



Platform EGG

Mechanical characterization of human oocytes

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October 19th, 2021



Context of this work

Infertility

- In France : 10 - 16 % of couples have conceiving difficulty

Assisted reproductive technology ART

- In the world : 1.5×10^6 ART per year
- ART represents in France 1 birth / 34

ICSI

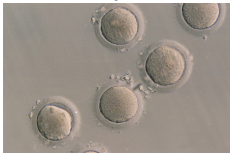
- France 2013 : 40 006 ICSI attempts
- Success rate **21.8 %**





ICSI procedure

- Ovarian stimulation
- Ovarian puncture within 36 hours



- Selection of oocytes on morphological criteria
- In vitro fertilisation and culture of embryos for 2-5 days



- Transfer to the uterus

Context of this work

Cause of miscarrying

- Technical problem
- Spermatozoon quality
- Oocyte quality

Oocyte choice criteria

- Morphological (classical)



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Context of this work

Cause of miscarrying

- Technical problem
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Oocyte choice criteria

- Morphological (classical)

Platform EGG

Produce objective mechanical criterion to help physicians determine which oocyte should be inseminated and transferred



Morphological selection criteria



VG



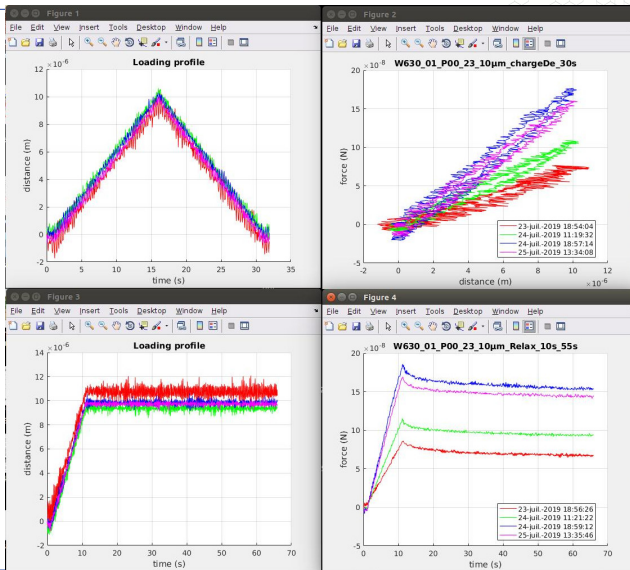
M1



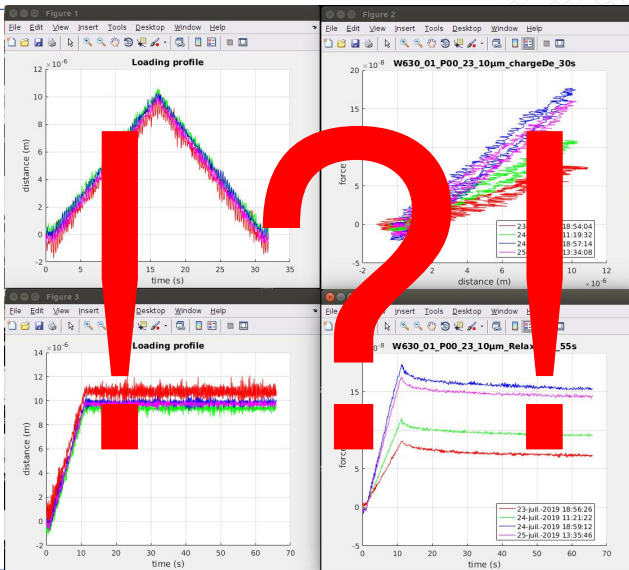
M2



Mechanical selection criteria

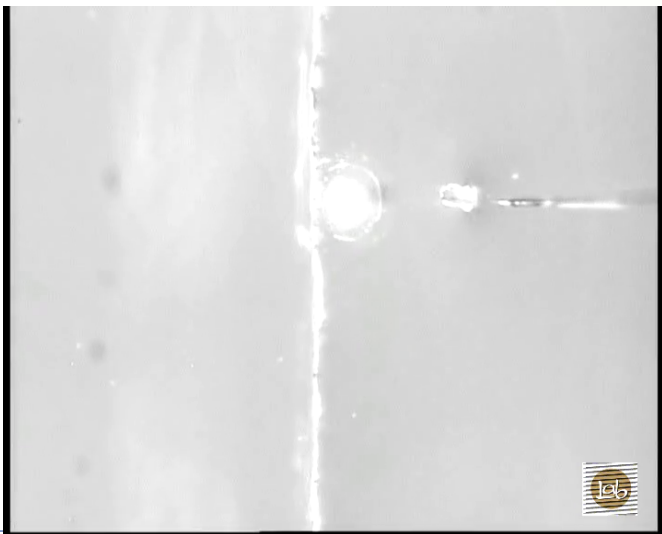


Mechanical selection criteria



A mechanical test

Our first Time





- 1 Concept of Magnetic spring
 - Simplest configuration
 - First force sensor design
 - Oocyte characterisation platform
 - Active magnetic springs
- 2 Capteur EGG
 - Global design
 - Magnetic springs design
- 3 Conclusion and perspectives



- 1 **Concept of Magnetic spring**
Simplest configuration
First force sensor design
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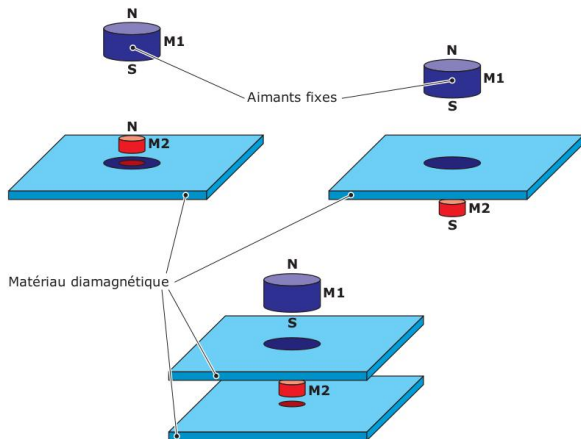
Magnetic levitation

Quite simple



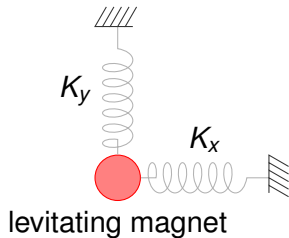
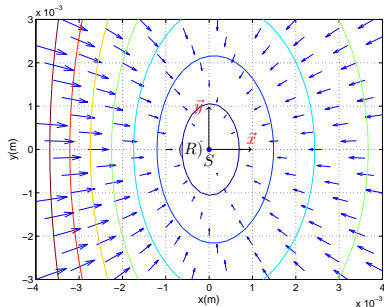
Magnetic levitation

Basics



Magnetic levitation

Horizontal behavior

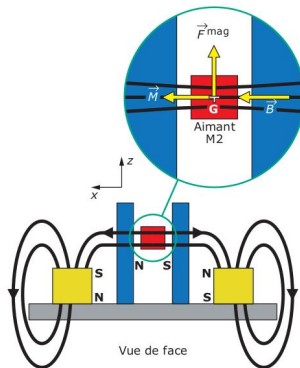
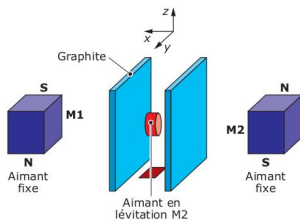


$$K_x \approx 0.01 \text{ N/m}$$

Displacement of $1 \mu\text{m} \implies$ force of 10 nN

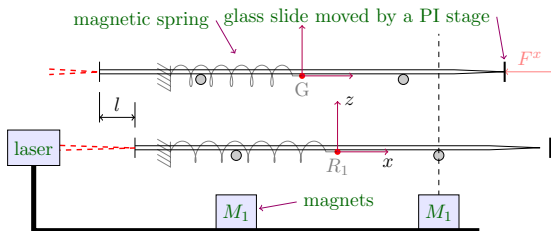
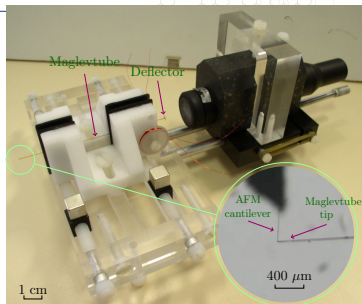
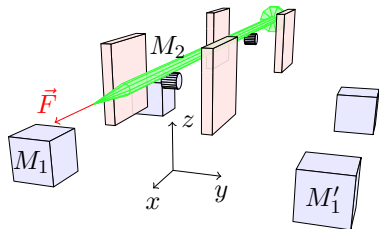
First force sensor design

Design modification



First force sensor design

Final design



First force sensor design

Application to oocyte characterisation

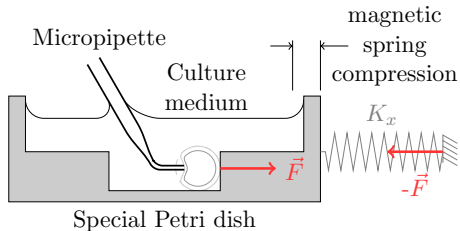
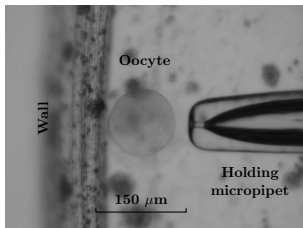


Problem of surface tension



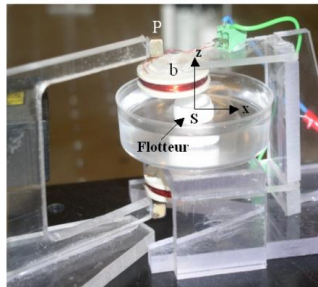
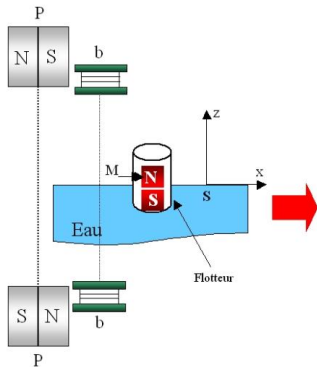
First oocyte characterisation platform

Force instrumented Petri dish



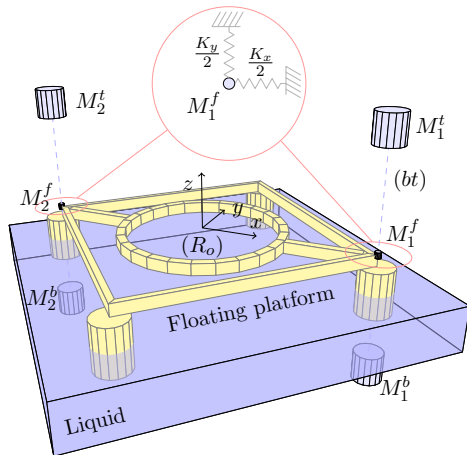
First oocyte characterisation platform

Using buoyancy



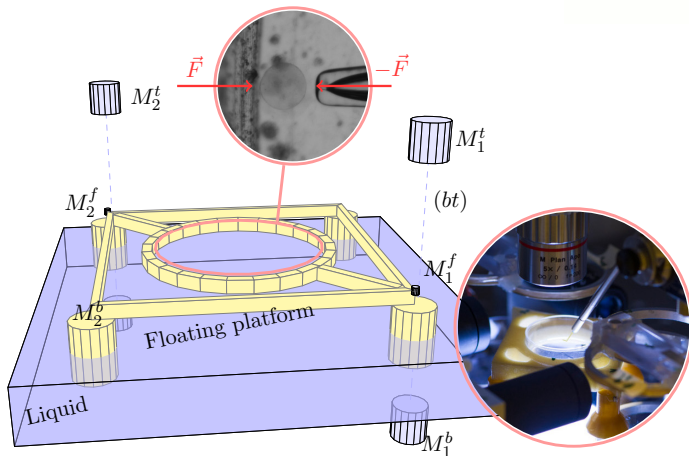
First oocyte characterisation platform

Entire platform



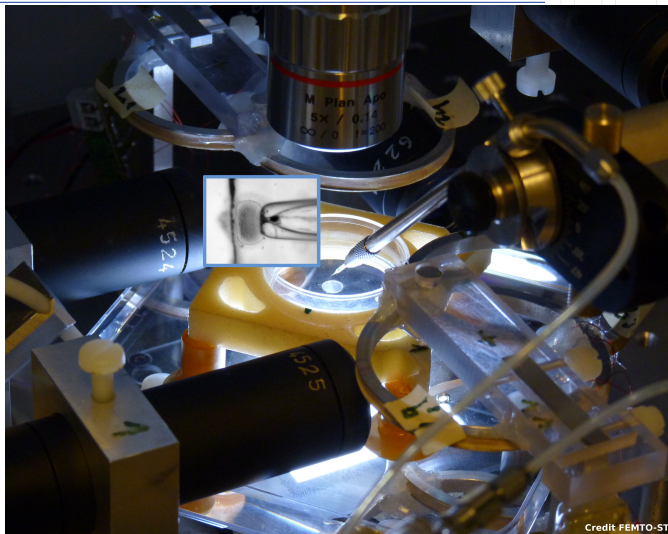
First oocyte characterisation platform

Entire platform



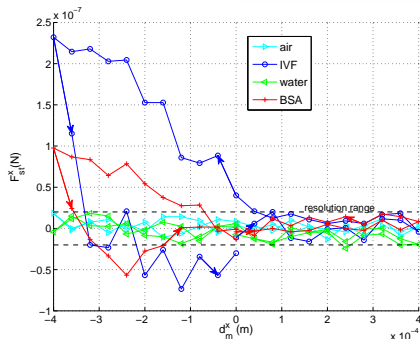
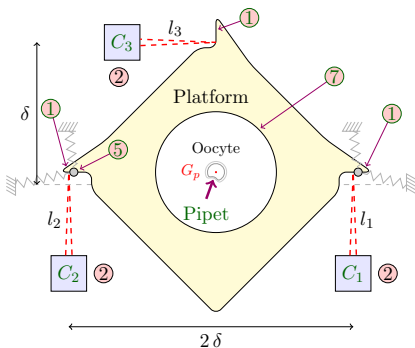
First oocyte characterisation platform

Entire platform



First oocyte characterisation platform

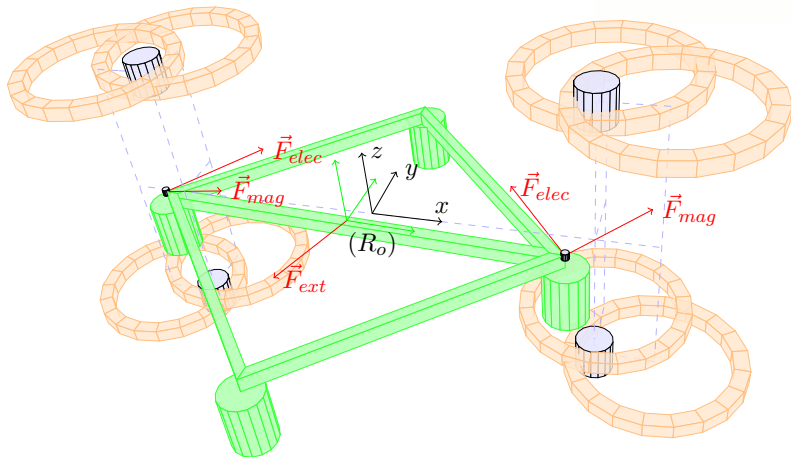
Results



Problem of surface tension remains

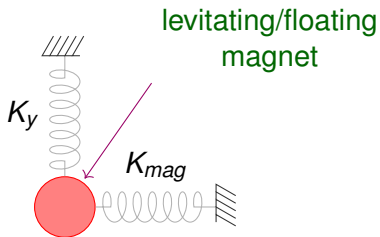
First oocyte characterisation platform

3 DOF active control



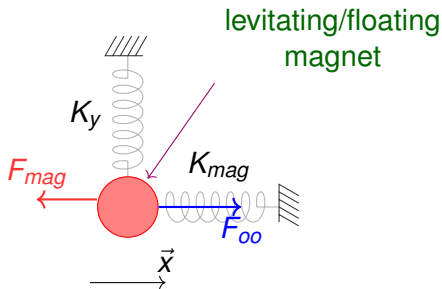
Active magnetic springs

Basics



Active magnetic springs

Basics



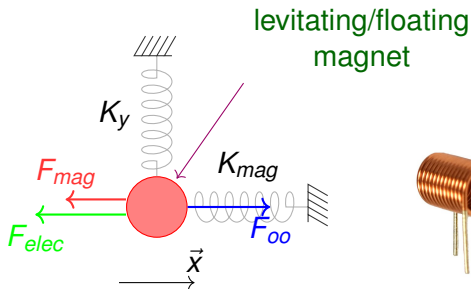
$$F_{mag} = K_{mag} \cdot x$$

 \implies

$$\hat{F}_{oo} = K_{mag} \cdot x_{measured}$$

Active magnetic springs

Basics

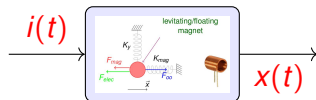


$$F_{mag} = K_{mag} \cdot x \quad \Longrightarrow \quad \hat{F}_{oo} = K_{mag} \cdot x_{measured}$$

$$F_{elec} = K_{elec}(x) \cdot i \quad \Longrightarrow \quad \hat{F}_{oo} = K_{elec} \cdot i_{measured} + K_{mag} \cdot x_{measured}$$

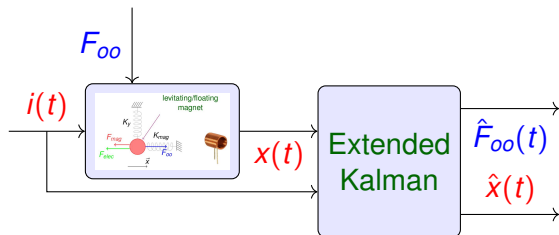
Active magnetic springs

Position active control



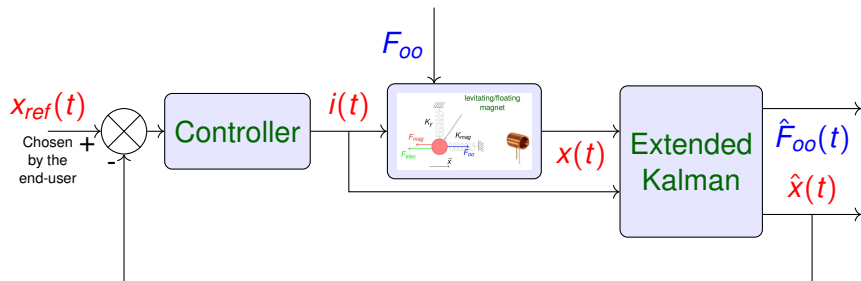
Active magnetic springs

Position active control



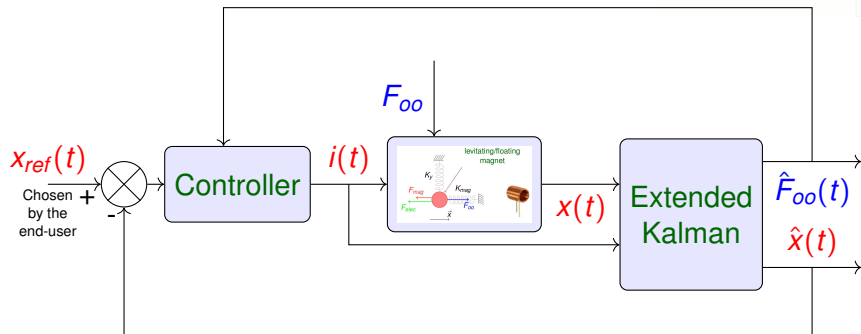
Active magnetic springs

Position active control



Active magnetic springs

Position active control



VIRCO : Virtual Input Rejection COntrol



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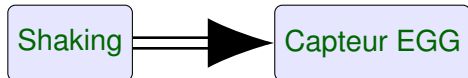
Capteur EGG

Design



- Magnetic levitation
 - Magnetic springs
 - Buoyancy
 - Active magnetic springs
 - Unknown input observer (Kalman or GELESO filters)
 - Advanced robust control law (VIRCO)
-

- Medical and biological requirements
- habits of ART centers



Platform EGG

In ART center of Besançon hospital





1. W02018172688 - DISPOSITIF POUR LA CARACTERISATION MECANIQUE D'UN ELEMENT D'INTERET PAR EXEMPLE UN OVOCYTE

[Données bibliographiques PCT](#) [Description](#) [Revendications](#) [Dessins](#) [ISR / WOSA / A17 \[2\] \(a\)](#) [Phase nationale](#) [Famille de brevets](#) [Notifications](#) [Documents](#)

[Lien permanent](#) [Traduction automatique ▼](#)

Numéro de publication

W0/2018/172688

Date de publication

27.09.2018

N° de la demande internationale

PCT/FR2018/050670

Date du dépôt international

20.03.2018

CIB

G01N 3/40 2006.1 G01N 3/42 2006.1

C12M 1/42 2006.1

CPC

C12M 35/06 C12M 41/48 G01N 2203/0078

G01N 2203/0085 G01N 2203/0087

G01N 2203/0089

Voir plus de classifications

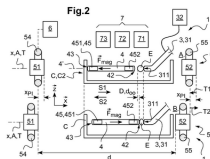
Déposants

CENTRE NATIONAL DE LA RECHERCHE

Titre

[EN] DEVICE FOR MECHANICALLY CHARACTERIZING AN ELEMENT OF INTEREST SUCH AS AN OOCYTE

[FR] DISPOSITIF POUR LA CARACTERISATION MECANIQUE D'UN ELEMENT D'INTERET PAR EXEMPLE UN OVOCYTE



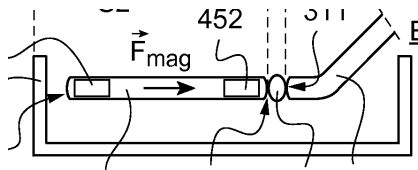
Abstré

[EN]

The invention relates to a device for mechanically characterizing an element of interest [E], for example an oocyte. The mechanical characterization device comprises: - support means [2] for receiving a container [C] suitable for containing a liquid medium, - holding means [3] for holding said element of interest [E], - an indenting member [4], - magnetic means [5] for generating a magnetic field in which said indenting member [4] is intended to move and which participates in suspending said indenting member [4] with an unstable horizontal direction (x) oriented coaxially to the longitudinal axis (z), - control means [6], intended to control said magnetic means [5] so as to maneuver the indenting member [4] in translation along the unstable horizontal direction (x), and - means [7] for

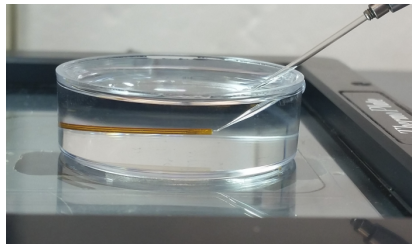
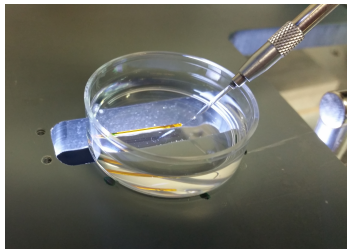
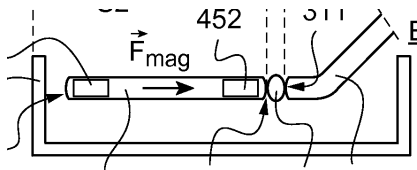
Platform EGG

Magnetic springs inside the Petri dish



Platform EGG

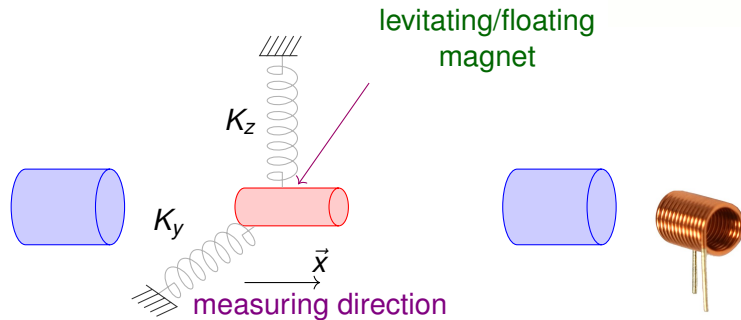
Magnetic springs inside the Petri dish



Single use magnetic glass indenter of 16 mm length and 0.8 mm diameter

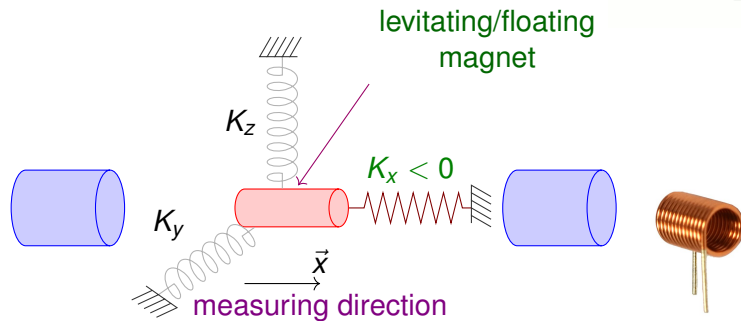
Platform EGG

Negative stiffness magnetic spring



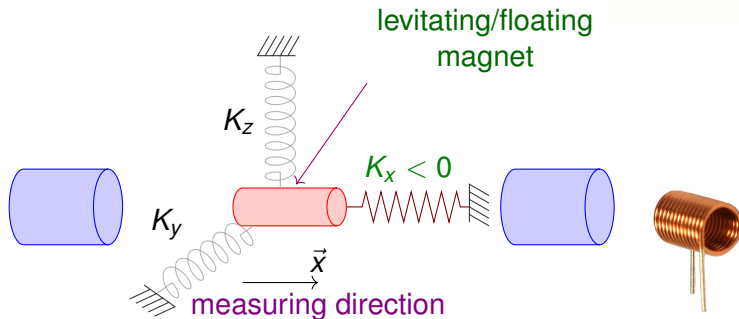
Platform EGG

Negative stiffness magnetic spring



Platform EGG

Negative stiffness magnetic spring



$$K_x \approx -0.001 \text{ N/m}$$

Unstable behavior along \vec{x}

Displacement of $1 \mu\text{m} \implies$ force of 1 nN

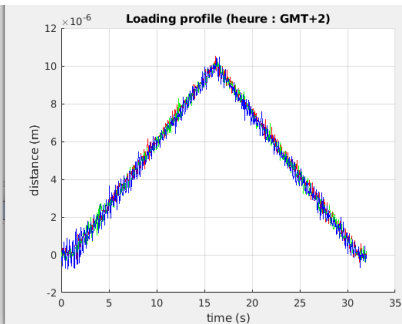
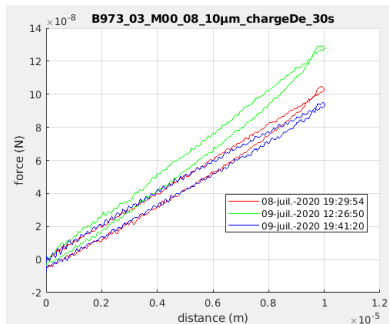
Platform EGG

Conducting experiments on oocytes



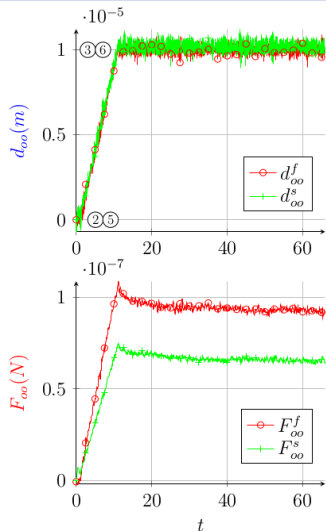
Oocytes mechanical properties

Loading and unloading tests



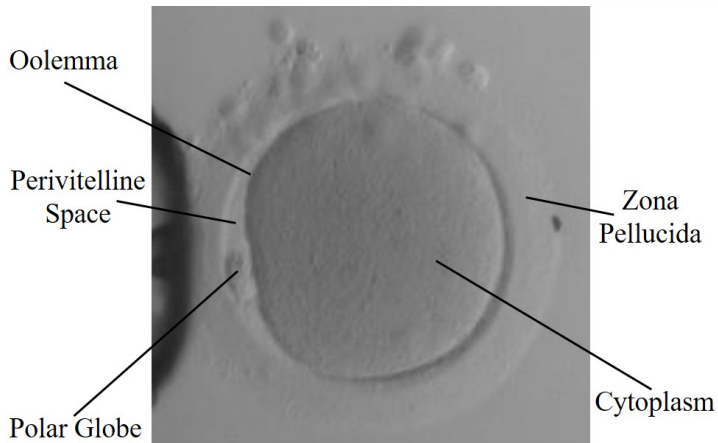
Oocytes mechanical properties

Relaxation tests with flat or sharp tip



Oocytes mechanical properties

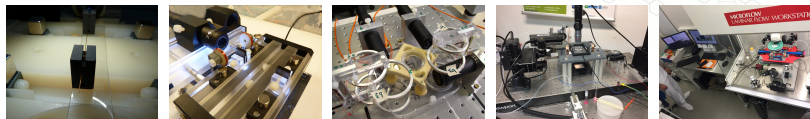
Oocyte constitution





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Conclusion and perspectives



- Approximately 80 supernumerary oocytes already tested
- Each oocyte exhibit a particular mechanical profile

Next steps

- Fine modelling of the different oocyte parts
- Clinical trial on 20 patients...

Thank you for your attention...



Contributors

Mehdi Boukallel, Ali Cherry, Stéphane Oster, Racha Gana, Juan Antonio Escareno, Margot Billod, Reda El Hirech, Fadoua Nana Najim, Jorge Andres Perez, Francois Vuillemin, David Purwins, Mickael Ohruh, Fawzia Amokrane, Mélanie Béduer, Romain Merillo, Danielle Lyne Cambou, Benjamin Heinzman, Ferdinand Schäffer, David Grams, Zhuldyzay Temirzhanova, Rachid Laydi, Adrien Drouot, ...