EPFL

Master in Electrical and Electronics Engineering

EE-517: Bio-Nano-Chip Design

Practical Sessions

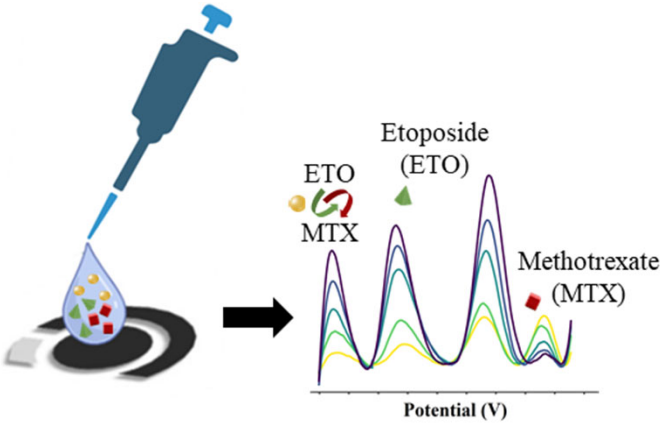
Every Tuesdays 11.00-13.00

- Projects
- Exercise

Francesca

francesca.rodino@epfl.ch

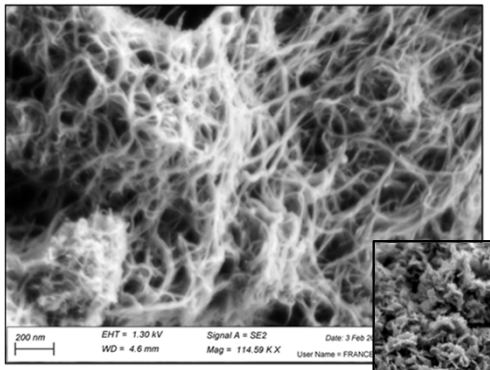
- PhD student at Bio/CMOS Interfaces Laboratory
- Education: Biomedical Engineering (B.Sc. and M.Sc.)
- **Ph.D. project**: Developing electrochemical biosensors and front-end electronics for therapeutic monitoring of multiple chemotherapeutic drugs for personalized medicine.



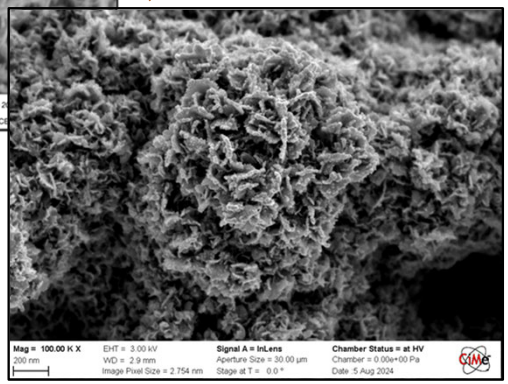
Bio

Specific functionalization (i.e., enzymes)

Nano Pt structures



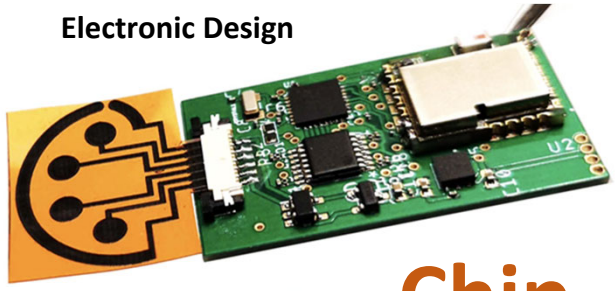
Carbon nanotubes (CNTs)



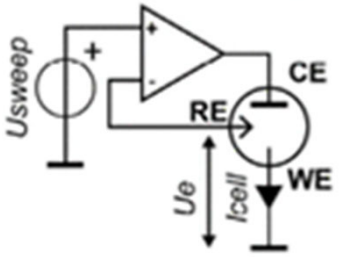
Nano

Portable PoC device

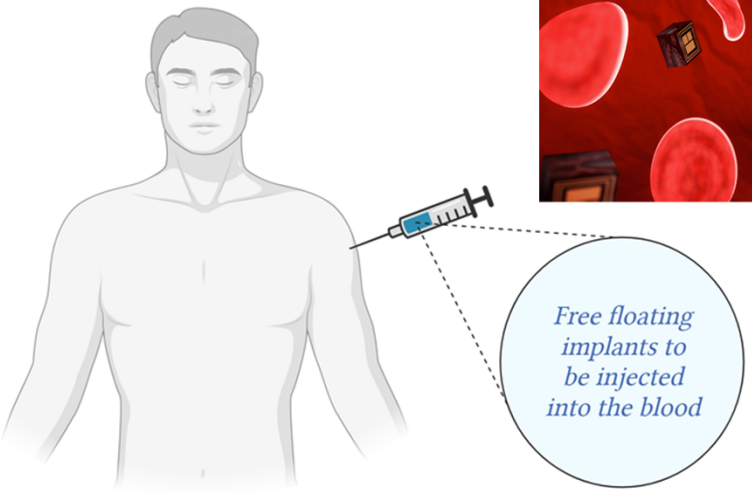
Electronic Design

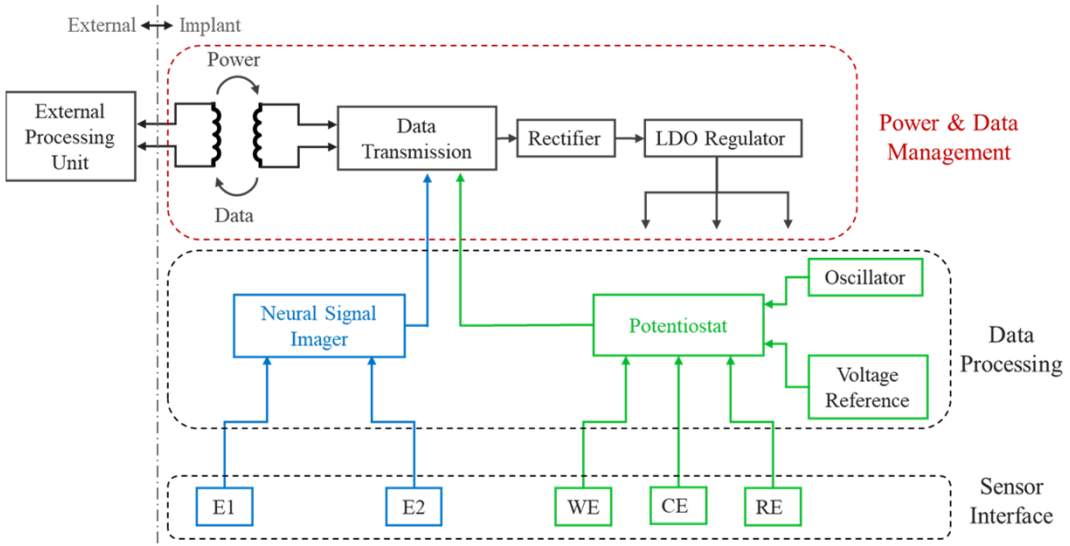


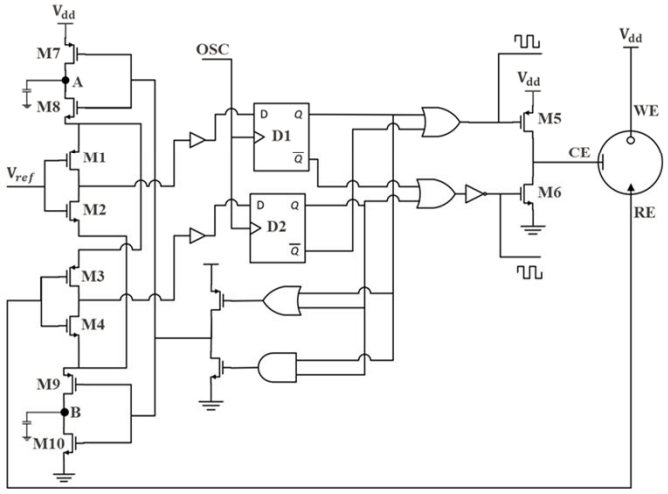
Chip

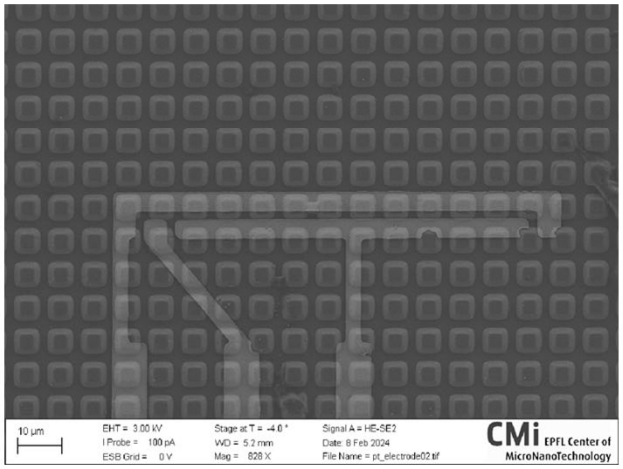


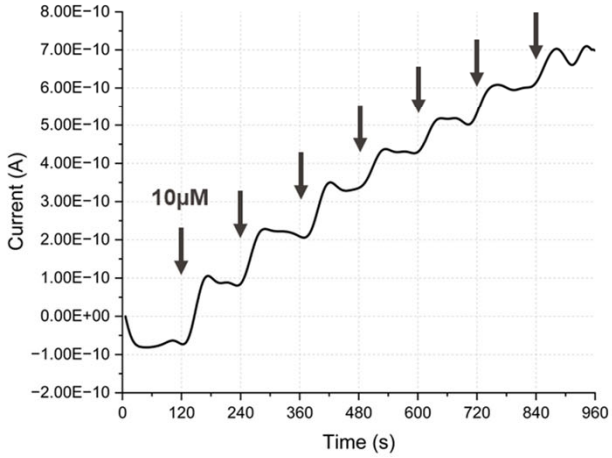
Ali


Free floating implants to be injected into the blood


External
Implant
Power
External Processing Unit
Data Transmission
Rectifier
LDO Regulator
Data
Power & Data Management
Oscillator
Neural Signal Imager
Potentiostat
Voltage Reference
Data Processing
E1
E2
WE
CE
RE
Sensor Interface


Vdd
M7
A
M8
OSC
D1
D2
M1
M2
Vref
M3
M4
M9
B
M10
M5
M6
CE
WE
RE

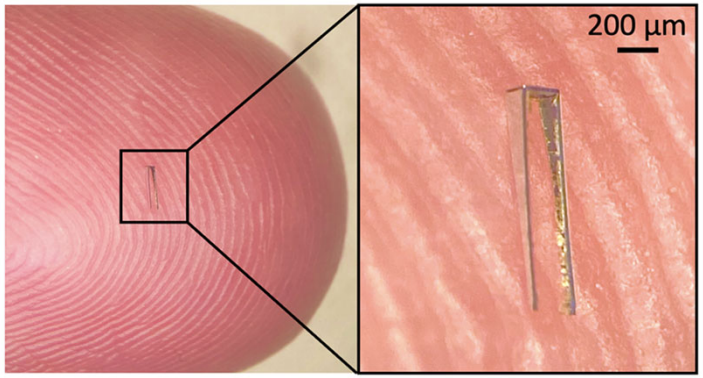

10 µm
CMi EPFL Center of MicroNanoTechnology

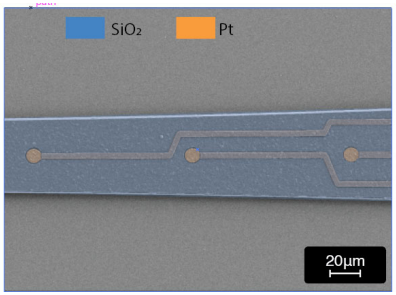

8.00E−10
7.00E−10
6.00E−10
5.00E−10
4.00E−10
3.00E−10
2.00E−10
1.00E−10
0.00E+00
−1.00E−10
−2.00E−10
Current (A)
10µM
0
120
240
360
480
600
720
840
960
Time (s)

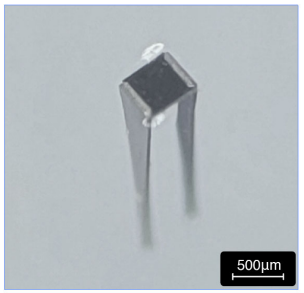
Junyan

- PhD candidate in Electric and Electronic engineering
- Bachelor and Master in Micronano technology

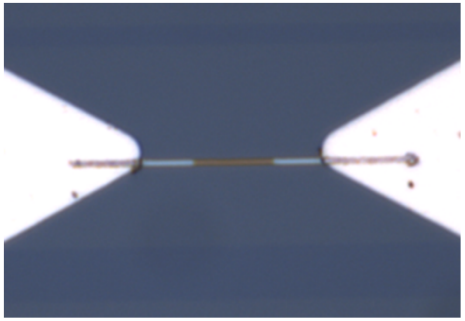
Foldable CMOS integrated probes

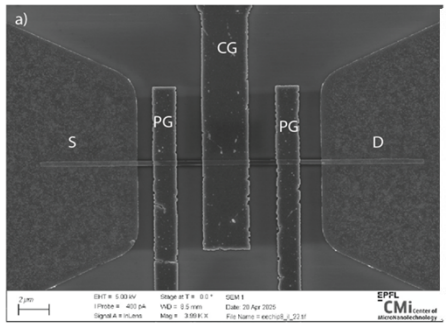


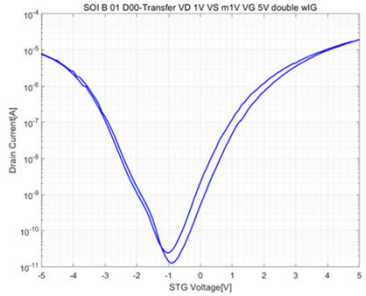


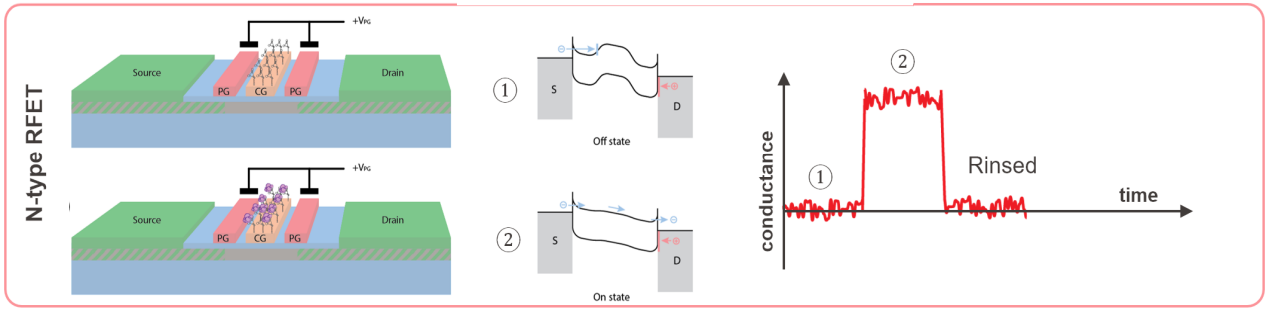


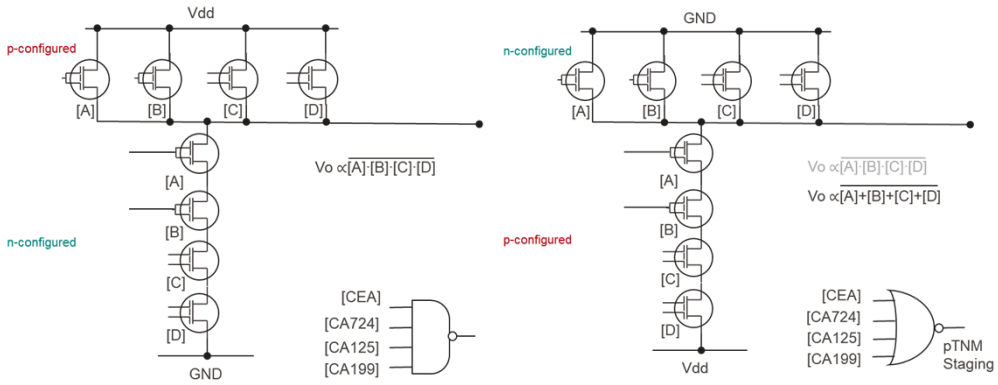
Reconfigurable FET based bio sensing platform



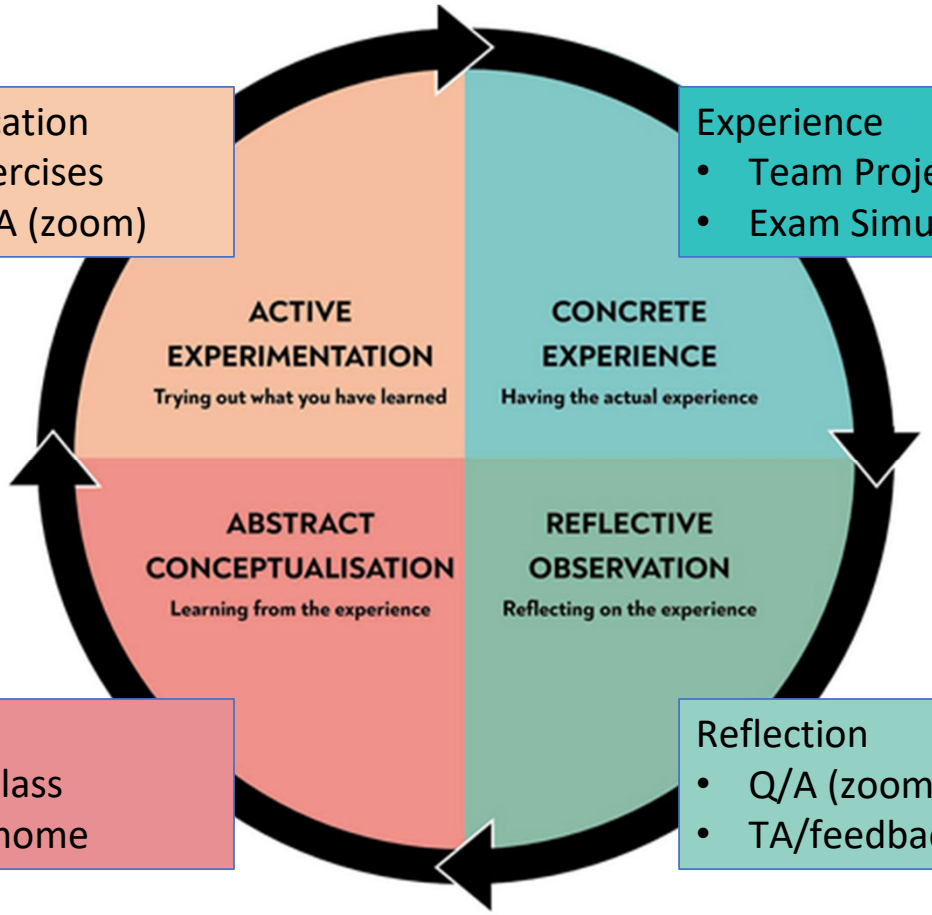








Kolb experimental learning cycle



Application
- Exercises
- Q/A (zoom)

Experience
- Team Projects
- Exam Simulation

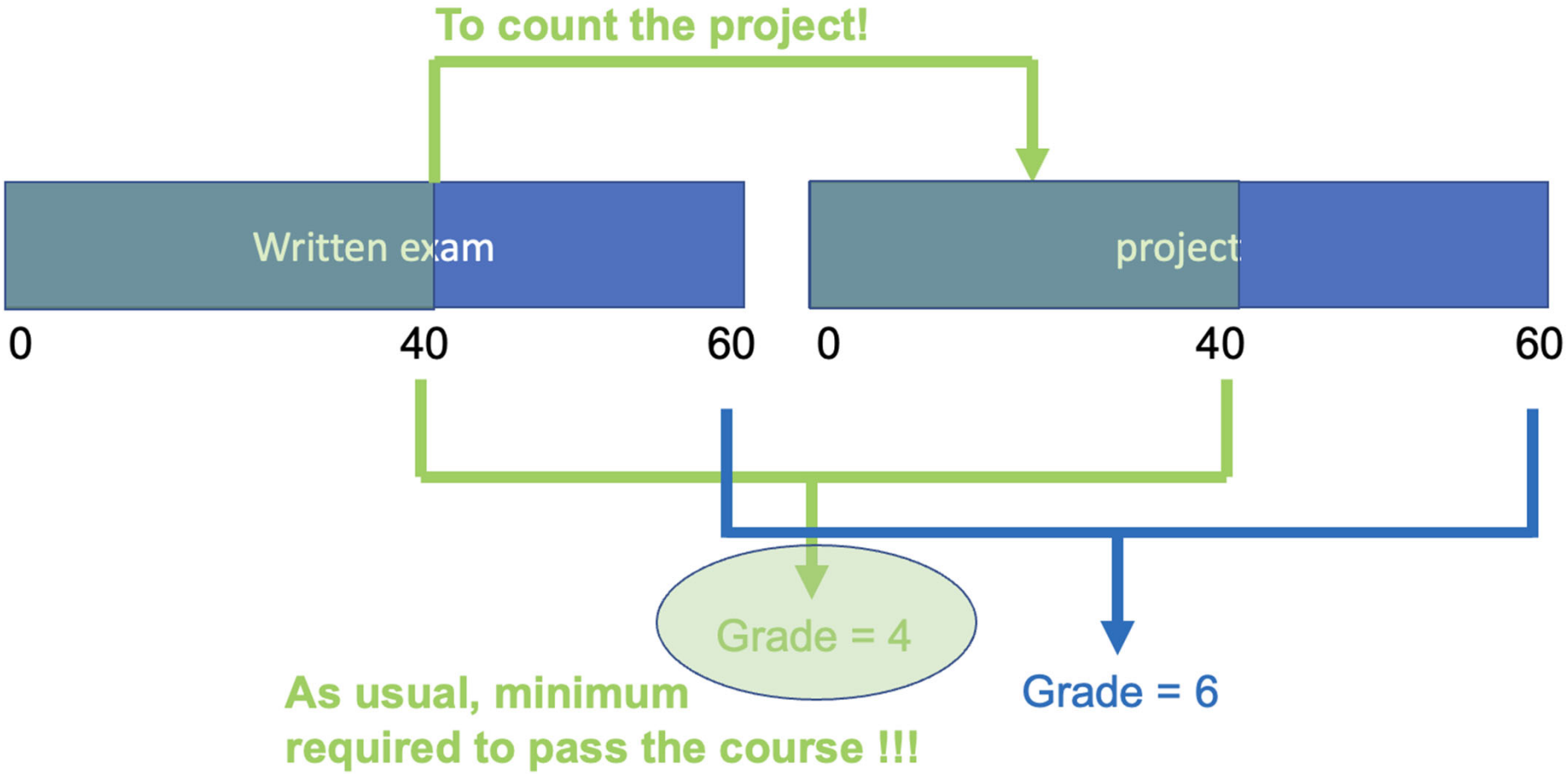
Theorization
- Front Lectures @class
- Personal Study @home

Reflection
- Q/A (zoom)
- TA/feedback

Project Rules

1. The project is **mandatory**.
2. The Project counts for 60 points maximum on the final grade.
3. The Written exam also counts for 60 point maximum on the final grade.
4. **Obtaining 40 points out of 60 in the written exam is mandatory for the project to be counted.**
5. **The Final Grade** is obtained by adding all the gained points on 120 points maximum, and then reporting the final grade on 6 maxima.

Project Rules



Project Rules

6. Students are **individually required to propose a project** (following the project template form) **within week 2 - Friday, 17.00**. Proposals must be sent to all TAs by email in pdf format (object: "EE-517 Project", filename: <EE-517_proposal_name_surname>)
7. The TAs will evaluate all the individual proposals based on the overall quality, final scope and timeline feasibility. The TAs will attribute final projects and working groups before week 3.
8. Every week, during the proejct session, the TAs will supervise the groups in their own project.
9. **Groups are required to submit one progressive report** following the guidelines and the template given by the TAs to receive their feedback and comments about the progress of their work. Progressive report must be sent to all TAs **within week 9 - Friday, 17.00** by email in .pdf format (object: "EE-517 Project", filename: <EE-517_group#XX_Progressive>)
10. **A final report** (following the IEEE template provided) **will be required from each group within 31st of December**. The final report must be sent to all TAs by email in .pdf format (object: "EE-517 Project", filename: <EE-517_group#XX_final>)
11. The TAs will evaluate the reports assigning the final mark, based on the technical quality, completeness, and clarity.
12. In order to obtain the maximum grade, the project needs to be focused on electrochemical detection only and needs to include at least the three aspects covered by the course: Bio, Nano, and CMOS.
13. The **Project evaluation** is done based on the following aspects**:**
    a. **Individual proposal: 15 points (individual mark)**
    b. **Progressive report: 15 points (group mark)**
    c. **Final report: 15 points (group mark)**
    d. **Individual queries during the sessions: 15 points (individual mark)**

    → **Oral presentation (last class, 16/12)**
14. **Motivational hint: exceptionally good-projects might have the possibility to be submitted to an internationally-recognized and peer-reviewed scientific conference.** At least, along the years, it happened sometimes! For example, the paper titled "Portable Breathalyzer for Exhaled Volatile Organic Compounds Monitoring in Lung Diseases" by Alec Chevrot, Justina Venckute, and Sarah Cuesta; published in the Proceedings of the IEEE Medical Measurements and Applications (MeMeA), held in 2022.
15. People of reference of project supervision and management:
    Francesca Rodino (francesca.rodino@epfl.ch) - Junyan Qian (junyan.qian@epfl.ch)- Ali Meimandi (ali.meimandi@epfl.ch)

Practical Sessions

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Ali Meimandi
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If you are interested in semester/master projects: https://www.epfl.ch/labs/bci/student-projects

Weekly schedule

	Objectives of the class	Deliverables	Deadline	TA
Week 1: Introduction to bio-nano-chip design and conductive solutions	- Finding a brilliant project idea! - Evaluating the novelty of the idea by literature search	Individual project proposal	Week 2 - Friday, 17h00	Fra, Junyan, Ali
Week 2: Probes/Targets Building Blocks	- Project proposal writing according to the template			Francesca
Week 3: Probe/Target interactions	- Contact and meet team members as soon as possible - Focus on literature review to identify similar published works - identify biomaterial to sense and from which biofluid and identify required specification			
Week 4: Probe Detection Principles (Faradaic Processes)	- identify the strategy to recognize the chosen target molecule - Identify method (CV, CA, ...)			
Week 5: Probe Detection Principles (with Antibodies and DNA)	- Calculate area/ sensitivity			
Week 6: Probes immobilisation	- Literature analysis - Identify nanomaterials that can be used for increasing sensitivity and creating selectivity			Junyan
Week 7: Checking Probes-layer quality (RM+SPR+SEM+AFM)	- Identify possible undesidered interactions with other molecules and think how to prevent it			
Week 8: Nanotechnology to prevent Electron Transfer	- Identify sensitivity and specificity of the designed bio-nano probe Calculate/Discuss the improved sensitivity			
Week 9: Nanotechnology to enhance Electron Transfer	- Wirte group progressive report for weeks 1 to 9 (follow provided template/guidelines)	Group progressive report	Week 9 - Friday, 17h00	
Week 10: CMOS Building Blocks	- Identify the adequate method for electrochemical sensing and design a signal conditioning unit for the proposed electrochemical sensor			Ali
Week 11: Circuits for metabolites detection in Fixed-Voltage	- Communication synthesis; - Unified system-level synthesis of Bio-Nano-CMOS-sensing device			
Week 12: Circuits for metabolites detection in Scanning Voltage	- Simulation of the proposed design using the LTspice software or similar			
Week 13: CMOS Circuits for DNA Detection	- Complete the front-end of the sensor at system level or transistor level by searching the market or designing the CMOS circuit - Presentation preparation	Group presentation	Week 14 - 11.00-13.00	
Week 14: Review	- Groups presentations - Final project report 3 pages (+1 page references) IEEE template	Final group project report	31.12.2024	Fra, Junyan, Ali

Summary – Deliverables over the semester

1. Individual project proposal
Deadline: week 2 - Friday, 17.00

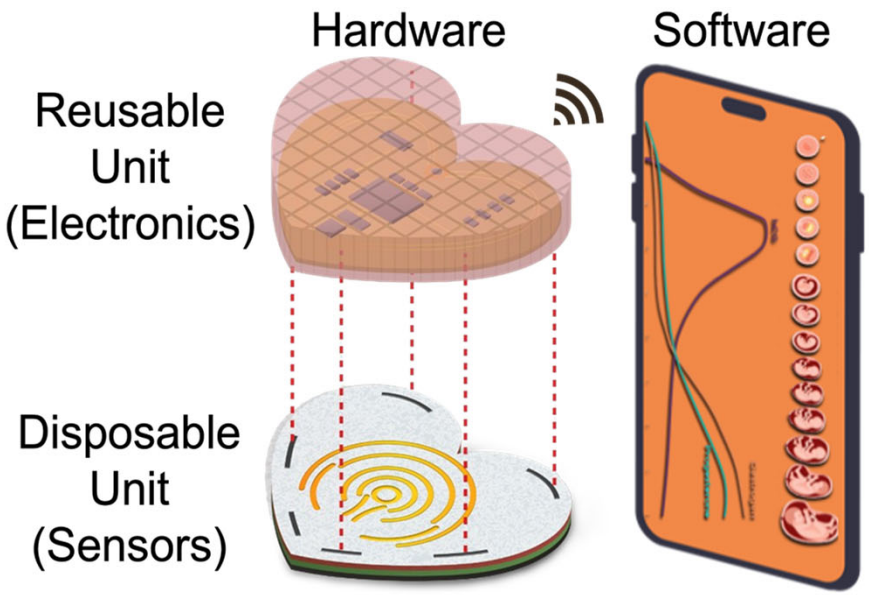
2. One progressive report
Deadline: week 9 - Friday, 17.00

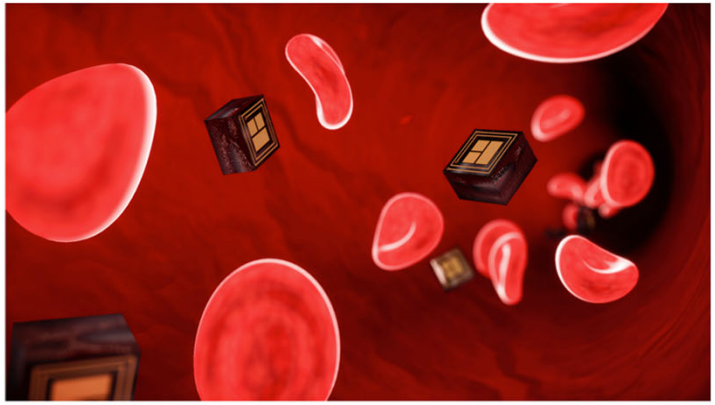
3. Oral presentation
Tuesday, 16th of December

4. A final report (following the IEEE template provided)
Deadline: 31st of December

- Bio
- Nano
- CMOS

Project Proposal





Point-of-care

Wearable

Implantable

Project Proposal

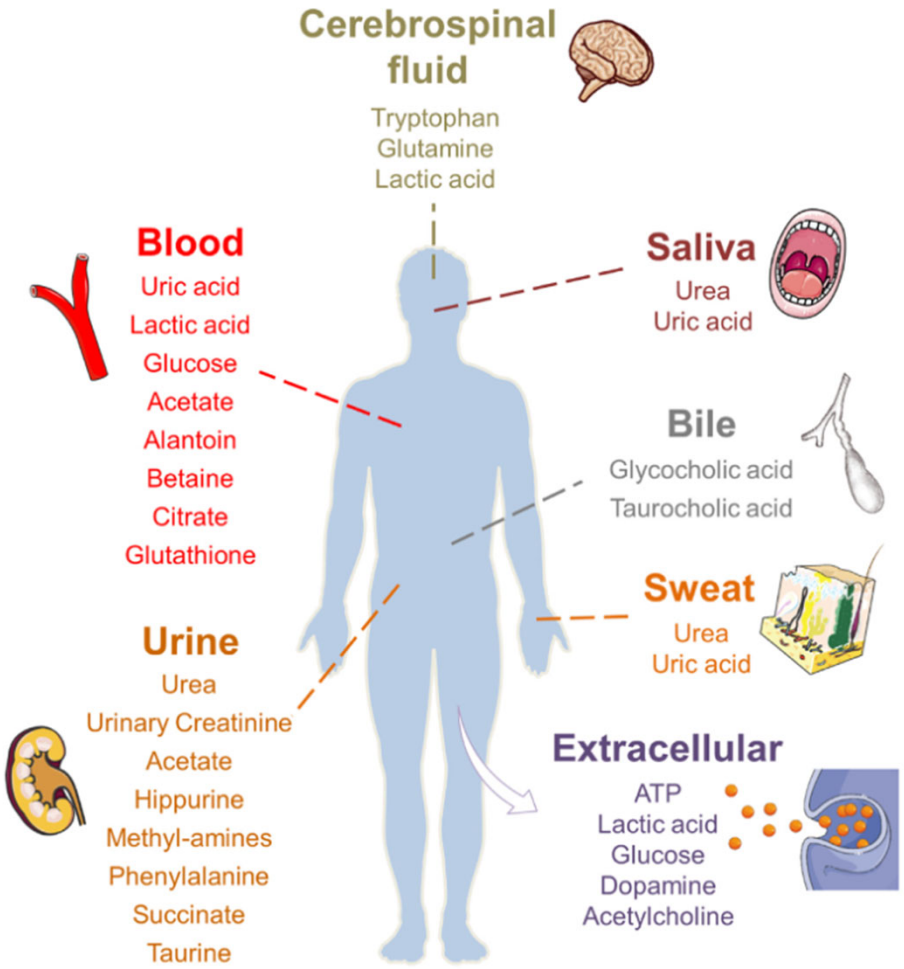


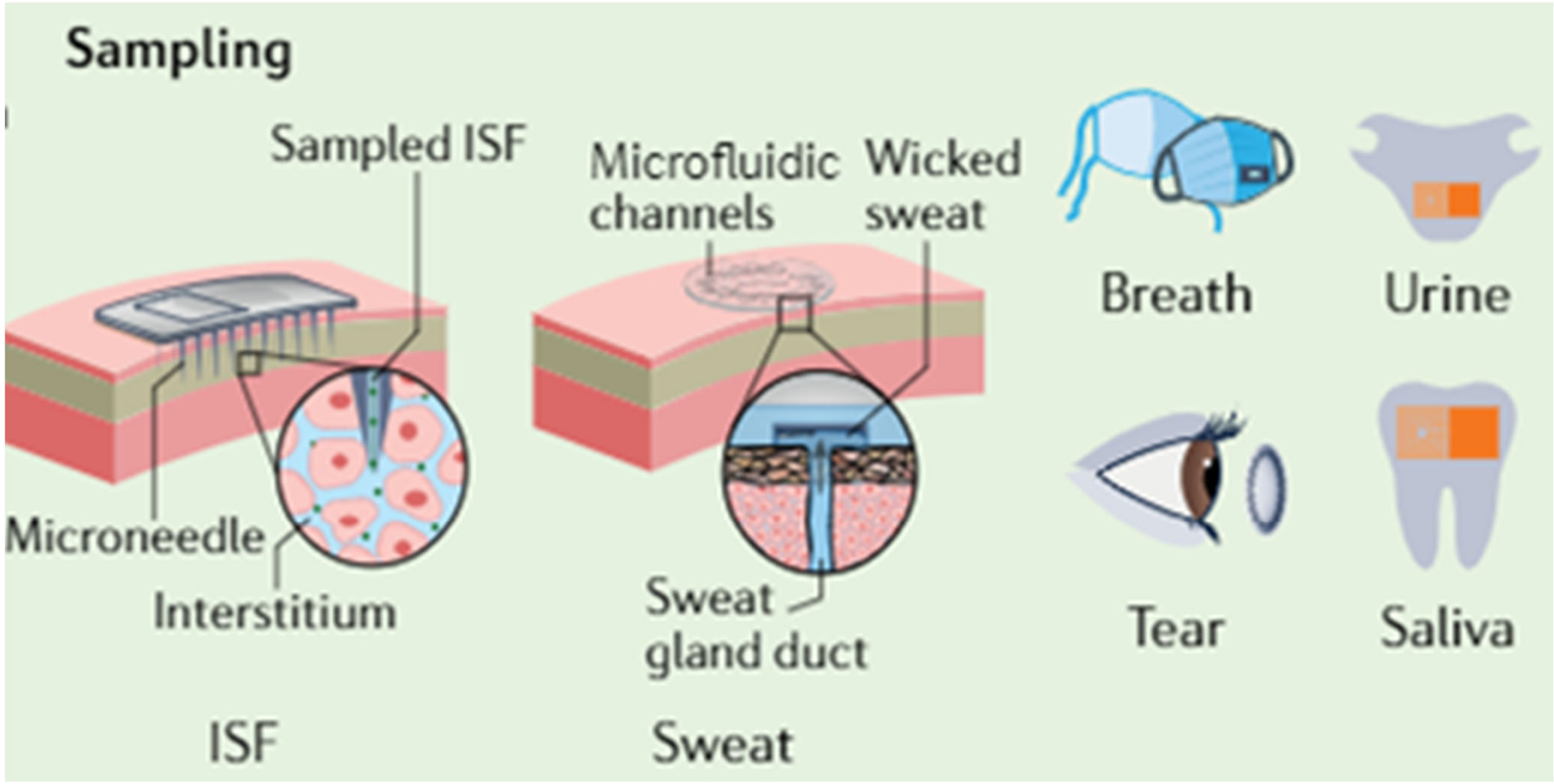
Fig. 1. Common metabolites from biofluids and cells.

Example

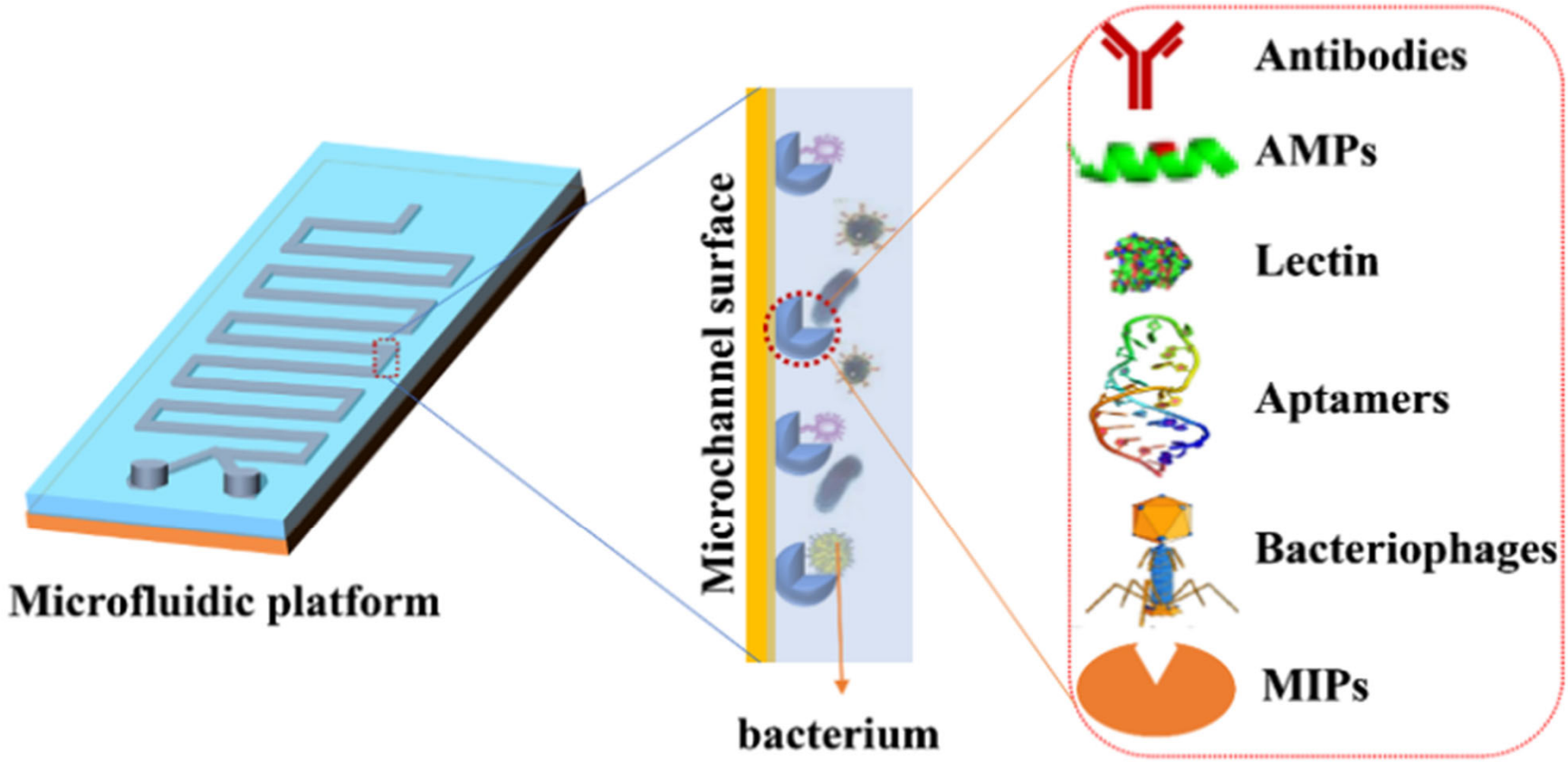
Urea:
Blood between 2.5 and 7.1 mmol/l
Sweat between 5 and 50 mmol/l
Saliva between 0.1 and 2 mmol/l
Urine: daily excretion of 342 ± 67 mmol in 490 to 2690 mL

Lu, Y., Lin, L. and Ye, J., 2022. Human metabolite detection by surface-enhanced Raman spectroscopy. *Materials Today Bio*, p.100205.

Project Proposal



Project Proposal



Mi, F., Hu, C., Wang, Y., Wang, L., Peng, F., Geng, P. and Guan, M., 2022. Recent advancements in microfluidic chip biosensor detection of foodborne pathogenic bacteria: a review. Analytical and Bioanalytical Chemistry, pp.1-20.

Project Proposal

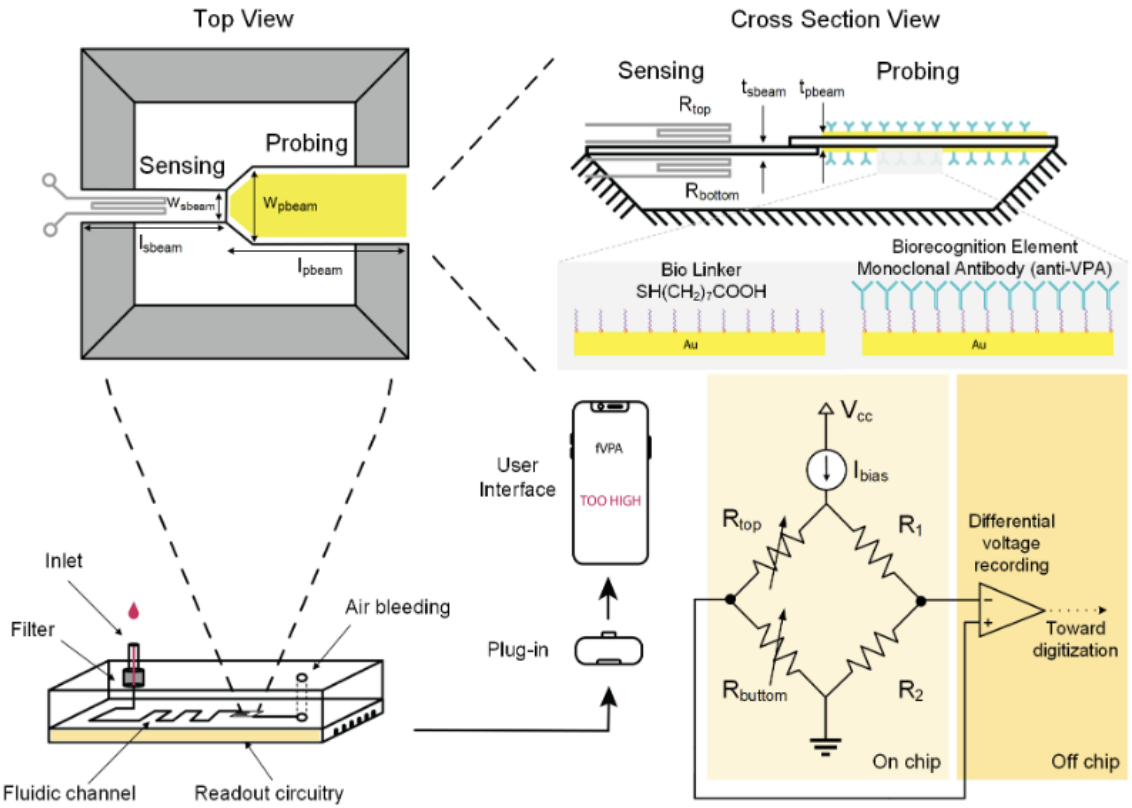


Fig. 1: On button left, the operational principle of the proposed free Valproic acid (fVPA) microcantilever biosensor in depicted. On the top left, the top view of the bioMEMS device, including the gold probing beam and the polyimide sensing beam is depicted. On top right, cross-section view of the biosensor highlighting the top and bottom serpentine gauges (R_{top} and R_{bot}) operating on the sensing beam and self-assembled monolayer (SAM) bio linker interacting with the bio-recognition antibodies in the probing beam acquired for selective biosensing measurement. On button right, the simplified block diagram of the readout system, including the on-chip biased half-Wheatstone bridge (yellow box) and the off-chip readout circuitry, including the amplification stage, the low-pass (LP) filter, and the analog to digital converter (ADC).

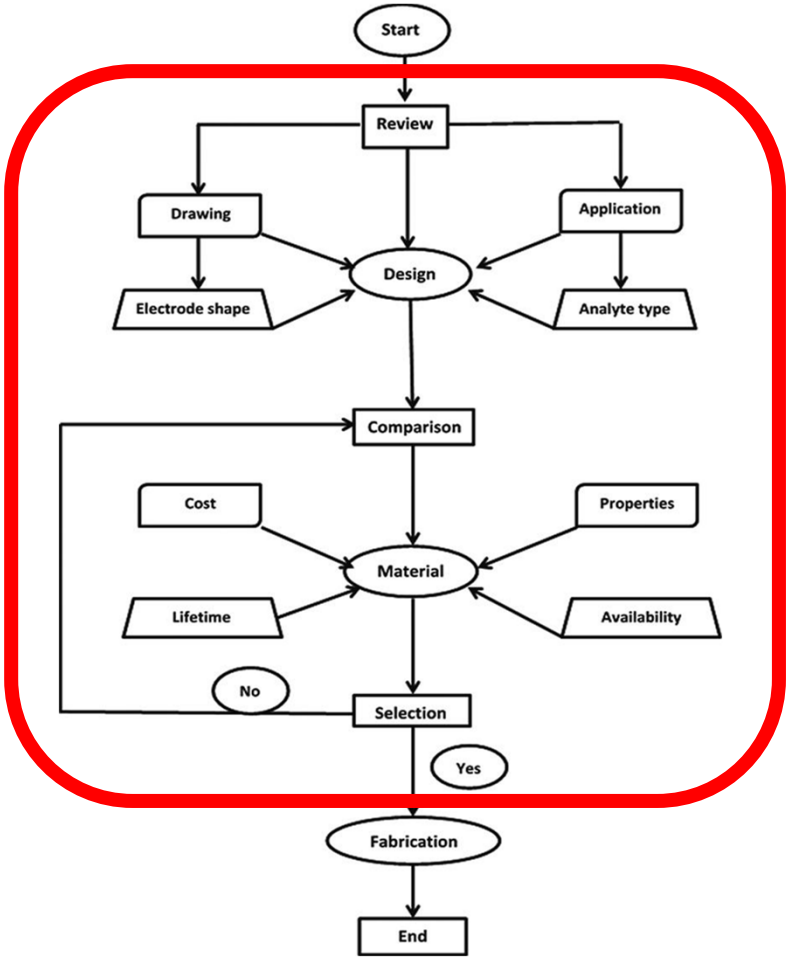
What is innovation? It's all about efforts.

1.Invent a fork, the tool is more hygiene and convenient than hands.

2.Use fork to brush hair is better than using just fingers

3. Find out a 2-tooth fork is particularly good for stabbing bread and dip it into fondu.

Project Proposal



Abdulbari, Hayder A., and Esmail AM Basheer. "Electrochemical biosensors: electrode development, materials, design, and fabrication." *ChemBioEng Reviews* 4.2 (2017): 92-105.

Project Proposal - Example

Portable Breathalyzer for Exhaled Volatile Organic Compounds Monitoring in Lung Diseases

Alec Chevrot, Justina Venckute, Sarah Cuesta, Ata Golparvar, Amar Kapic, Gian Luca Barbruni*, Sandro Carrara
Integrated Circuit Laboratory (ICLAB), École Polytechnique Fédérale de Lausanne (EPFL), Neuchâtel, Switzerland

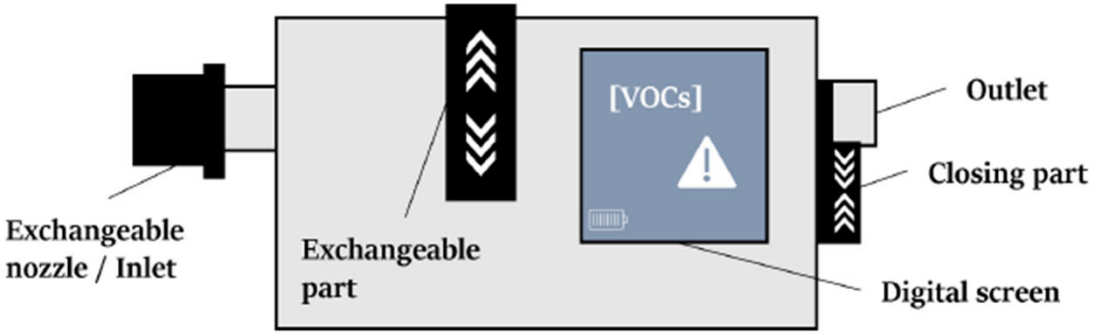


Fig. 1. The overall scheme of the proposed portable breathalyzer includes the exchangeable nozzle to enable sanitary operation by multiple users, an interchangeable sensor compartment for quick adjustment to enable different biomarker detection, and a segment display to ensure low-power operation when displaying the readings.

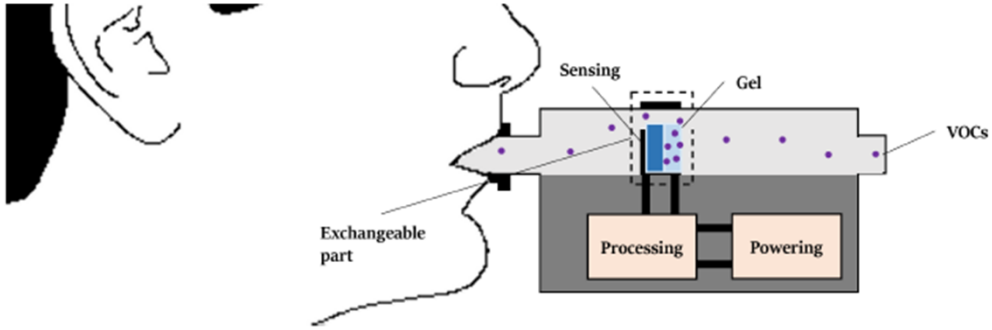


Fig. 2. The overall structure of the handheld breathalyzer during a typical application scenario. VOC are directly exhaled into the device. The flow is directed onto the exchangeable sensor compartment composed of agarose gel to collect VOC and an aptamer-based bio-nano interface to selectively detect the particular VOC of interest.

Projects Proposal - Hints

- Find idea based on personal interests/motivation
- Check recent review paper about the topic
- Search papers by keywords
- Check paper relevance: abstract/conclusions/tables
- Check references of the paper: might be useful!
- Tools: Google Scholar/IEEE Xplore/Nature

Project Proposal – Template

Name and Surname:
Section:

Previous Experience Checklist (put a check mark if applicable):

- Transistor-level Bi/CMOS circuit design
- PCB-level circuit design
- SPICE simulation software use (e.g., LTspice)
- Multiphysics finite element analysis software use (e.g., COMSOL)
- Microfluidics (design or simulation or fabrication)
- Development/simulation of a sensor (if yes, which one?):

Title of the project (provisional):

Short description/summary (250 words max):

Sections you want to implement (1 sentence description for each):

- Bio:
- Nano:
- CMOS:
- Other (if applicable):

More Details:

- Application (250 words max):
- Motivation (250 words max):
- Problems to be solved (250 words max):
- Proposed solutions (250 words max):
- How will you implement your solution? (250 words max):
- Novelties of your proposal (Bullet points, each point 1 sentence):