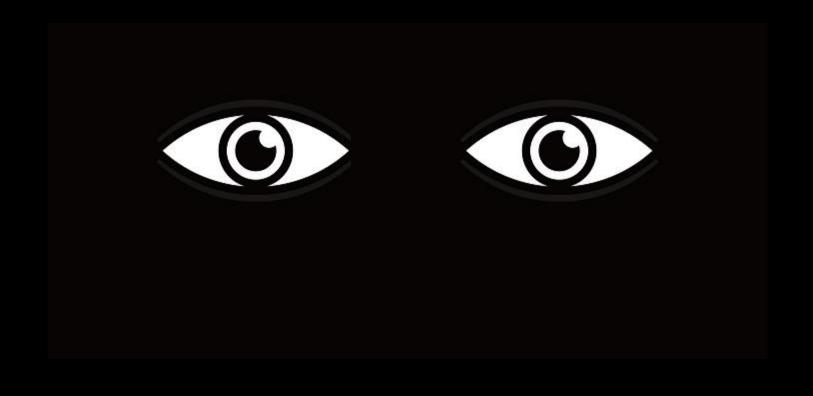
## Mechano-Wolbachia

Uncovering the mechanisms of action of an antiviral bacterium

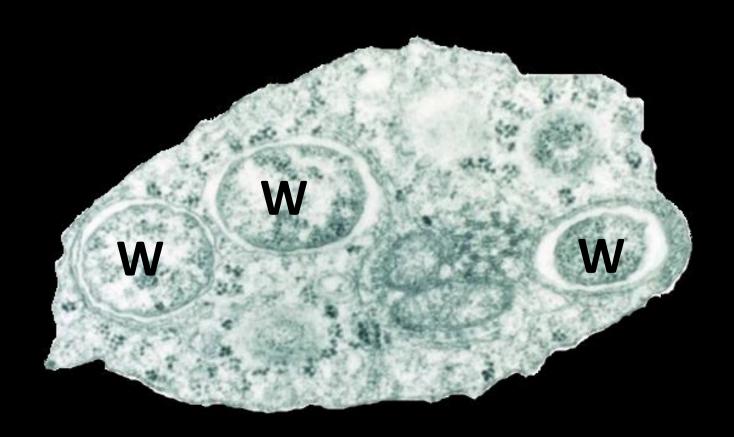


Ewa Chrostek
Jagiellonian University

# I have a vision of how to transform Wolbachia research



# Wolbachia Intracellular bacterium



# Wolbachia

# vvoibuciliu

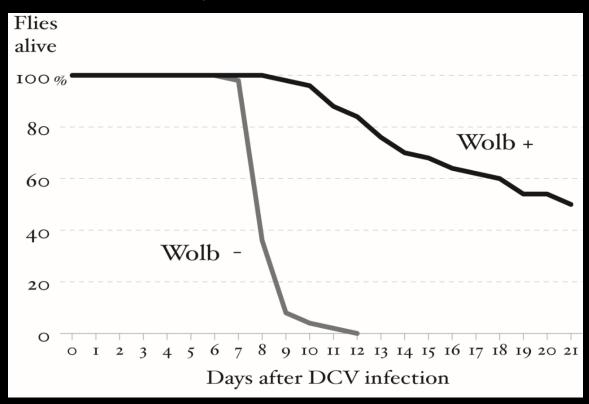
# Intracellular bacterium



### Wolbachia protects flies from viruses



#### Drosophila C virus infection

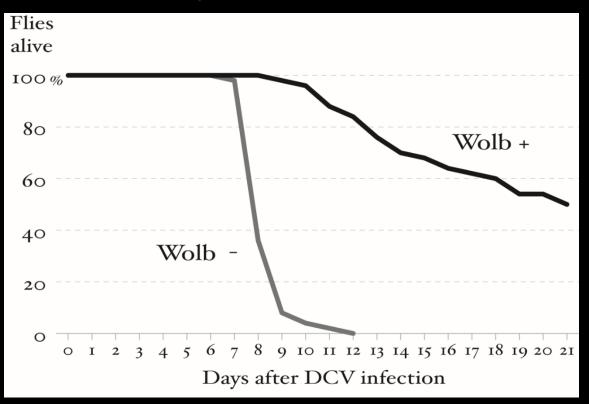


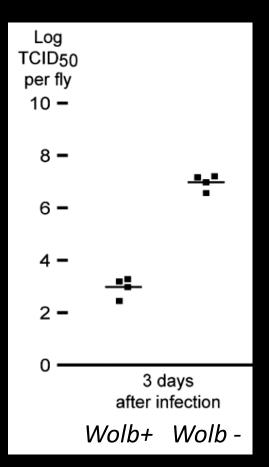
Modified from: Teixeira L et al., PLoS Biol 2008

### Wolbachia protects flies from viruses



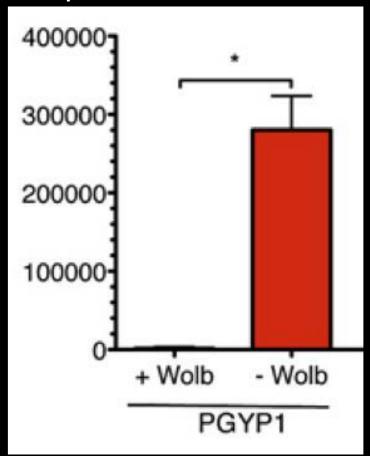
#### Drosophila C virus infection





# Wolbachia protects mosquitoes from viruses

Dengue virus RNA copies



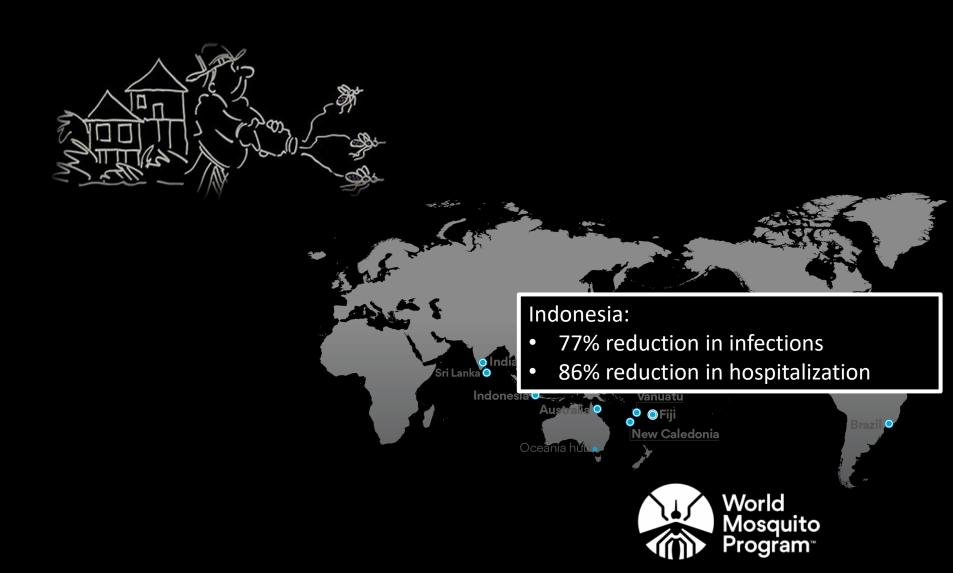


Moreira et al., 2009. Cell

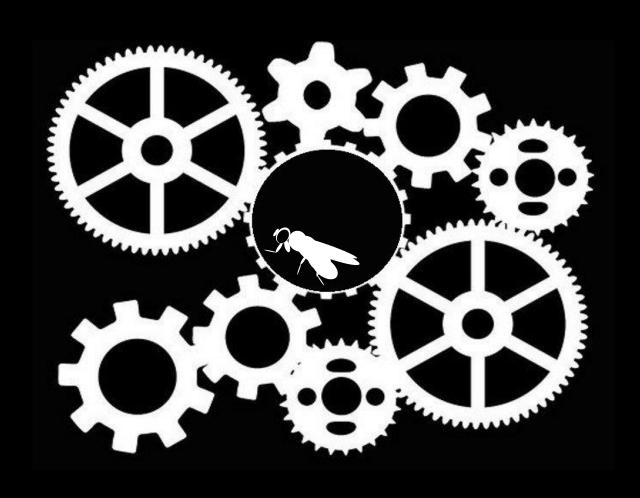
# Wolbachia-infected mosquito releases to fight dengue



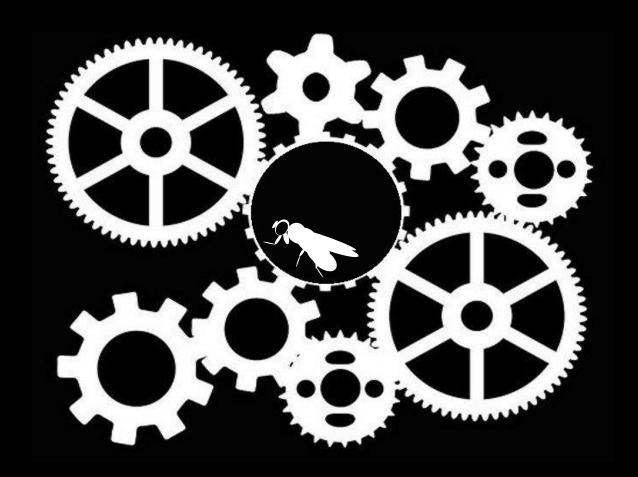
# Wolbachia-infected mosquito releases to fight dengue



# What is the mechanism of Wolbachia-conferred antiviral protection?



# What is the mechanism of Wolbachia-conferred antiviral protection?



None of the known antiviral mechanisms...

### ERC StG Mechano-Wolbachia

#### Obj. 1. Establish model system.

Obj. 1.1. Determine when and where protection starts. Discover which genes are involved.

Obj. 1.2. Validate cell culture model.

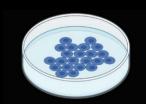
#### Obj. 2. Discover molecular bases of antiviral protection.

Obj. 2.1. Use novel tools to pinpoint genes causing protection.

Obj. 2.2. Develop new tools.











# 1.1. Determine when and where protection starts

**State-of-the art** 



Whole organisms



Late timepoints

# 1.1. Determine when and where protection starts

### **State-of-the art**

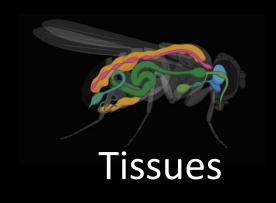


Whole organisms



Late timepoints

#### Mechano-Wolbachia





Early timepoints

### 1.1. Discover which genes are involved

#### **State-of-the art**



Whole organisms



Late timepoints

#### Mechano-Wolbachia





Early timepoints



Genes involved in protection

### ERC StG Mechano-Wolbachia

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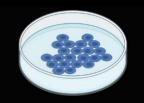
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Previously, no tools to study *Wolbachia* gene functions directly.

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Previously, no tools to study *Wolbachia* gene functions directly.

# I developed tools to do it!



## Long-term gains

#### **Scientific:**

- Discover mechanisms of Wolbachia phenotypes
- Develop tools to study other unculturable microbes
- Use Wolbachia as a vector in insects





# Long-term gains

#### **Scientific:**

- Discover mechanisms of Wolbachia phenotypes
- Develop tools to study other unculturable microbes
- Use Wolbachia as a vector in insects

#### <u>Societal:</u>

- Improve vector control efforts
- Preserve health of beneficial insects
- Use beyond Wolbachia or insects...











4+1 year

**Gulbenkian Institute** 

FCT PhD Fellowship

First genotype-phenotype link in unculturable *Wolbachia* 





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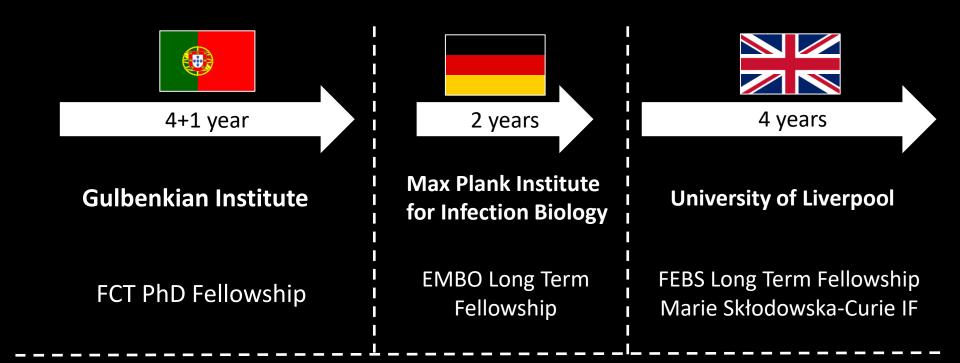


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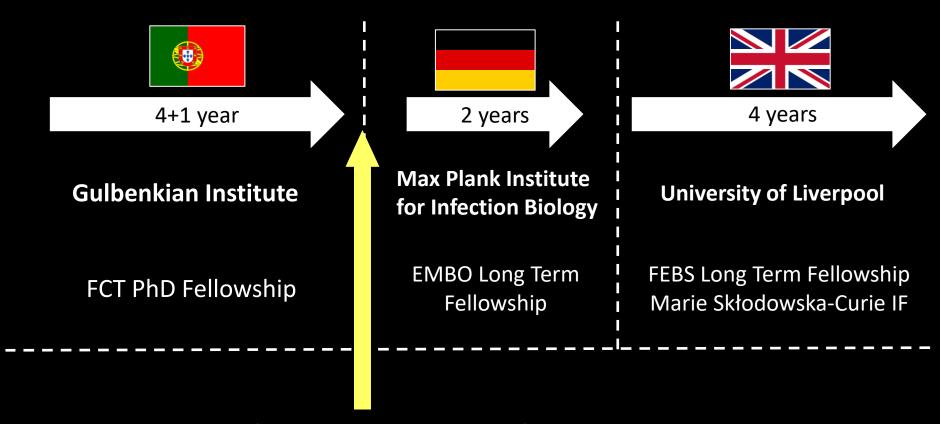
Tools to study Wolbachia gene function!!!



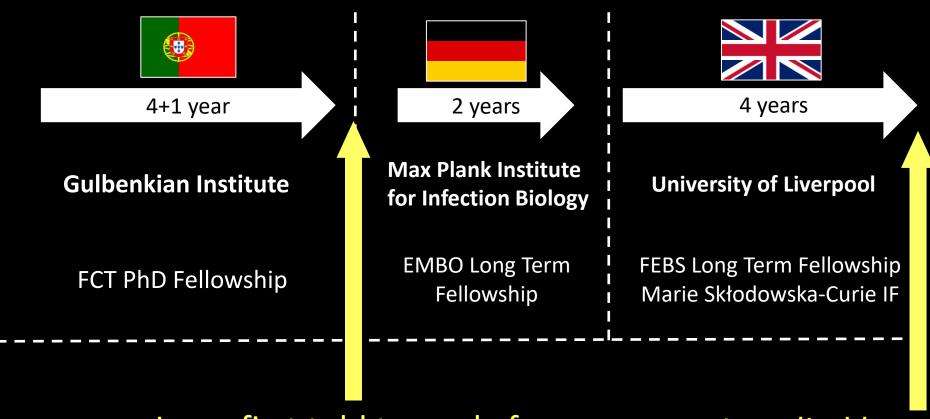


4 individual fellowships
10 first-author publications
8 invited talks

7 awards for professional achievements



I was first told to apply for an ERC StG on the day of my PhD Thesis defence!



I was first told to apply for an ERC StG on the day of my PhD Thesis defence!

I applied here

# What did I do for those 6 years?

### Who can apply?

Researchers of any nationality with 2-7 years of experience since completion of PhD, a scientific track record showing great promise and an excellent research proposal can apply.

### ERC is a grant for the best scientists:

Students of the best

Collaborators of the best

Mentees of the best

### Get an idea

### Combination of:

 Absolutely safe (but laborious and tedious) work no one has thought of doing before,

- Innovative/groundbreaking experiments.

# What has helped me get the grant?

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National Contact Point and the lessons learned

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National Contact Point and the lessons learned

**B1** 

#### National Contact Point and the lessons learned

**B1** 

CV & track record should tell a story

Stress on independence

International outlook

#### National Contact Point and the lessons learned

**B1** 

CV & track record should tell a story

Write to generalists

Stress on independence

Stress impact (if successful)

International outlook

Ask as many people as possible to read/provide feedback

#### National Contact Point and the lessons learned

**B1** 

**B2** 

CV & track record should tell a story

Write to generalists

Stress impact

(if successful)

Write to people in your field – you've got this!

Stress on independence

Ask as many

International outlook

people as possible to read/provide feedback

National Contact Point and the lessons learned

Interview

Think about it as a "ted talk"

Practice!

Questions: 3 sentences and stop talking

Mentors and people who care

Seek people who are "good at getting grants"

Help others so that you get help when needed

Network, network!

Ask, ask, ask!

The best science is brave; it breaks the convention and investigates new realms

Proposal must be innovative BUT you have to prove it is feasible.

Choose your panels wisely

PHYSICAL SCIENCES AND ENGINEERING (PE) LIFE SCIENCES (LS) SOCIAL SCIENCES AND HUMANITIES (SH)

#### Choose your panels wisely

LIFE SCIENCES (LS)

Integrative Biology: From Genes and Genomes to Systems (LS2)

Immunity, Infection and Immunotherapy (LS6)

Prevention, Diagnosis and Treatment of Human Diseases (LS7)

Environmental Biology, Ecology and Evolution (LS8)

#### Choose your panels wisely

LIFE SCIENCES (LS)

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Prevention, Diagnosis and Treatment of Human Diseases (LS7)

Environmental Biology, Ecology and Evolution (LS8)



Do not underestimate your CV

No Nature, Science, Cell papers

Stress on international outlook

Stress every achievement

# Acknowledgements







#### Team:

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-----

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**Angel Masaki** 

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**Andrea Betancourt** 

**Kayla King** (Oxford)

Raquel Oliveira (Oeiras)

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**Andrew Jackson** 

**Marcus Blagrove** 

**Alistair Darby** 

**Ben Makepeace** 

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Dominic Ferrandon (Strasbourg)

Luis Teixeira (Lisbon)

Frank Jiggins (Cambridge)

Casey Bergman (Athens, GA)











European Research Council
Established by the European Commission





# Thank you for your attention &

good luck with your proposals!