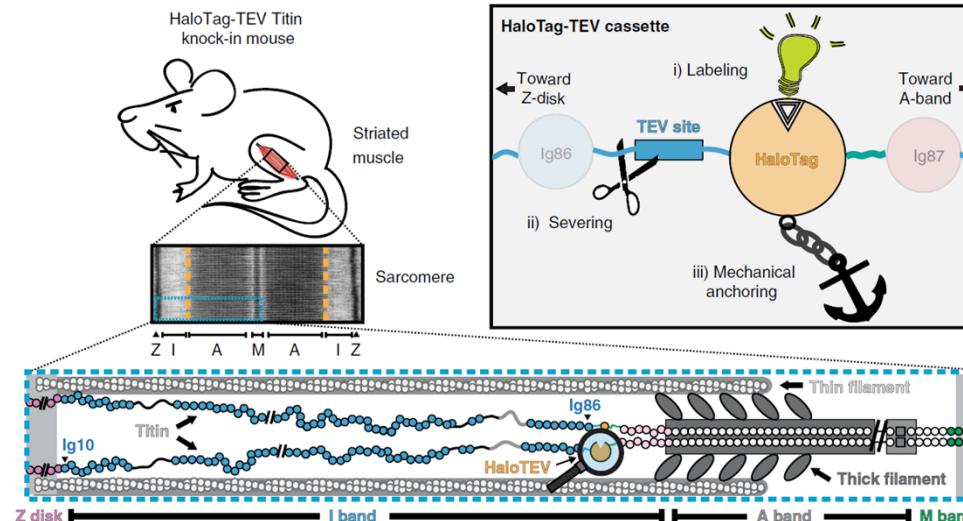
De Bragança et al *Cell reports* 2023

Labs involved: Moreno-Herrero and Llorca



## B. Application of new technologies to the study of biological nanomachines at the multiscale level

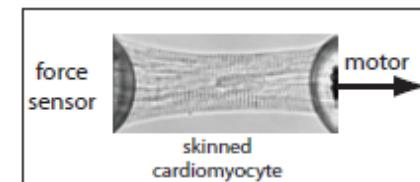
### Mechanical studies of cells and proteins



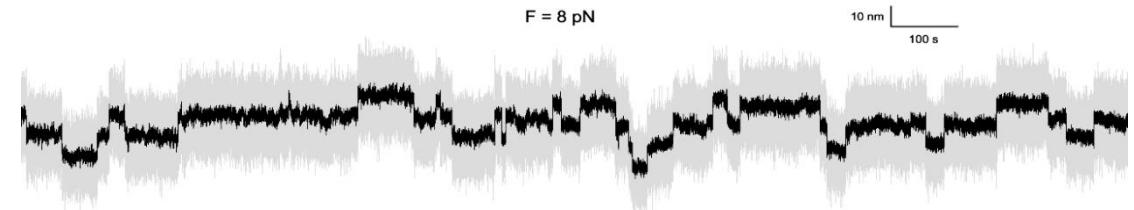
Micropipette cell aspiration:



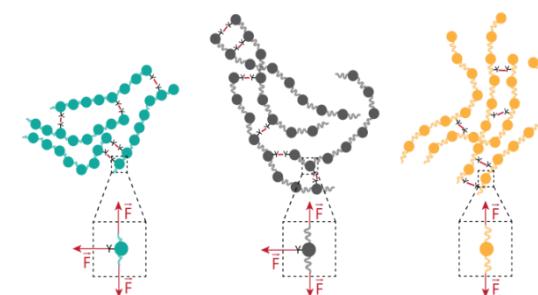
AURORA Cell stretcher:



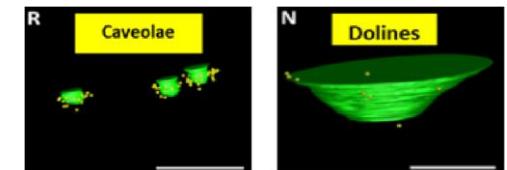
### Magnetic Tweezers:



### Protein hydrogels:



### Modelado 3D de estructuras caveolares:



Rivas-Pardo et al. *Nature Comm* 2020

Sen-Martin et al. *bioRxiv* 2024

Huerta-López et al. *Sci. Adv.* (under review)

Garcia-Garcia et al *Nature Comm*. 2022

Lolo et al *Nature Cell Biol* 2023



## Tec4Bio in numbers

- More than **63 publications** from the 6 groups involved in the consortium.
- >94% of them in open Access.

Famelis et al Nature 2019    Wilkinson et al Nucleic Acids Res. 2020    Zimmermann et al Science Adv. 2020    Perez-Rigueiro et al Molecules 2021  
Suay-Corredora et al ACS Nano 2021    Garcia-Garcia et al Nature Comm. 2022    Rivas-Pardo et al Nature Comm. 2022    Hormeño et al PNAS 2022  
Unfried et al. Cancer Research 2021    Lolo et al Nature Cell Biol 2023    De Bragança et al Cell reports 2023    Coloma et al Nucleic Acids. Res. 2023

- More than **360 presentations in conferences, meetings and invited talks**.
- **17 Master Thesis** and **26 PhD Thesis**.
- Numerous awards and recognitions to the PIs.
  - **J. Alegre-Cebollada**. Michele Auger Award. ERC-CoG Awardee. Academia Joven member.
  - **F. Moreno-Herrero**. Bruker Prize. CEDACT member. Promoted to CSIC Professor. ERC-SyG finalist.
  - **M.A. del Pozo**. EMBO member. Member of Faculty Opinions, SAB Member F. GADEA-Ciencia
  - **R. Garcia**. CAM Miguel Catalán Prize to the scientific career.
  - **O. Llorca**. Panel member ERC, ICREA, Boehringer, AECC, RyC program. ERC-SyG finalist.
  - **G. Plaza**. Vocal ANECA. Subdirector Alumnos y Extensión Univ.





## Tec4Bio in numbers

- **Human resources**

- 45 undergrads and master students.
- 14 predoctoral researchers.
- 7 postdoctoral researchers.
- 1 visiting professor.
- 6 technicians.
- SAB (Scientific Advisory Board) made of 3 recognized professors.
- 18 internships from lab members (adding up to 30 months).
- 23 activities in postgraduate courses and degrees.
- 36 seminars and/or conferences organized by Tec4Bio members.

- **Funding**

- Up to **10M€** secured by Tec4Bio Principal Investigators.
  - **ERC CoG 2020.** J. Alegre-Cebollada
  - **La Caixa Health Research.** O. Llorca, F. Moreno-Herrero, M.A. del Pozo.
  - **CAM. Sinergicos 2018.** O. Llorca & F. Moreno-Herrero.
  - **MICINN-AEI.** All PIs have secured grants from the National Research System.





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# ¿Cómo hemos continuado?

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This was a new consortium in 2019

Now – consolidated and aiming for a **new proposal** in 2024!

- ➡ Novel methodologies
- ➡ To tackle key problems in DNA repair, Cardiovascular diseases and mechanobiology
- ➡ New member. Prof. Cristina Flors (IMDEA Nanociencia). Superresolution Microscopy



Tec4Bio Meeting May 2022

