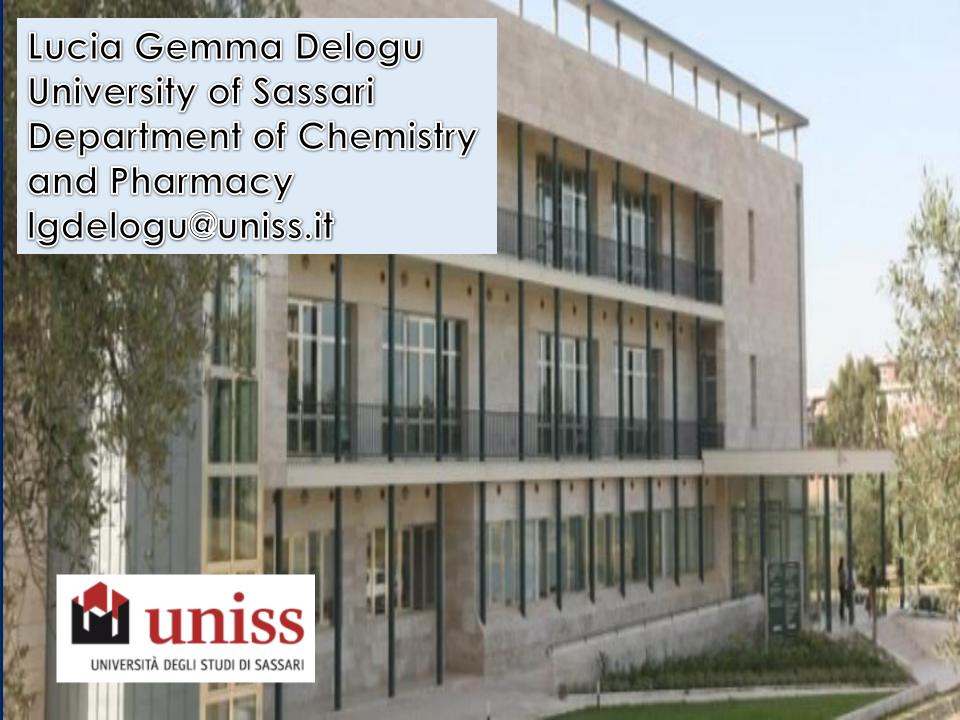


"Characterization of graphene immuneimpacts through omics approaches and genotoxic analysis"

FLAG-ERA JTC 2015 Project - Budapest 13/04/2016







#### Consortium Partners

UNIVERSITÀ DEGLI STUDI DI SASSARI	University of Sassari (Partner 1) Chemistry and Pharmacy Department (UNISS - Italy)	Dr. LUCIA GEMMA <b>DELOGU</b>
DEGLI JULIO DO TRIE	University of Trieste (Partner 2) Chemical and Pharmaceutical Sciences Department (UNITS - Italy)	Prof. MAURIZIO <b>PRATO</b>
cnrs	Centre National de la Recherche Scientifique (Partner 3) Immunopathologie et Chimie Thérapeutique (CNRS - France)	Prof. ALBERTO BIANCO
MANCHESTER 1824 The University of Manchester	<u>University of Manchester (Partner 4)</u> Institute of Inflammation and Repair (UNIMAN – United Kingdom)	Prof. KOSTAS KOSTARELOS
T946	Ankara University (Partner 5) Biomedical Engineering Department (UNIANK – Turkey)	Dr. ACELYA <b>YILMAZER</b>
	<u>University Hospital Cologne (Partner 6)</u> Institute for Genome Stability in Ageing and Disease ( <b>UKK – Germany</b> )	Prof. BJORN SCHUMACHER



# Graphene for biomedical applications

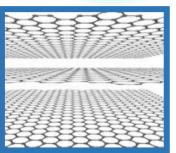
T cells

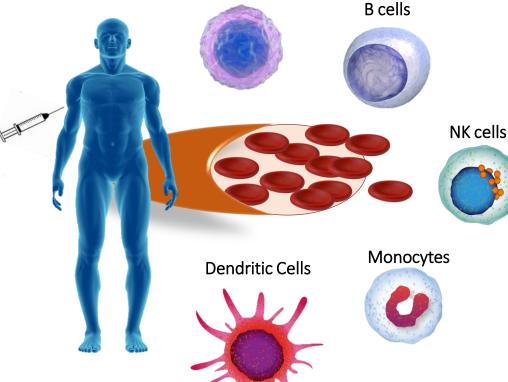
Gene delivery





Sensors

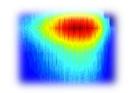




A STATE OF THE STA

**Photoacoustic** 

Diagnosis & Imaging

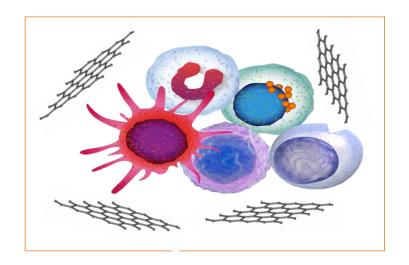


Ultrasonography





## **Overall Objective**



To complement Flagship research on graphene safety with immunogenomic and proteomic data.

G-IMMUNOMICS will provide new insights on the immune potential impact of several types of graphene.



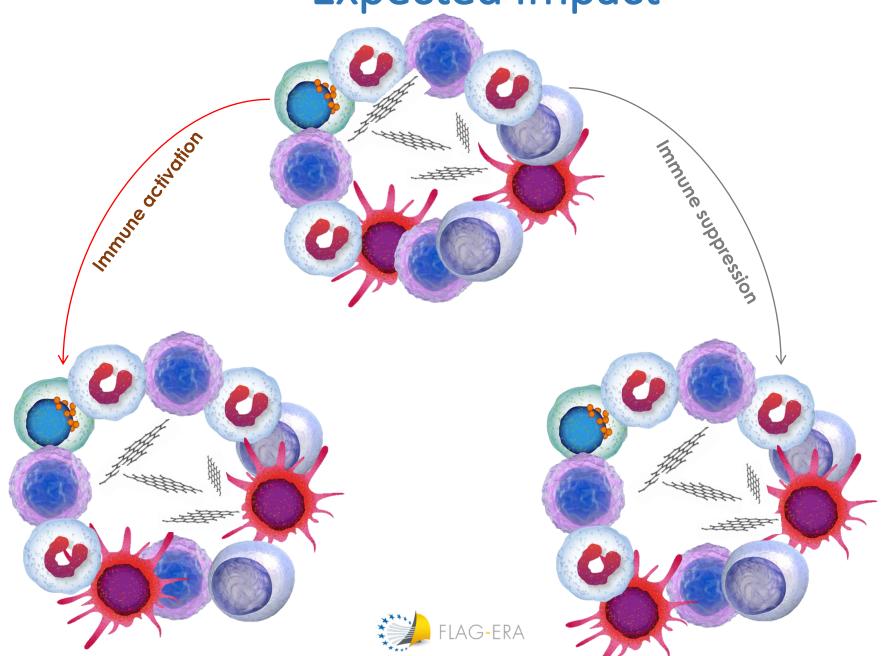
## **Expected Impact**

To Minimize the risk for human health raising from graphene usage.

To Identify the best candidate materials for translational applications in nanomedicine.

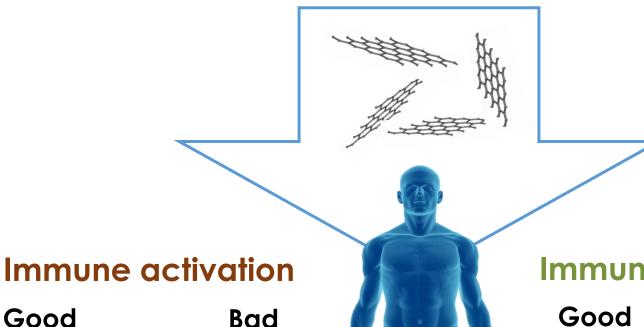


# **Expected Impact**





# **Expected Impact**



Good

Vaccine adjuvant **Immunotherapy** of cancer

Bad

**Hyper sesitivity** 

Inflammations

Immune suppression

**Anti inflammatory** action

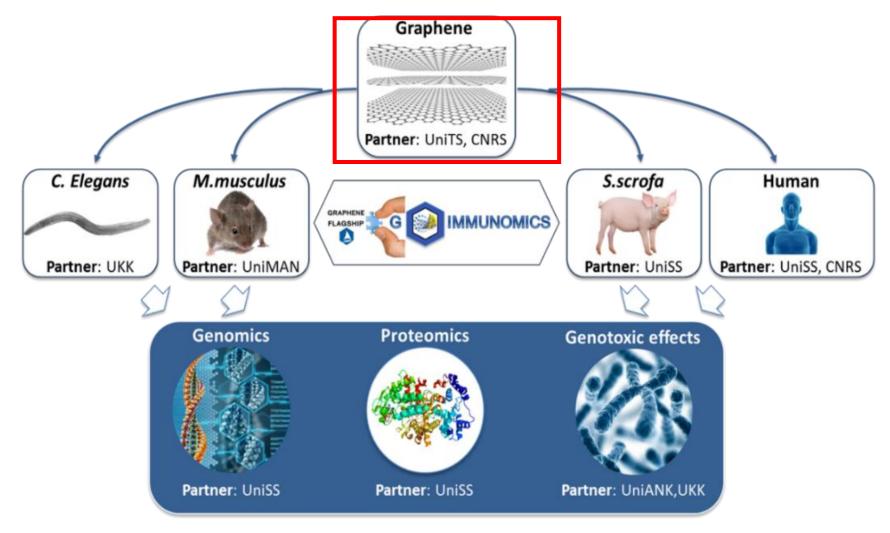
> No allergic response

Bad

Susceptibility to Pathogen infections



#### **Project Scientific Objectives**

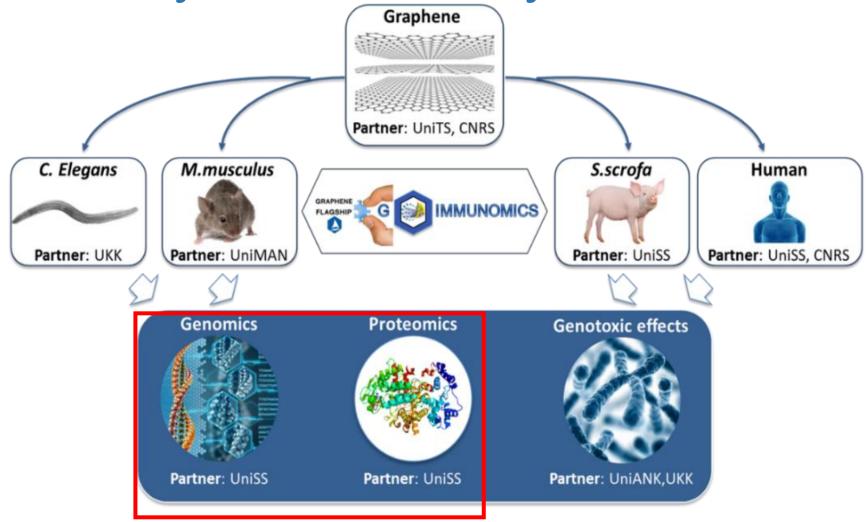


1) <u>Produce highly stable and dispersible pristine and functionalized GFMs</u> such as Exfoliated Graphene (G) and Graphene Oxide (GO) with different lateral size and appropriate functionalizations

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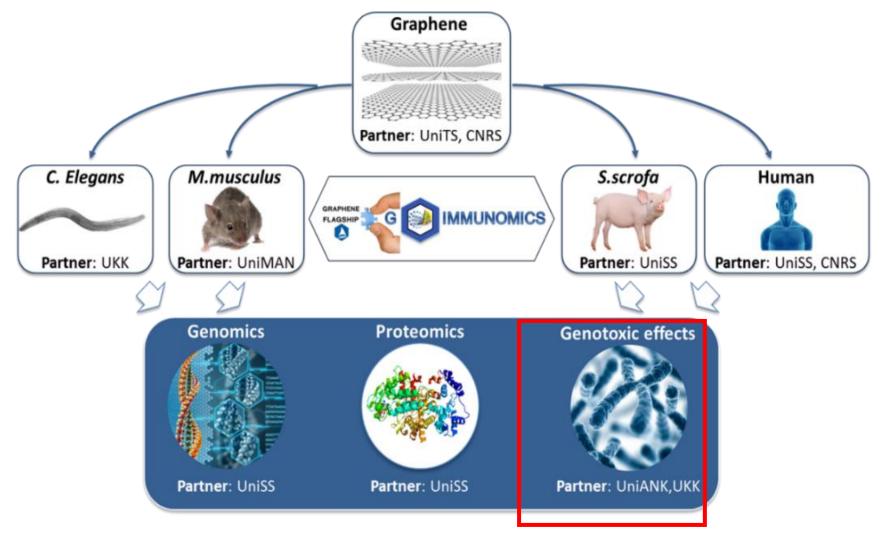
**Project Scientific Objectives** 



2) <u>Characterize</u> through high throughput functional immunogenomics and proteomics approaches the immune cell response induced by functionalized graphene on different cell lines (from human, mouse and swine) and primary cells (human, swine, mouse and nematodes)



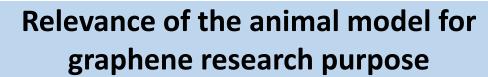
#### **Project Scientific Objectives**



3) Evaluate the genotoxic effects of the selected materials on different species: nematodes (*C. elegans*) and mammalians (*M. musculus, S. scrofa* and human).







- Remarkable consistency between gene expression profiles and transcriptional regulators in the mouse and human immune systems (conservatively estimated at 80%)
- Exceptional easiness to produce cancer models in comparison to other species in view of future applications



- Highly conserved genetic mechanism regulating the immune response and excellent anatomical similarities with human
- Compatibility with human ultrasound probes and waves and MRI systems suitable for the development of graphene-based imaging diagnostic tools



- The innate immune system response of *C. elegans* is initiated by conserved MAP kinase signaling cascades
- The DAF-16/FOXO mediated DNA damage response program induce an innate immune response which is conserved in higher species
- Fast cycle of reproduction

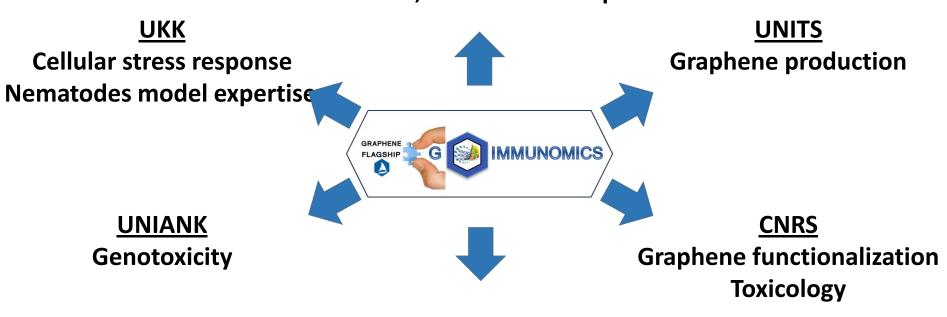




#### Consortium Expertise

#### <u>UNISS</u>

Immuno - Genomics / Proteomic Human cells; Swine model expertise



#### <u>UNIMAN</u>

Pharmaco-kinetics
Mouse model expertise



13/04/2016

## Partnership Disciplines

The partnership covers multidisciplinary expertise that combines

Genetics

**Bioinformatics** 

**Immunology** 

Pharmacology

Chemistry

Nanotechnolgy

**Biochemistry** 

Toxicology

Cell biology

Toxicology

Biology

Haematology

Molecular Biology

**Cell Biology** 

Molecular Biology

15



#### **G-IMMUNOMICS Work Plan**

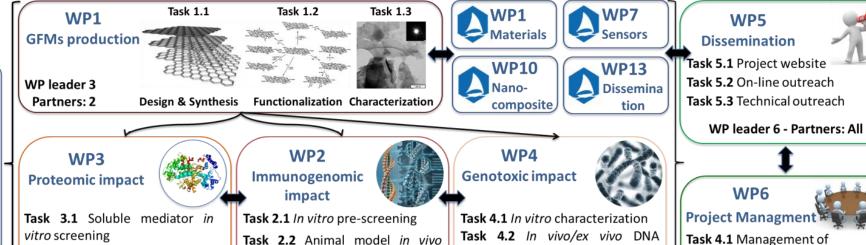


Task 3.2 In vivo screening

profiling

Task 3.3 Ex vivo human protein

WP leader 1 - Partners: 3, 4, 5



screening

Task 2.3 Human ex vivo screening

WP leader 1 - Partners: 3, 4, 6

damage screening

immunity

Task 4.3 DNA damage and innate

WP leader 5 - Partners: 1, 3, 4, 6

Consortium

Flagship

Task 4.2 Collaboration with the

WP leader 1 - Partners: All

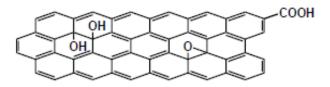


# Synergies with Flagship Core Project

FLAGSHIP WPs	Collaborations with flagship partners	
	A. Bianco, CNRS (FR)	
WP2- Health and Environment	K. Kostarelos, UNIMAN (UK)	
Environment	M. Prato, UniTS (IT)	
	E. Vazquez, UCLM (SP)	
WP10 – Nano-composite	V. Palermo, CNR-ISOF (IT)	
	X. Feng, TU DRESDEN	
WP13 Dissemination	Whole Flagship consortium	





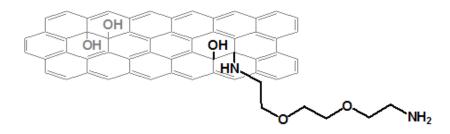


Graphene oxide= GO









Graphene oxide+ amidation= GONH<sub>2</sub>



#### Dissemination



Graphene 2016 Conference 19-22 April *Genoa Italy* 

Graphene Week
13-17 June Warsaw Poland