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Leveraging a better tomorrow

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Tutorial 4

Low-Noise Impedance Sensing: Circuits and Micro- Technological Applications

Marco Carminati



Politecnico di Milano - Dipartimento di Elettronica, Informazione e Bioingegneria

marco1.carminati@polimi.it

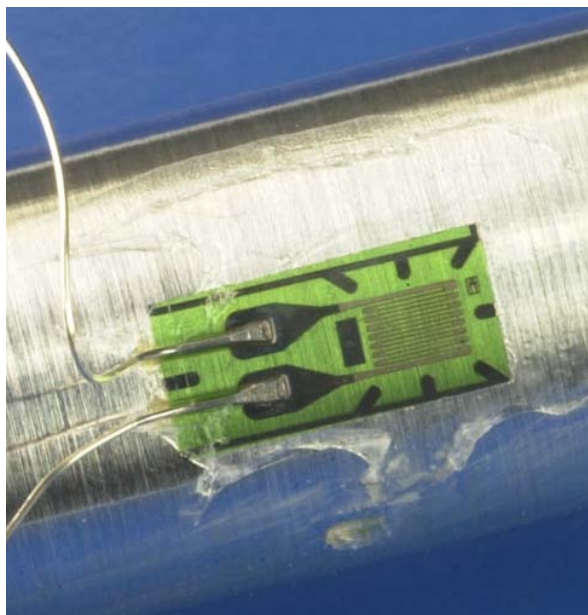


Motivation: Impedance is Everywhere

Thanks to its versatility, impedance is traditionally leveraged in **several industrial sensing applications**

Strain/Pressure

Resistance



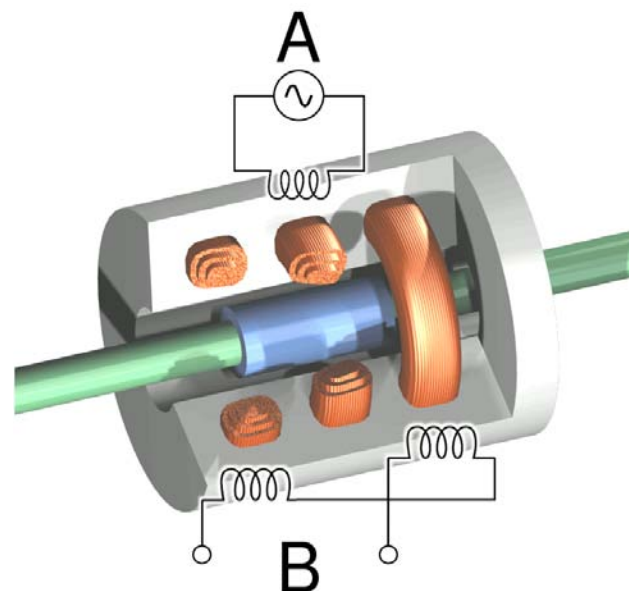
Level

Capacitance



Displacement

Inductance



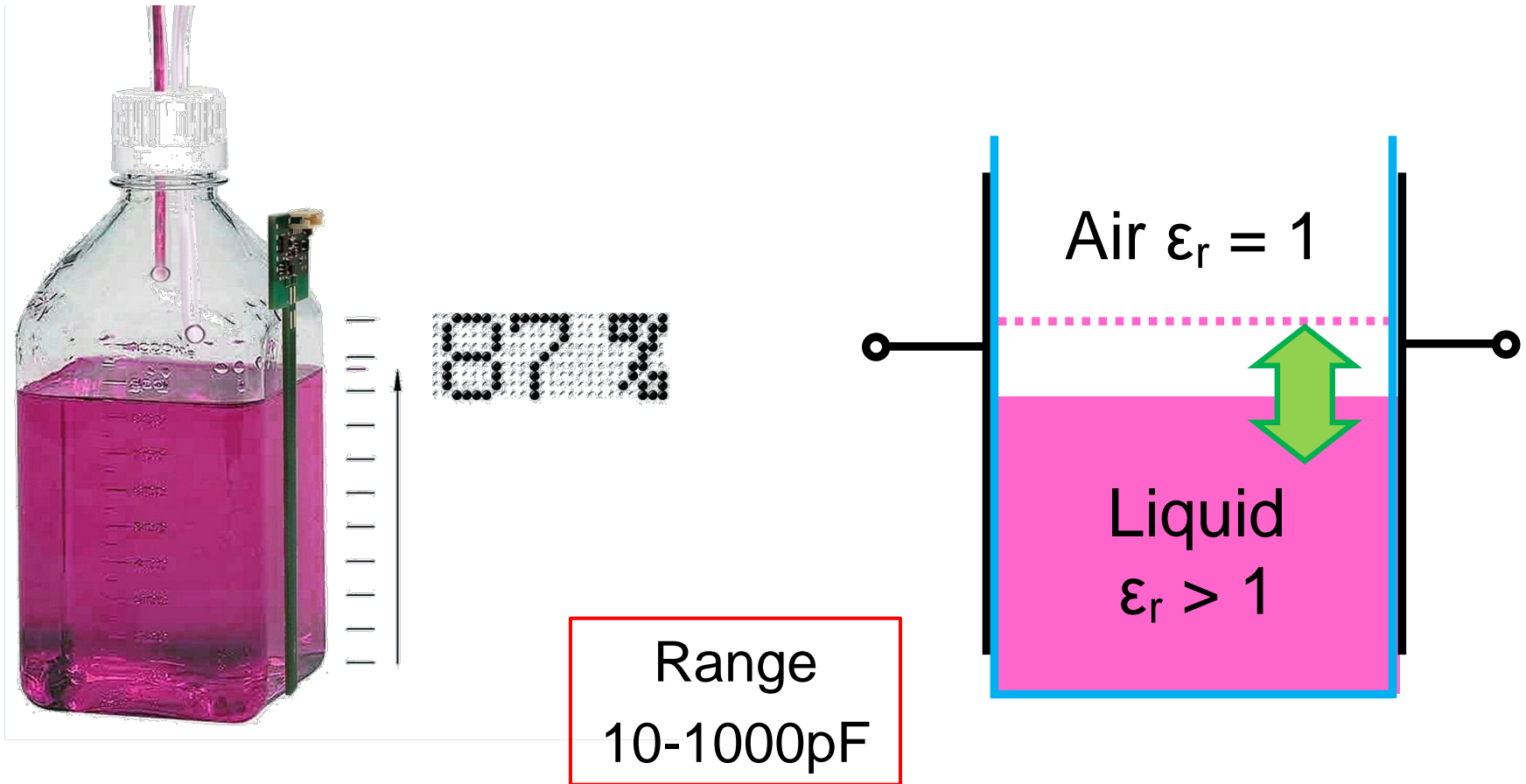


Focusing on the example of
contactless capacitive sensing,
let's see the scaling trend
from macro to micro size
applications



Macro: Contactless Liquid Sensing

Electrodes are protected from the liquid, the capacitance increases linearly with the tank level





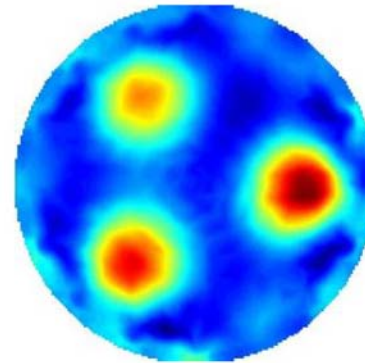
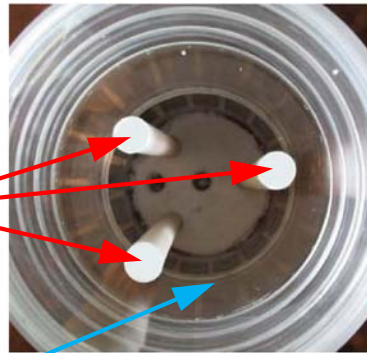
Macro: Contactless Capacitance Tomography

Impurities inside pipes (or droplets in microfluidic channels) can be detected with several electrodes properly arranged

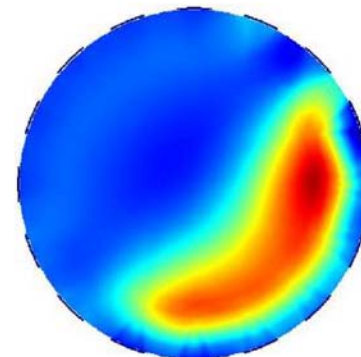
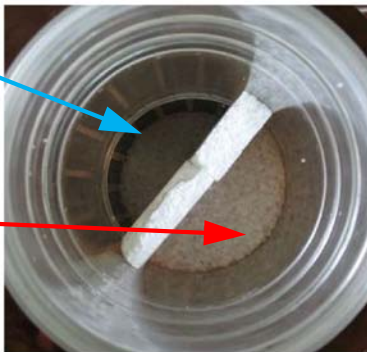
Test structures

Obtained Maps

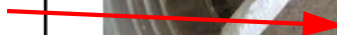
Plastic bars
 $\epsilon_r = 3$



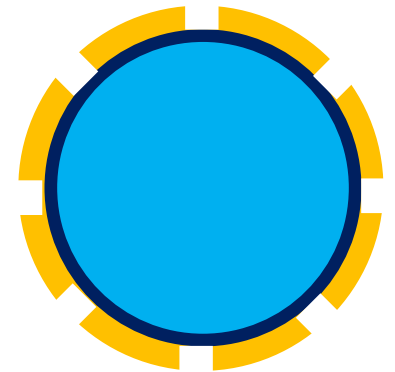
Air
 $\epsilon_r = 1$



Plastic beads
 $\epsilon_r = 3$

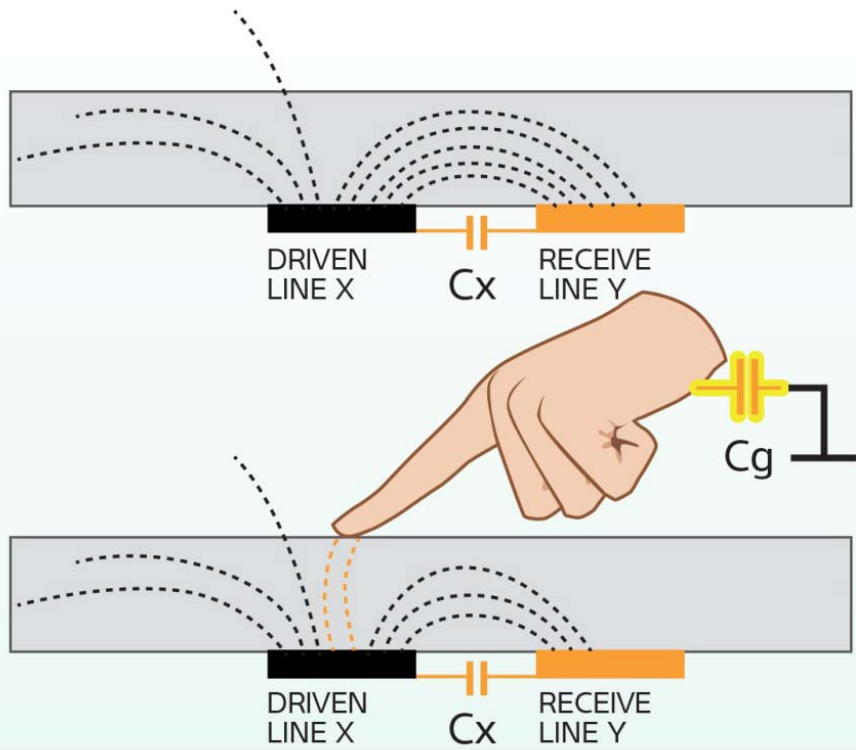


Pipe cross-section

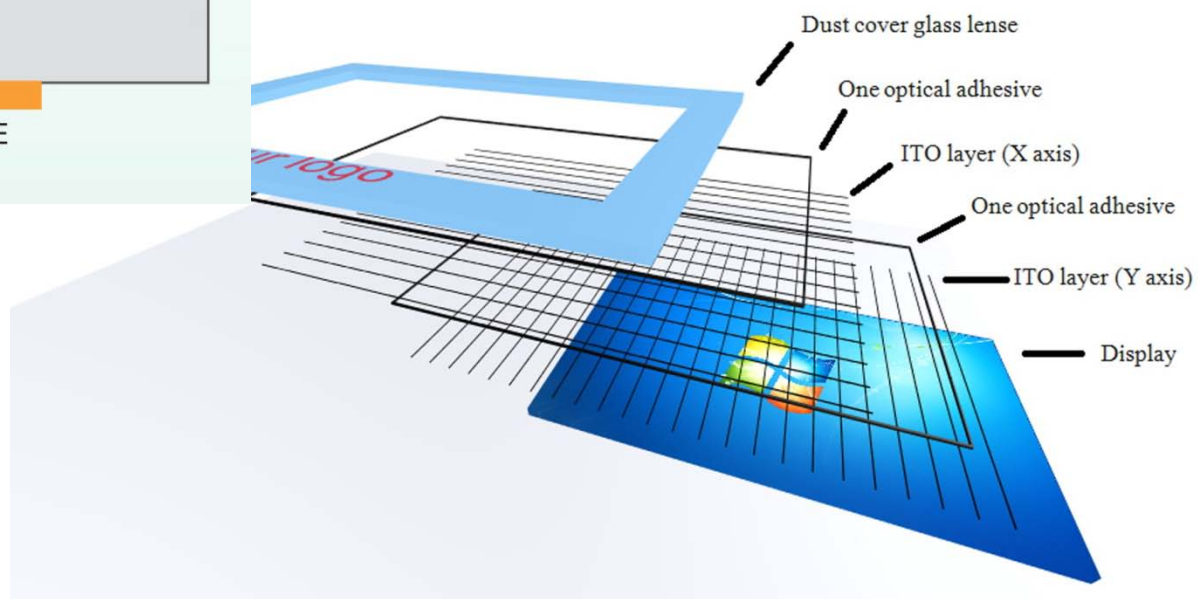


Range
10-100pF

Mini: Capacitive Touchscreen



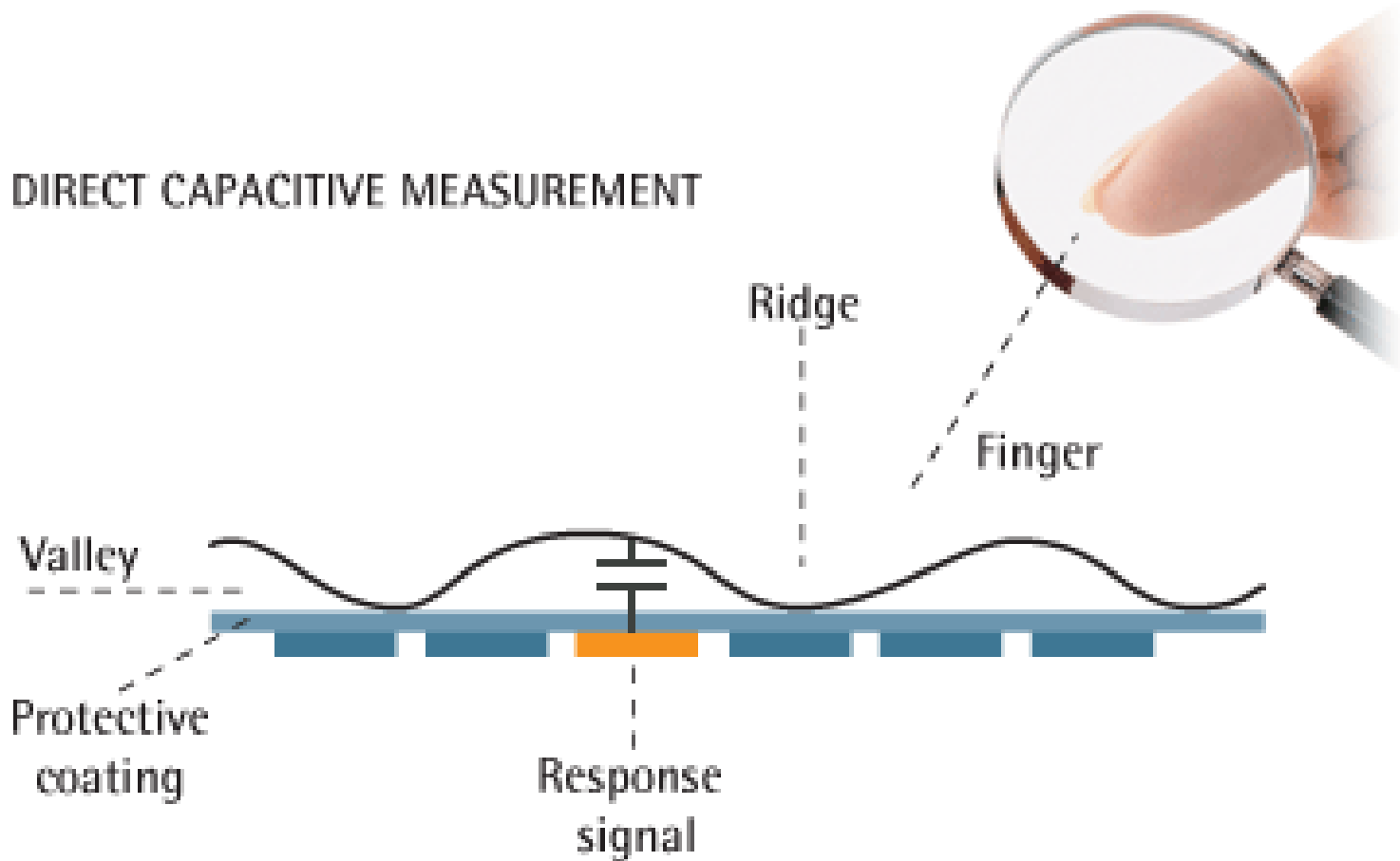
Range
10-100fF





Milli: Capacitive Fingerprint Scanner

DIRECT CAPACITIVE MEASUREMENT

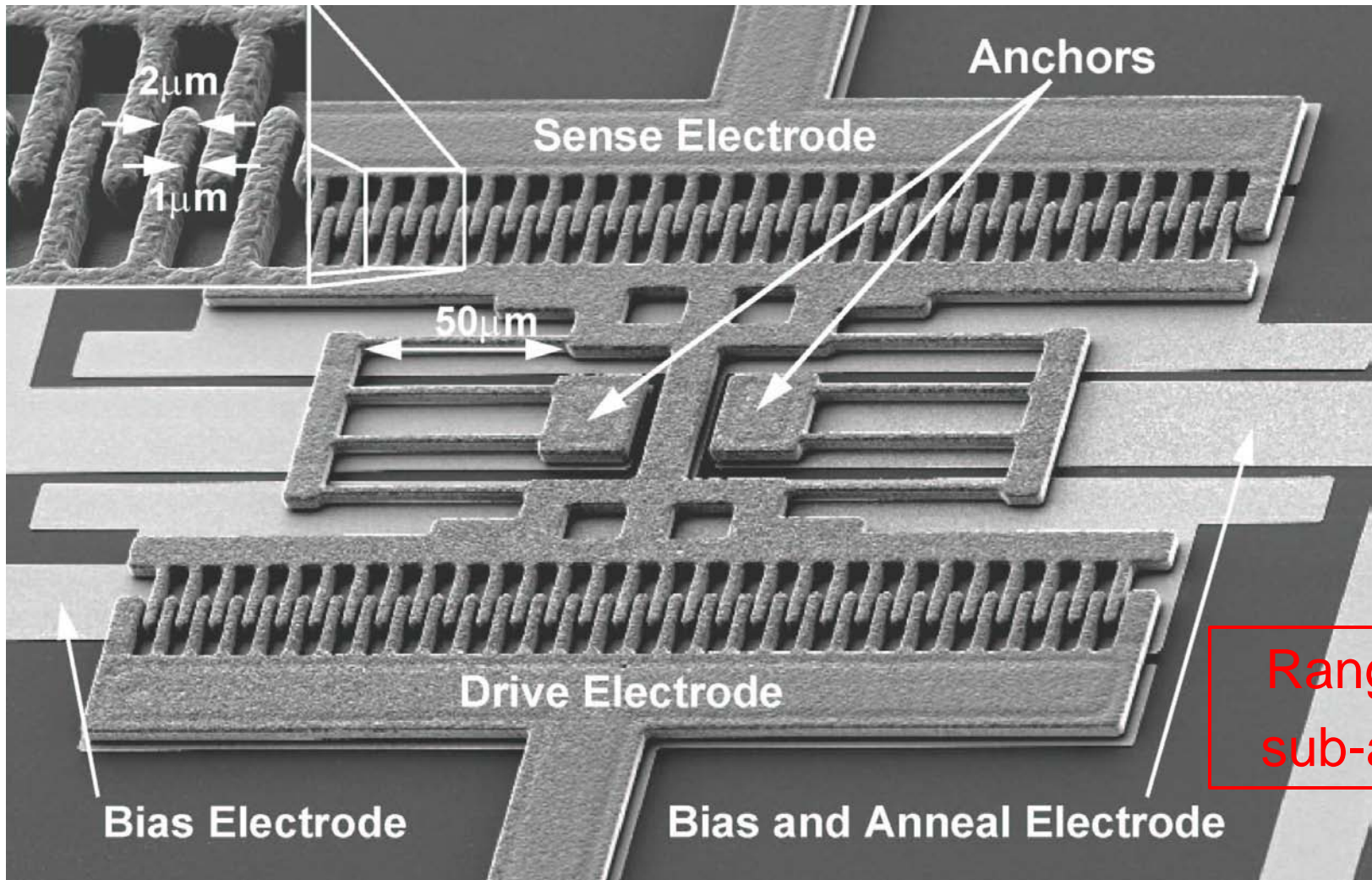


Range
5-50fF



Micro: Inertial MEMS

Capacitive sensing of displacement of micro proof masses

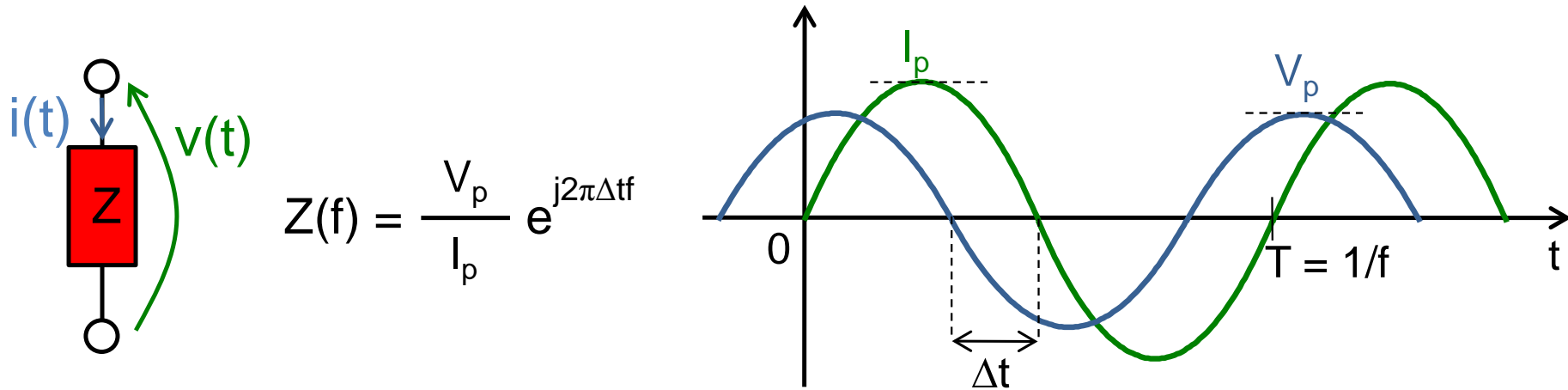




Tutorial Outline

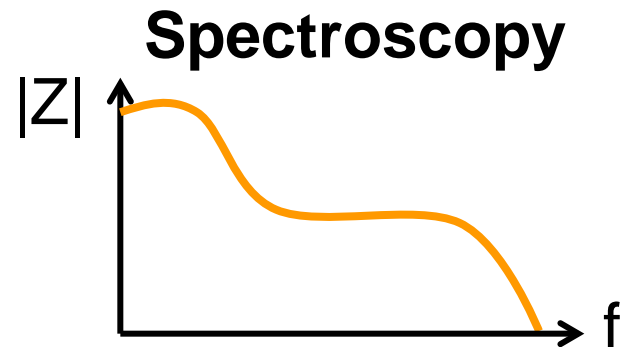
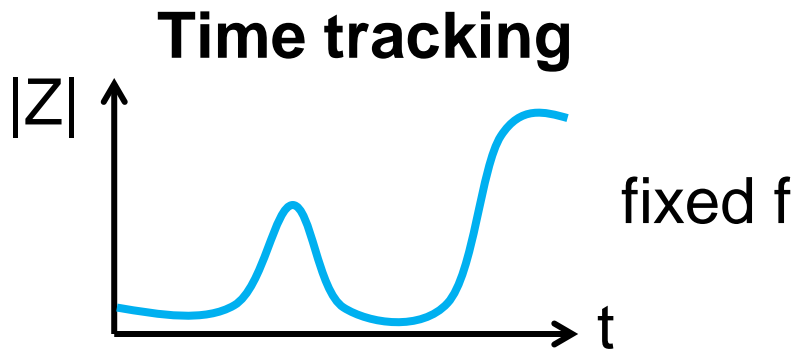
- Definitions and spectra diagrams
- Microscale impedance measurement
- Introduction to electronic noise
- Low-noise current sensing
- Transimpedance amplifier and lock-in
- Role of the input (stray) capacitance: cables and substrates
- Advanced circuit topologies
- **Example of applications:**
 - Biological cell counting
 - Particulate matter detection
 - Non-invasive light power metering

Impedance is a complex quantity changing with frequency f :



$\text{Re}\{Z\}$: energy dissipation, $\text{Im}\{Z\}$: energy storage

Impedance can be measured:

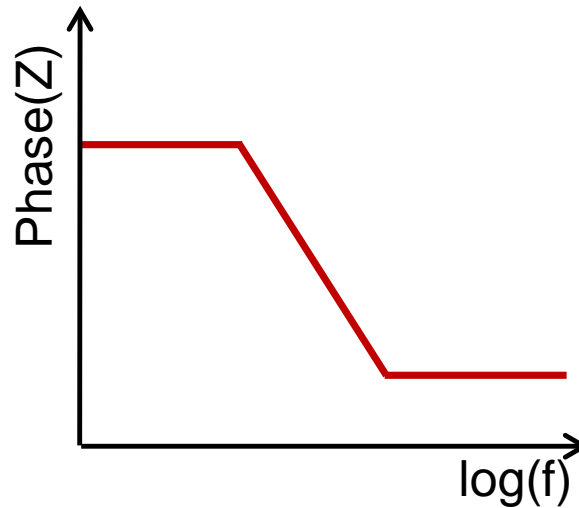
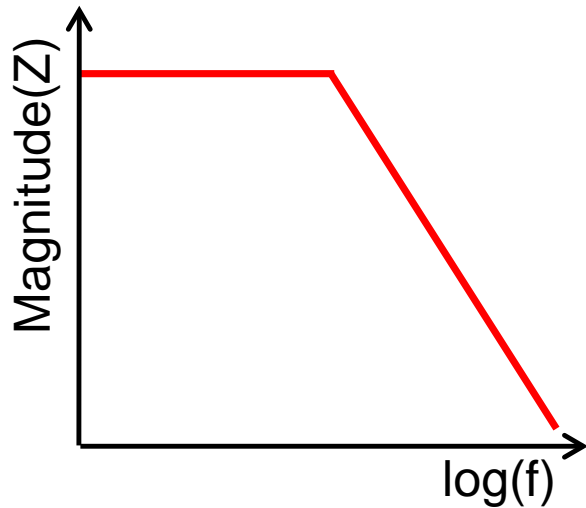




Plotting Impedance Spectra

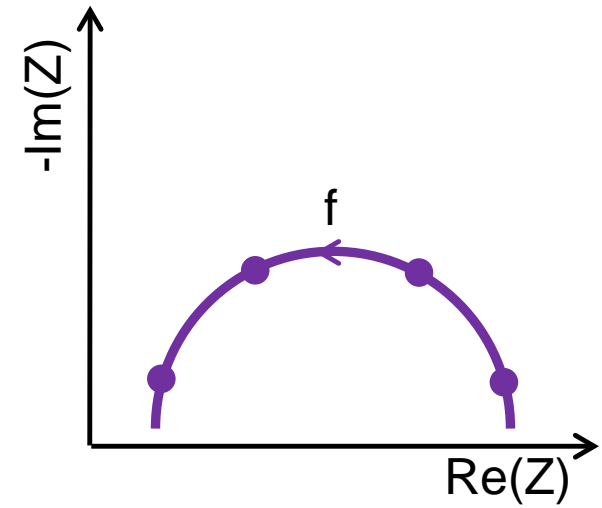
Two alternative ways to present impedance spectra:

Bode Plots



Electronic engineering and
automation/control science

Cole-Cole Plot

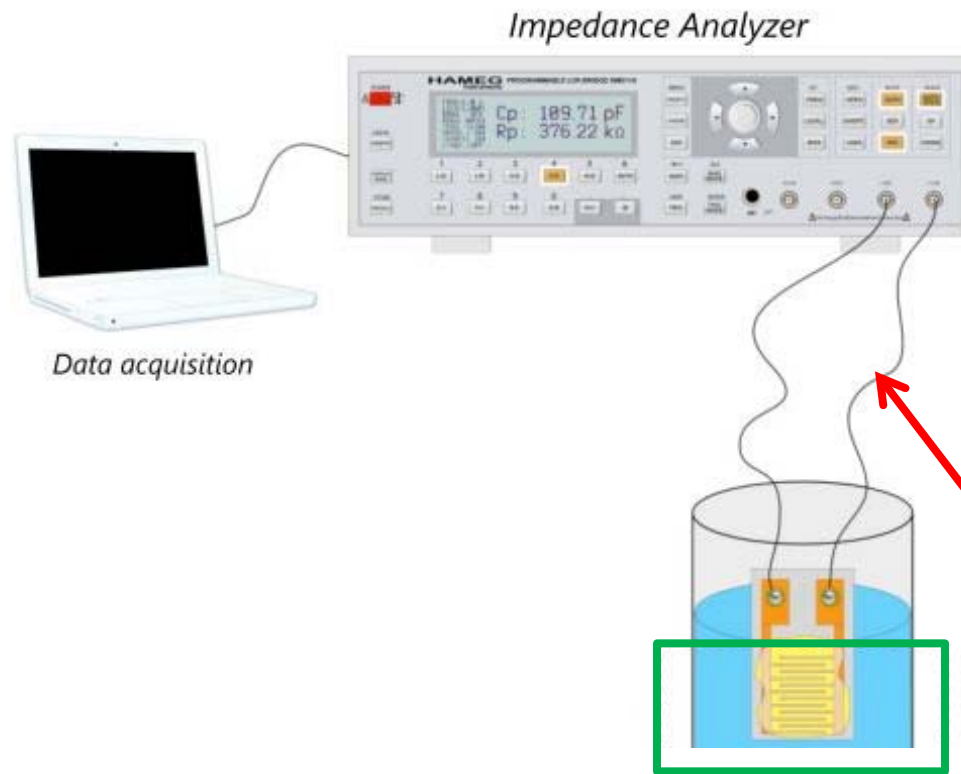


Electrochemical
& biosensors
community

Equivalent!

Microscale Impedance Sensing

Microfabrication allows the realization of **microelectrodes** to interface with microscale samples (such as biological cells).



The impedance of the micro sample **scales** with the size

while

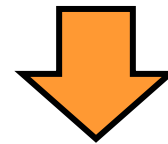
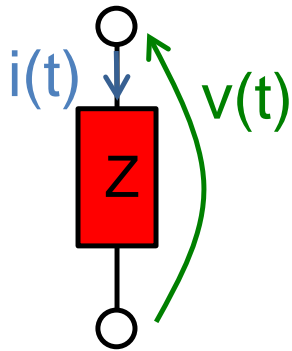
the parasitic impedance of the macroscopic **connections** stays fixed.



Noise Becomes Important

If Z increases, given the same excitation **voltage** (limited by biology or *dielectric breakdown*), the **current** flowing through the system **decreases**:

It becomes harder to measure a smaller current



The noise of the instrument (current reader) becomes significant!

The noise is a property of the detection instrument but might depend on the sample and on the connections.



What is Noise?

Any electrical signal is affected by **disturbances** and **noise**:

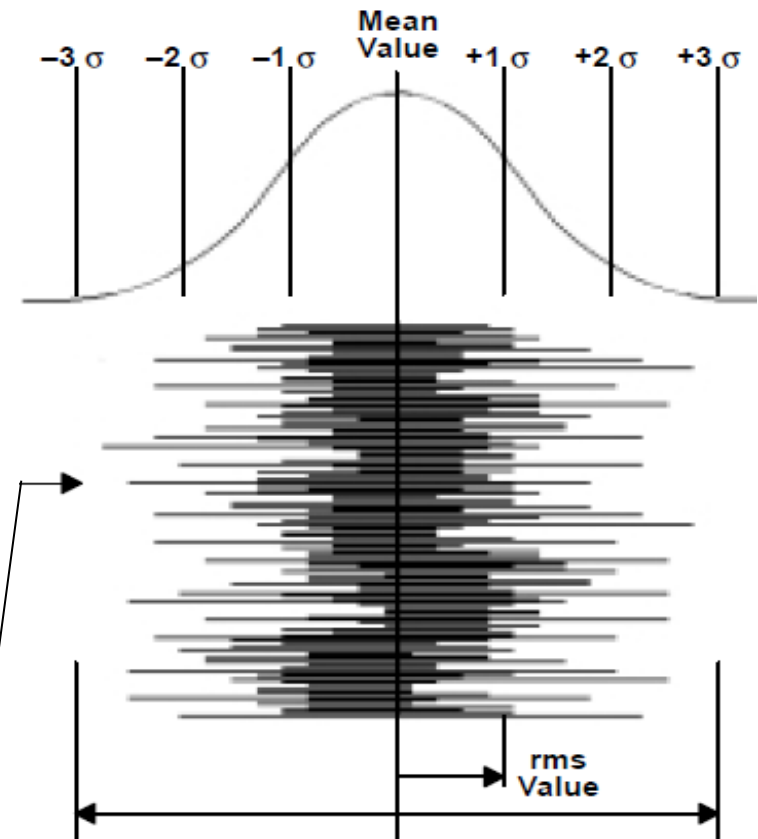
$$i(t) = \text{signal}(t) + d(t) + n(t)$$

Disturbances are signals of external origin that can be filtered.

Noise is a **random fluctuation** of the electrical variables due to the physical behaviour of *internal* components of the circuit.

Described by standard deviation σ

$$SNR = \frac{\text{Signal Amplitude}}{\text{Noise Amplitude}} = \frac{\text{Signal}}{\sigma_{rms}}$$





Noise Power Spectrum

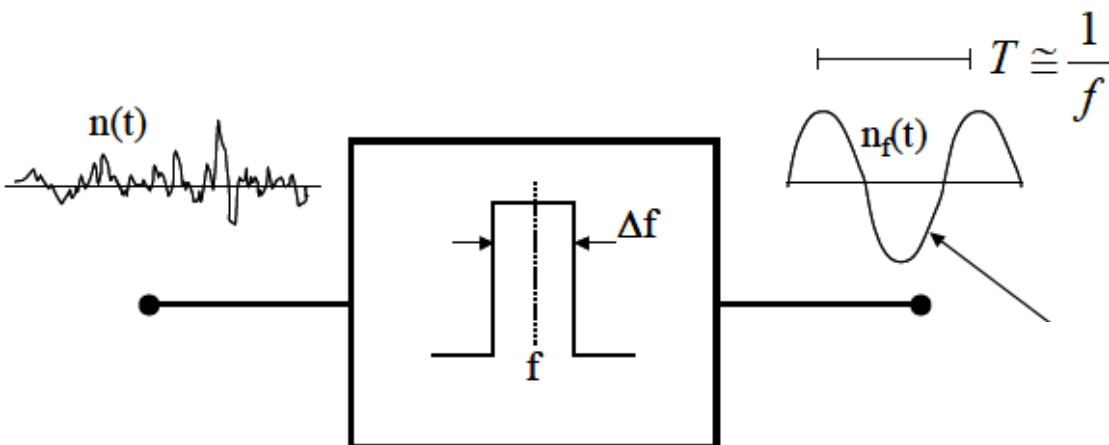
Description in the frequency domain:

Noise spectral density

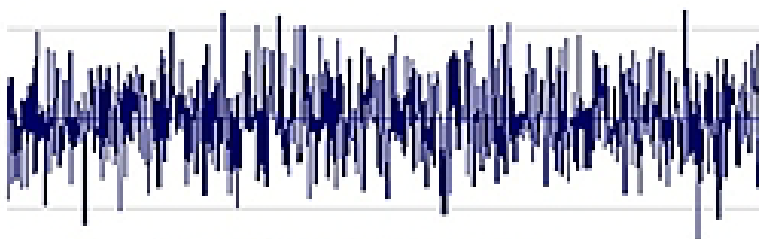
$$S(f) = \overline{n_f^2(t)} / \Delta f$$

$$\overline{n^2(t)} = \int_0^\infty S(f) df$$

$$\text{RMS} = \sqrt{\overline{n^2(t)}} = \sqrt{\int_0^\infty S(f) df}$$



Flat-band or White Noise



Time Domain Waveform



Spectral Density

1/f flicker noise





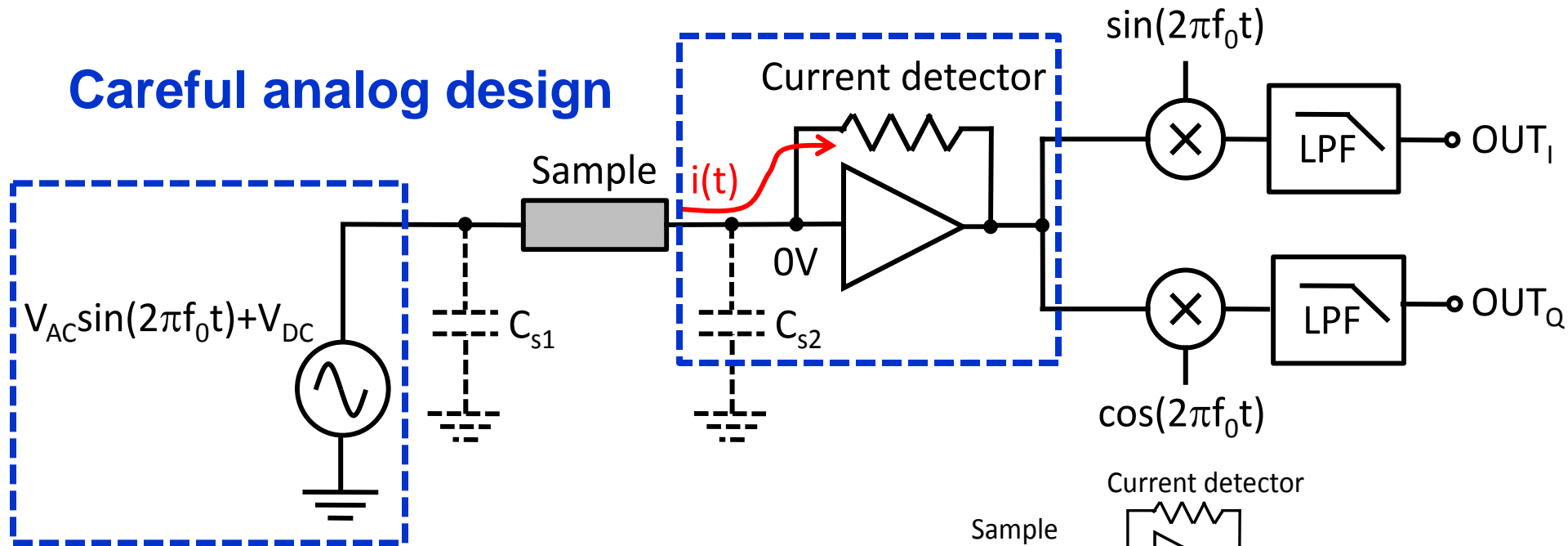
Impedance Measurement Techniques

- **Sinusoidal stimulation:** **(Measurand)**
 - Wheatstone bridge (voltage)
 - Ratiometric (voltage)
 - Resonant (frequency)
 - **Current sensing + lock-in (current)**

most versatile approach, better rejection of parasitics
- **Non sinusoidal stimulation:**
 - Multi-sine or pseudo noise FFT-based (current)
 - Time-domain fitting of step response (current/voltage)
 - Capacitance charge and discharge (current/time)

Impedance Detector Architecture

Careful analog design

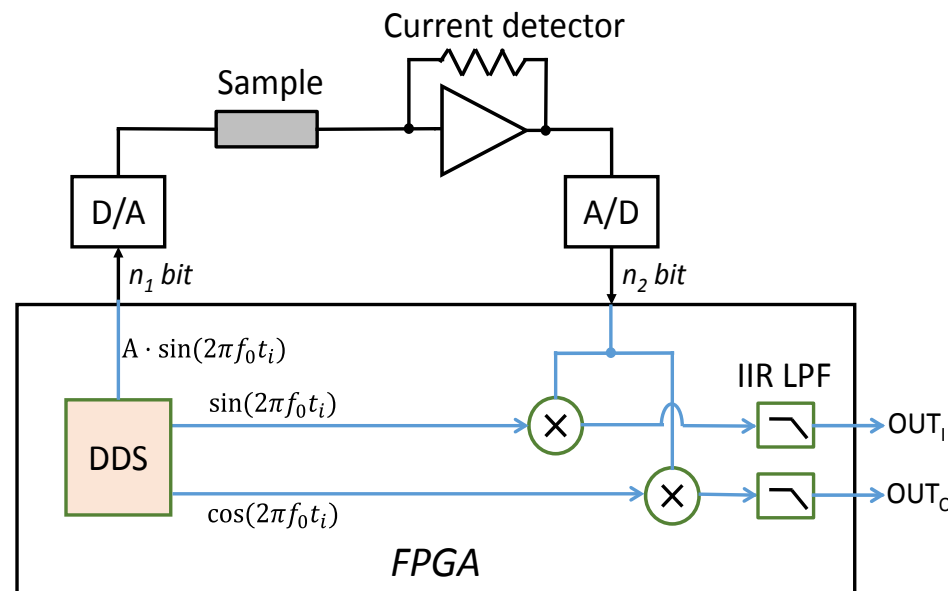


Lock-in digital implementation:

FPGA embedded processing for:

- speed
- versatility (multi-freq.)

M. Carminati, *IEEE I2MTC*. 2012

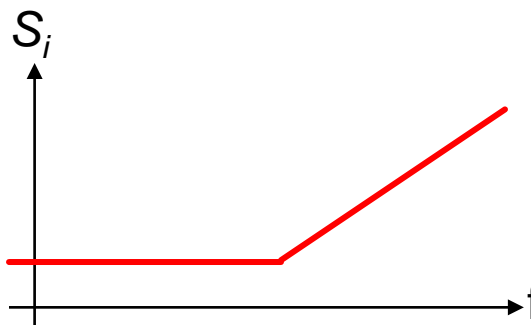
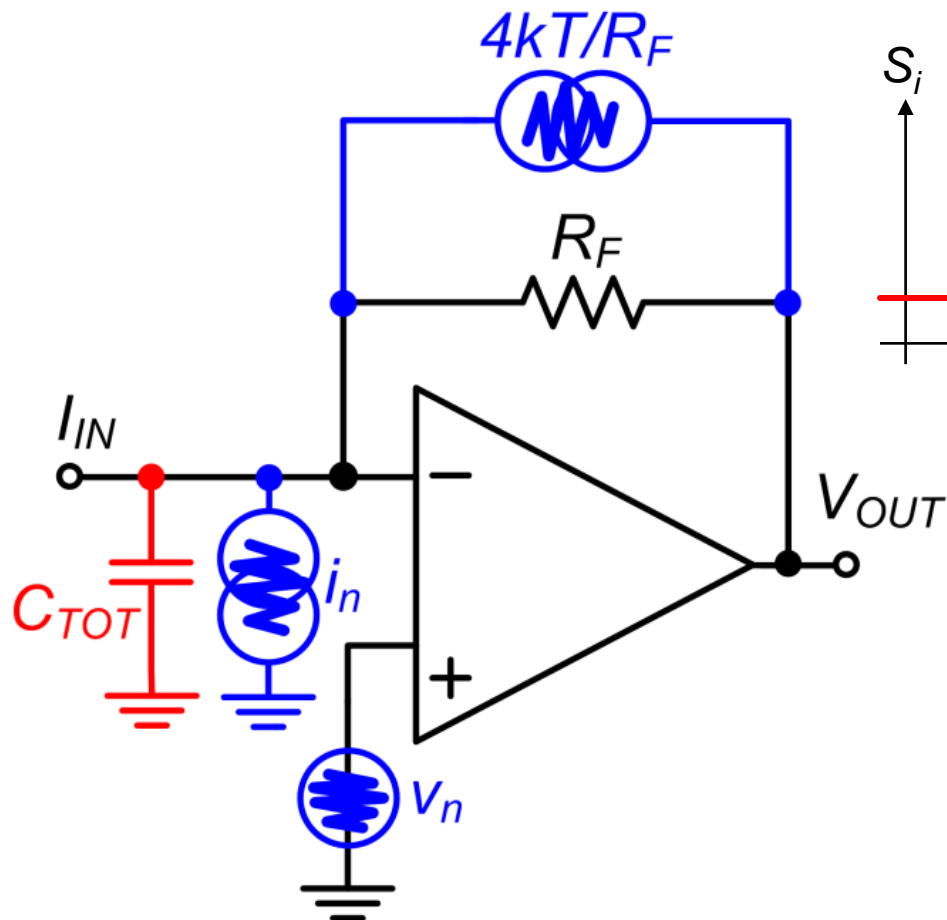




Transimpedance Amplifier: Noise Analysis

Input-referred total current noise:

$$S_i^2(f) = i_n^2 + \frac{4kT}{R_F} + \frac{v_n^2}{R_F^2} + v_n^2(2\pi f C_{TOT})^2$$

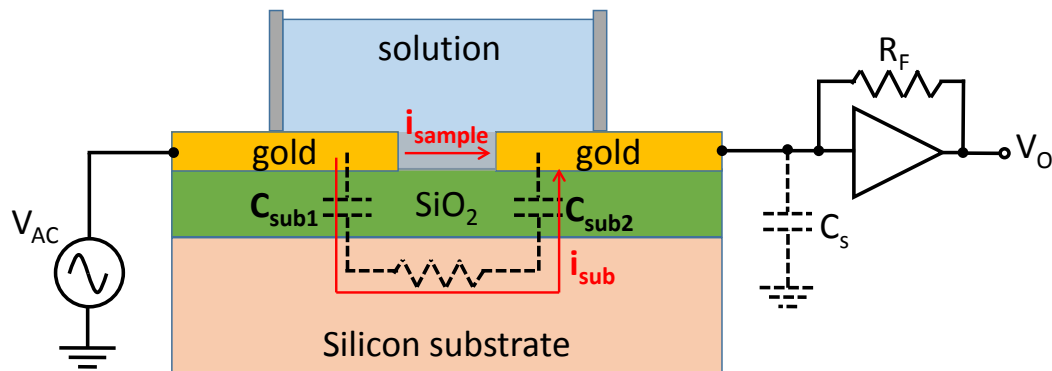
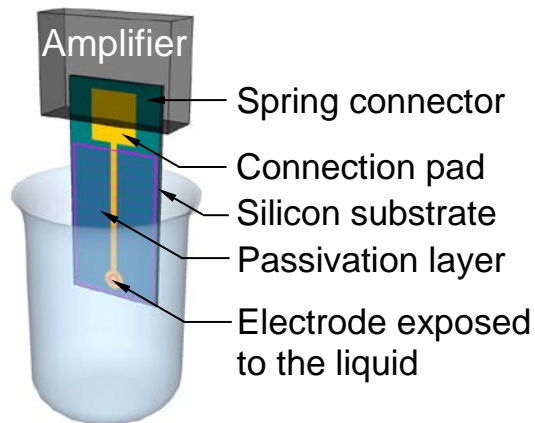


Minimize C_{Stray} !

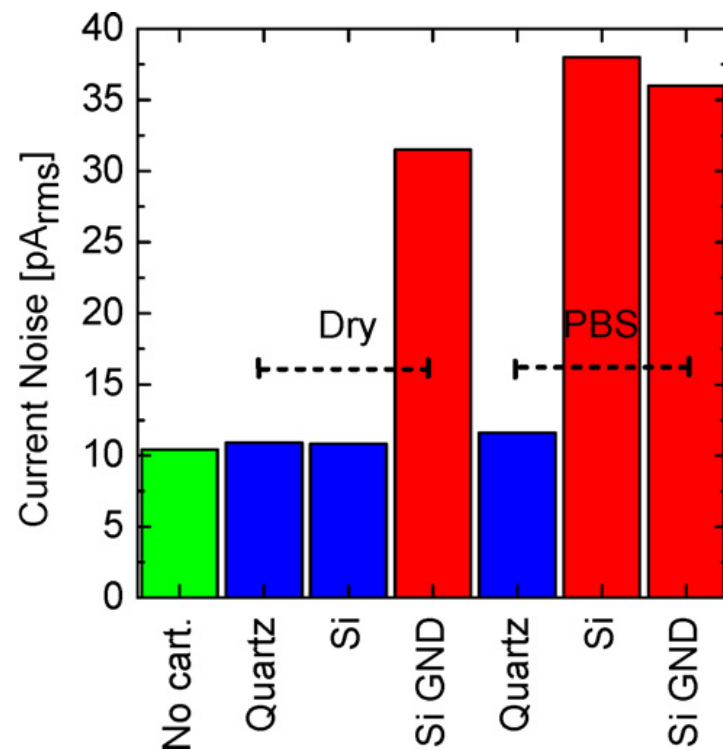
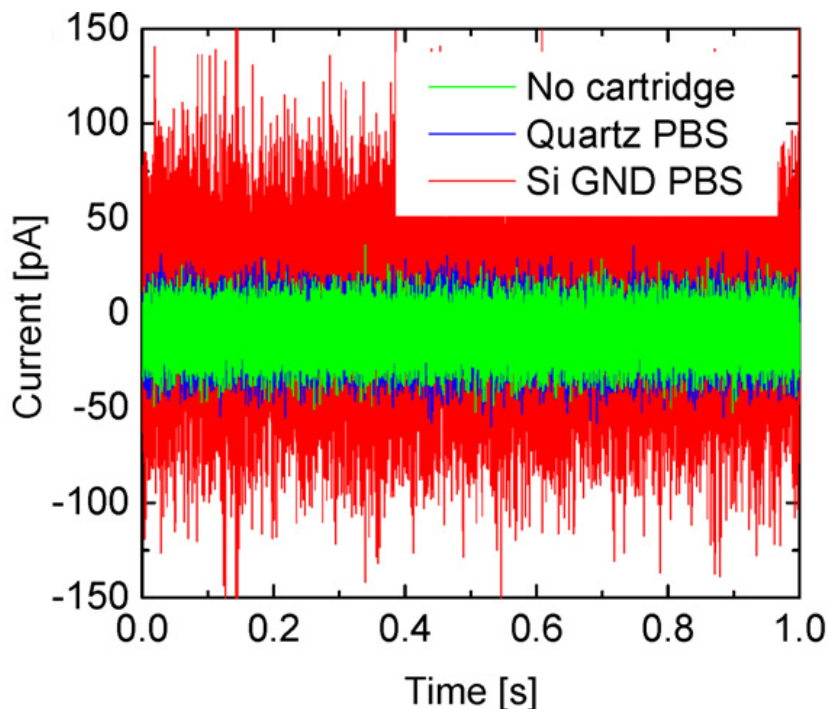
- Minimization of parasitics
- Major motivation for front-end integration
- Accurate modeling of the sensor impedance



Getting Rid of Silicon When Not Needed



Substrate comparison

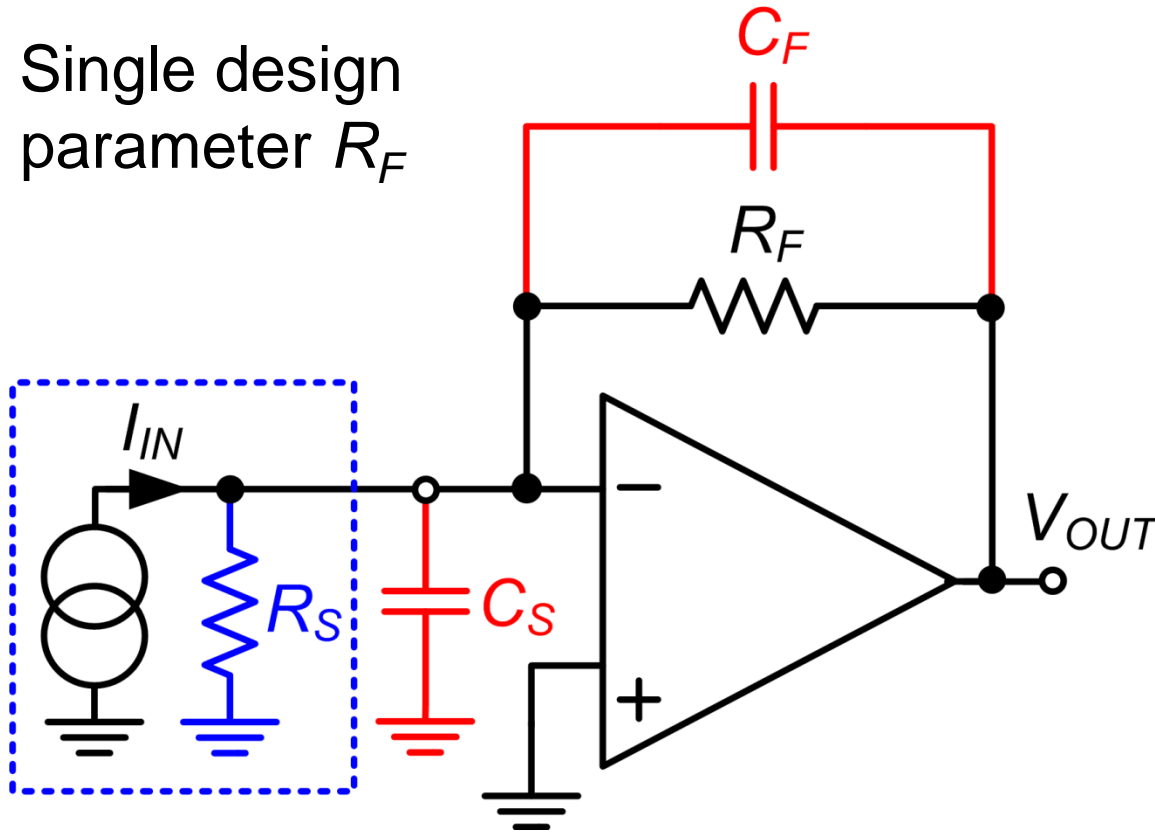


M. Carminati, "Accuracy and resolution limits in quartz and silicon substrates with microelectrodes for electrochemical biosensors", *Sens. Act. B*, **174**, 2012.

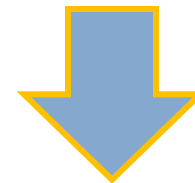


Transimpedance Amplifier: Limits

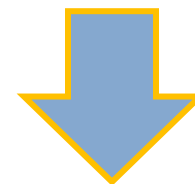
Single design parameter R_F



- gain $\sim R_F$
- noise $\sim \sqrt{4kT/R_F}$
- BW $\sim 1/(R_F C_F)$



Resolution/speed trade-off



Need for enhanced topologies

C stray SMD package

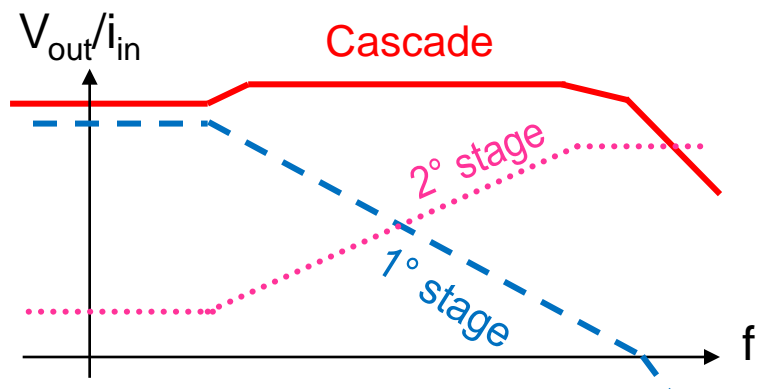
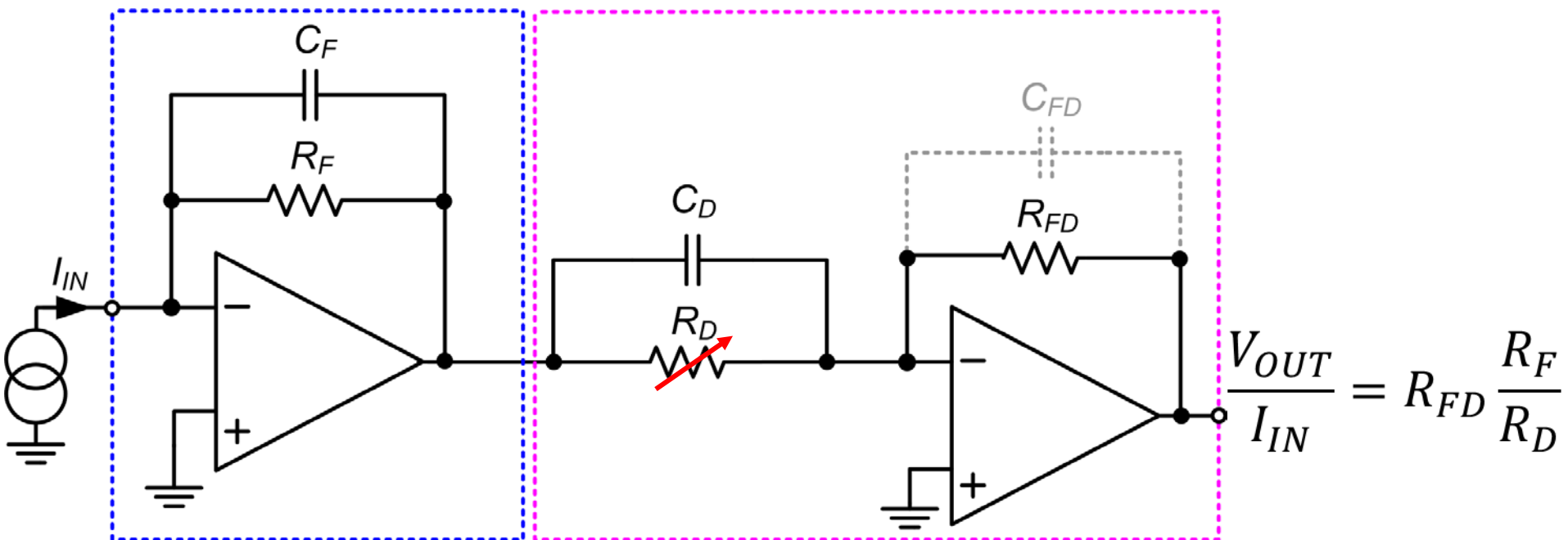


For example: $R_F = 1\text{G}\Omega$ $C_F = 0.2\text{pF}$ $\rightarrow S_i = 4\text{fA}/\sqrt{\text{Hz}}$ BW = 800Hz



1 - Compensated TIA

Introduce a zero that cancels the pole: $R_F \cdot C_F = R_D \cdot C_D$



☺ Wider bandwidth, low noise

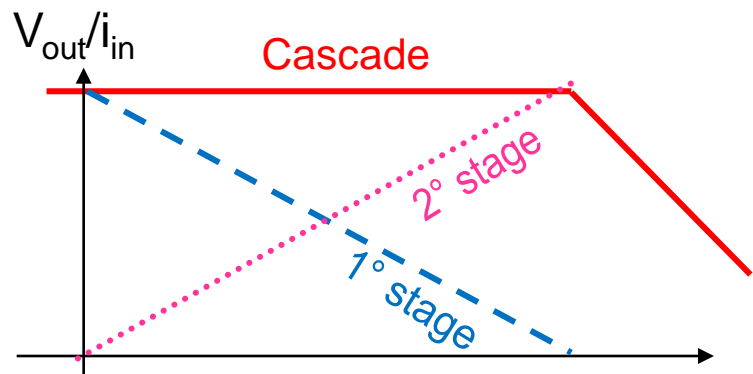
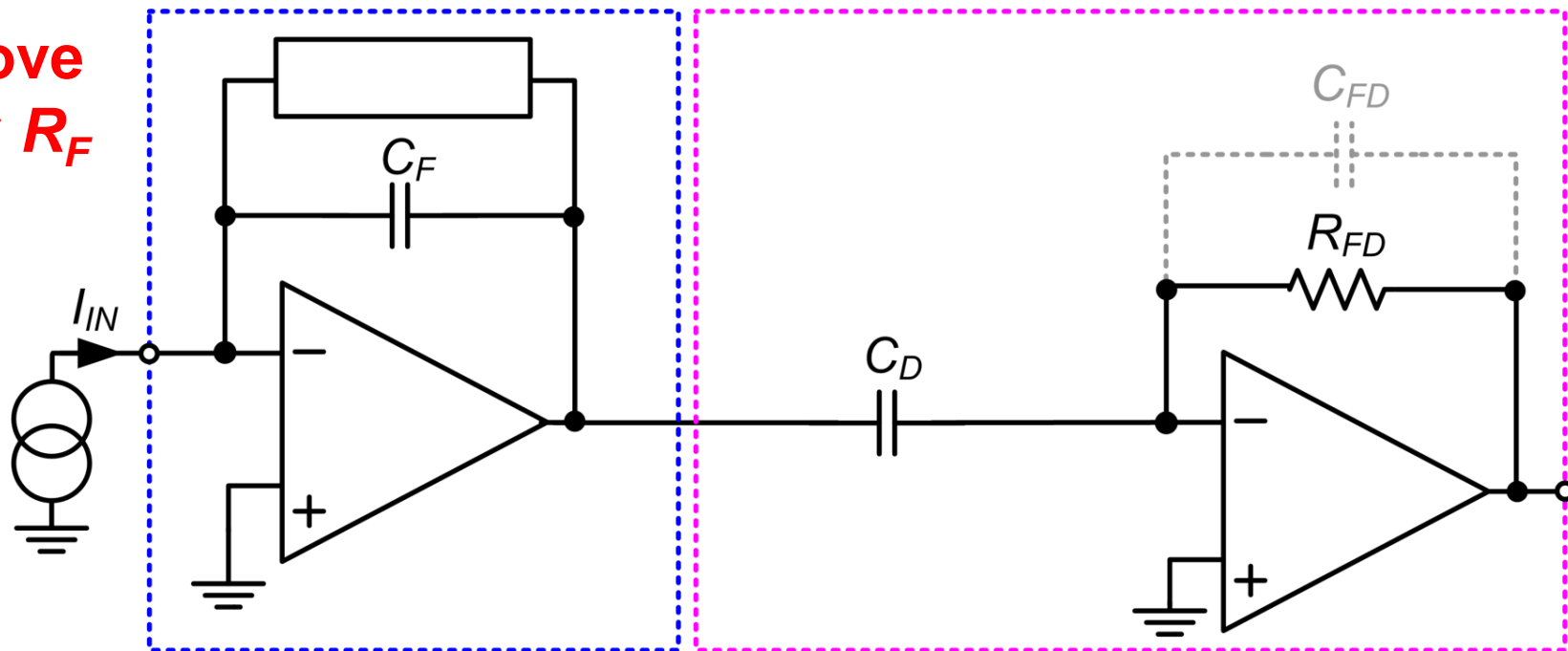
☹ Manual adjusting of the zero

Ciofi, *IEEE Instr. & Meas.* **55** 2006, M. Carminati, *Analog IC Sig. Process.* 2013



2 - Integrator-Differentiator Cascade

Remove
noisy R_F



$$\frac{V_{OUT}}{I_{IN}} = R_{FD} \frac{C_D}{C_F}$$

- 😊 Robust, linear, no calibration
- ☹ Handling input DC current

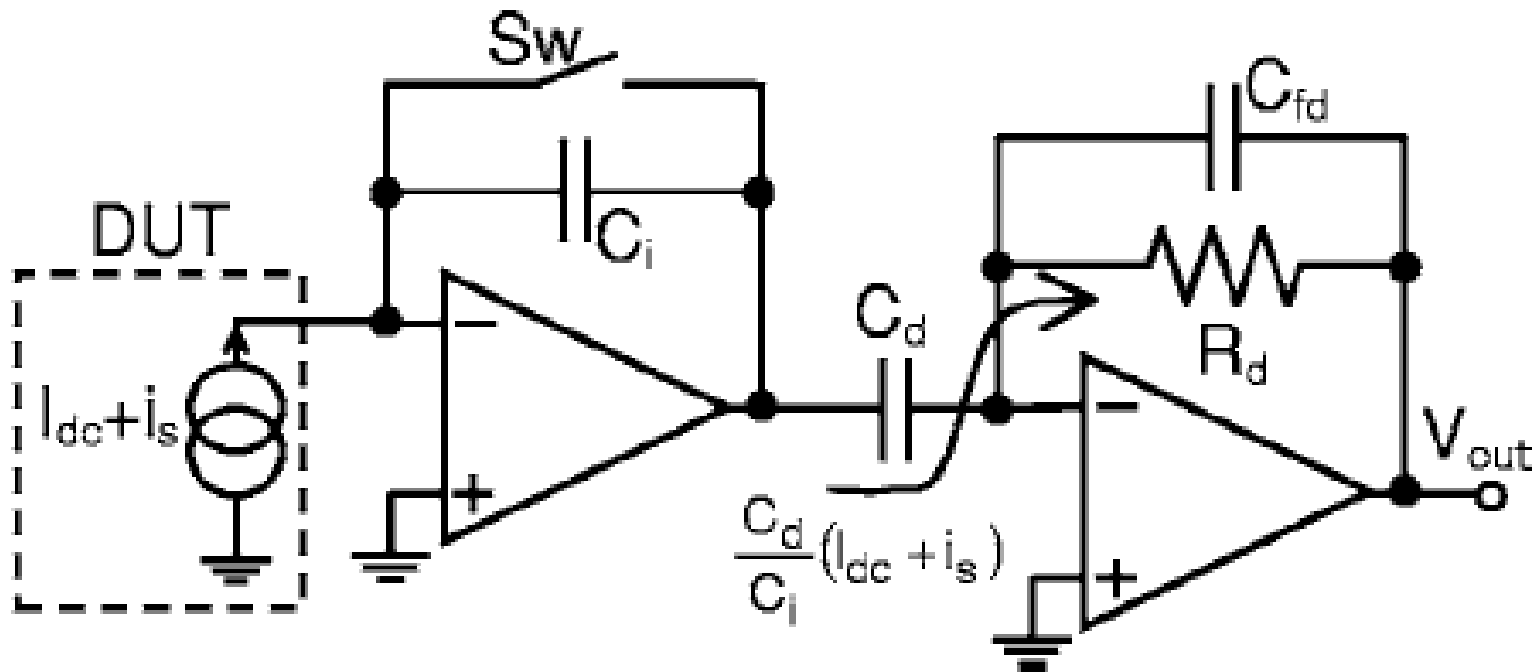
2.1 - Discrete Time: Reset Switch

Switch periodically closed to discharge the capacitor:



Limits the maximum measurement time

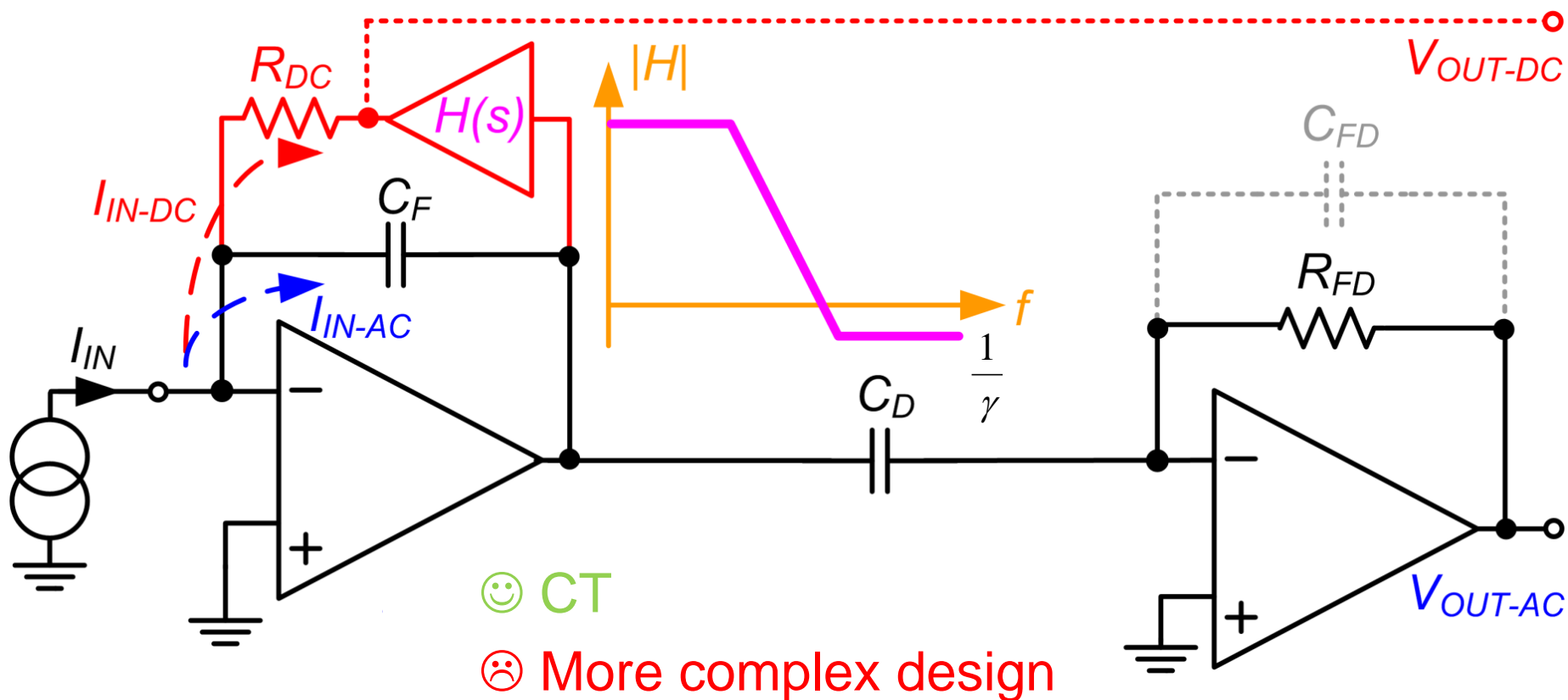
For example: $I_{IN-DC} = 1\text{nA}$, $C_F = 1\text{pF} \rightarrow (5\text{V}) T_{MAX} = 5\text{ms}$





2.2 - Continuous Time: Active Reset

To achieve continuous-time operation: additional feedback branch $H(s)$ with high gain at low frequency:





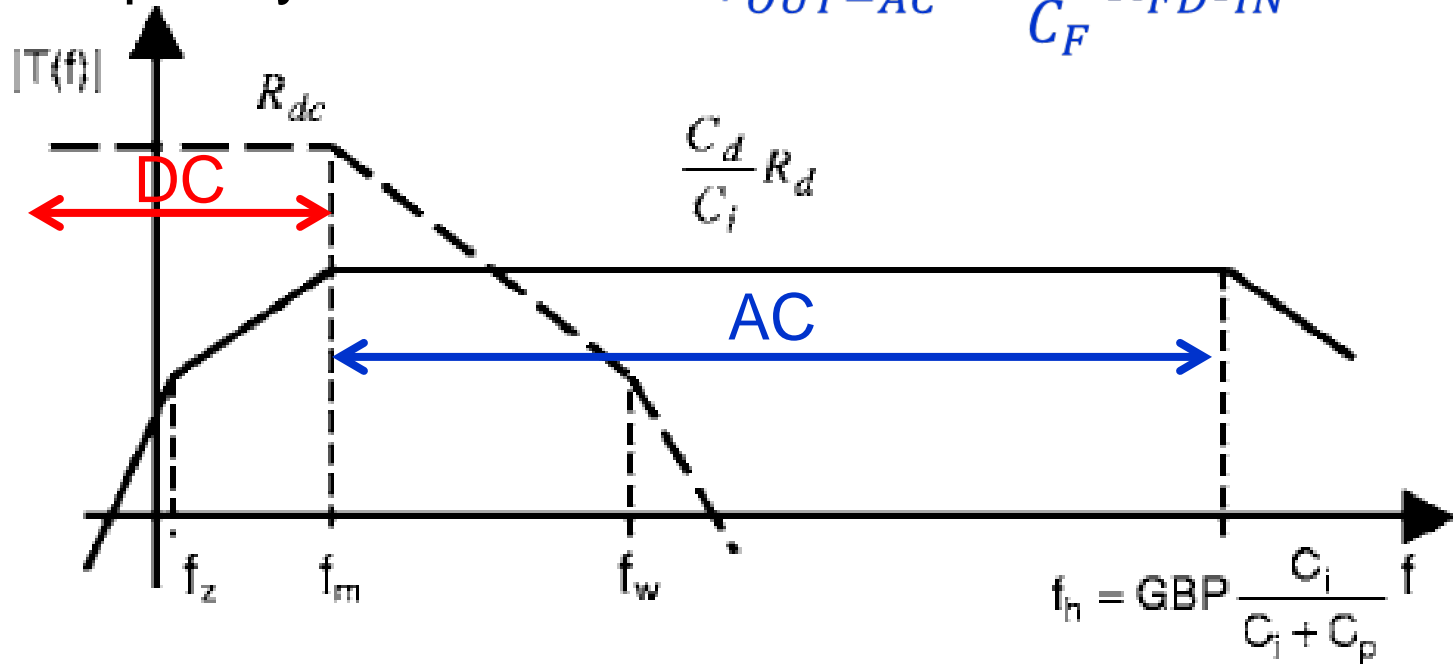
Transfer Functions

Two separate paths and gains:

- DC and low frequency
- High frequency

$$V_{OUT-DC} = -R_{DC} I_{IN}$$

$$V_{OUT-AC} = \frac{C_D}{C_F} R_{FD} I_{IN}$$

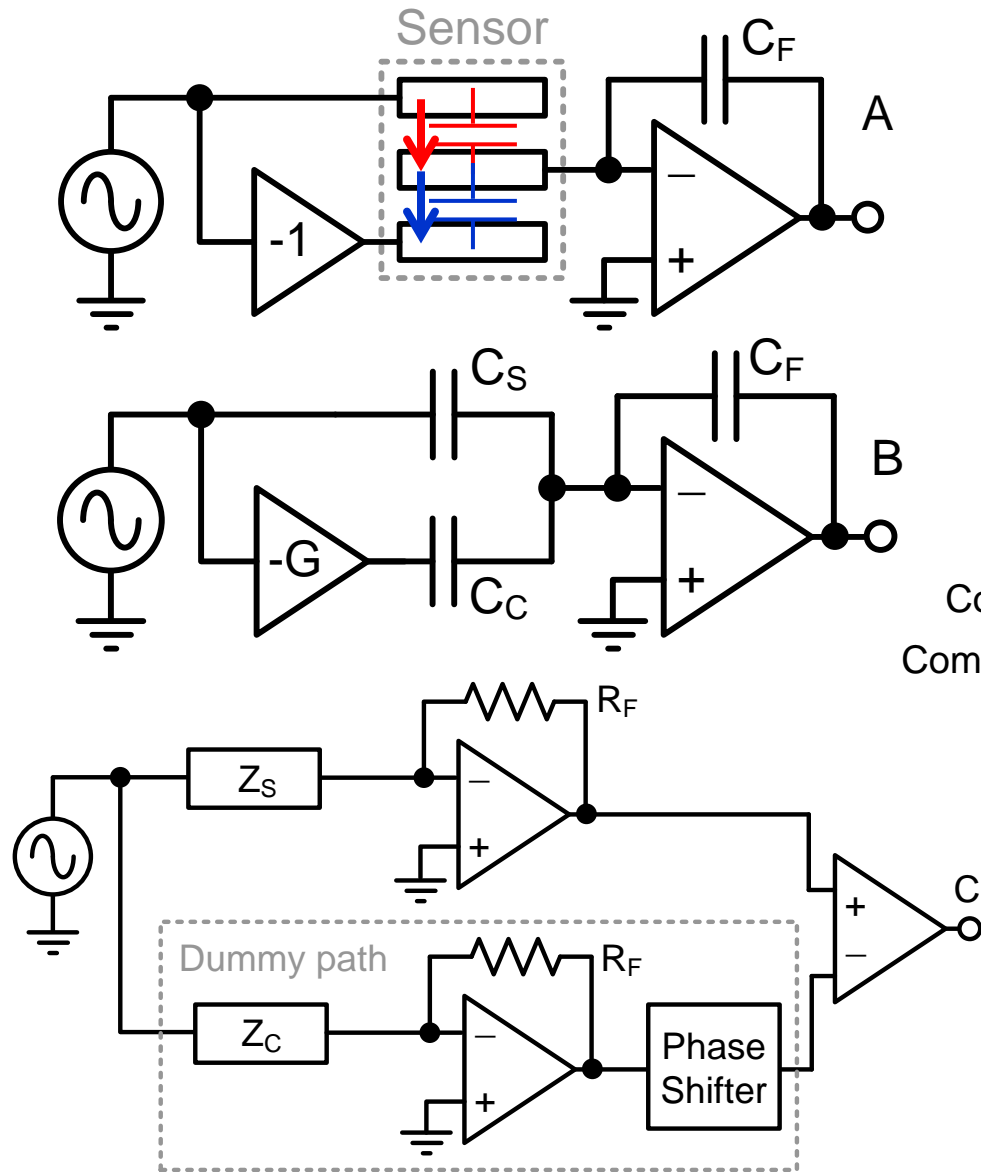


$$f_m = \frac{1}{2\pi C_F R_{DC} \gamma}$$

For wideband impedance spectroscopy,
large R_{DC} is required



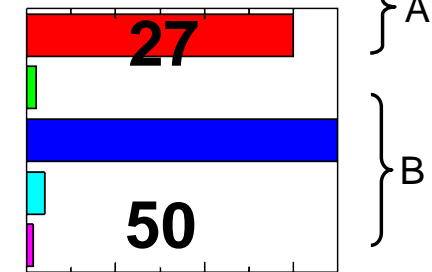
Mitigation of Source Noise: Differential Sensing



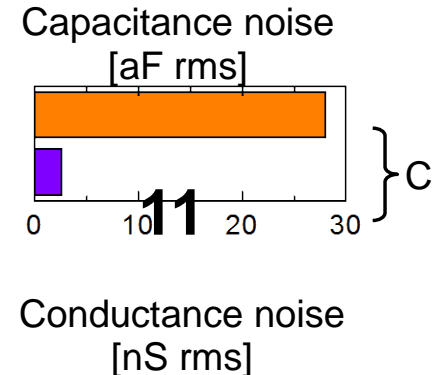
Limited by matching!

Significant noise reduction

Single-ended } A
 Differential }
 No compensation }
 Compesanted (same range) } B
 Compesanted (range reduced) } C



No compensation } C
 Compesanted }



M. Carminati, *Proc. IEEE SSD 2014* (Best Paper Award).

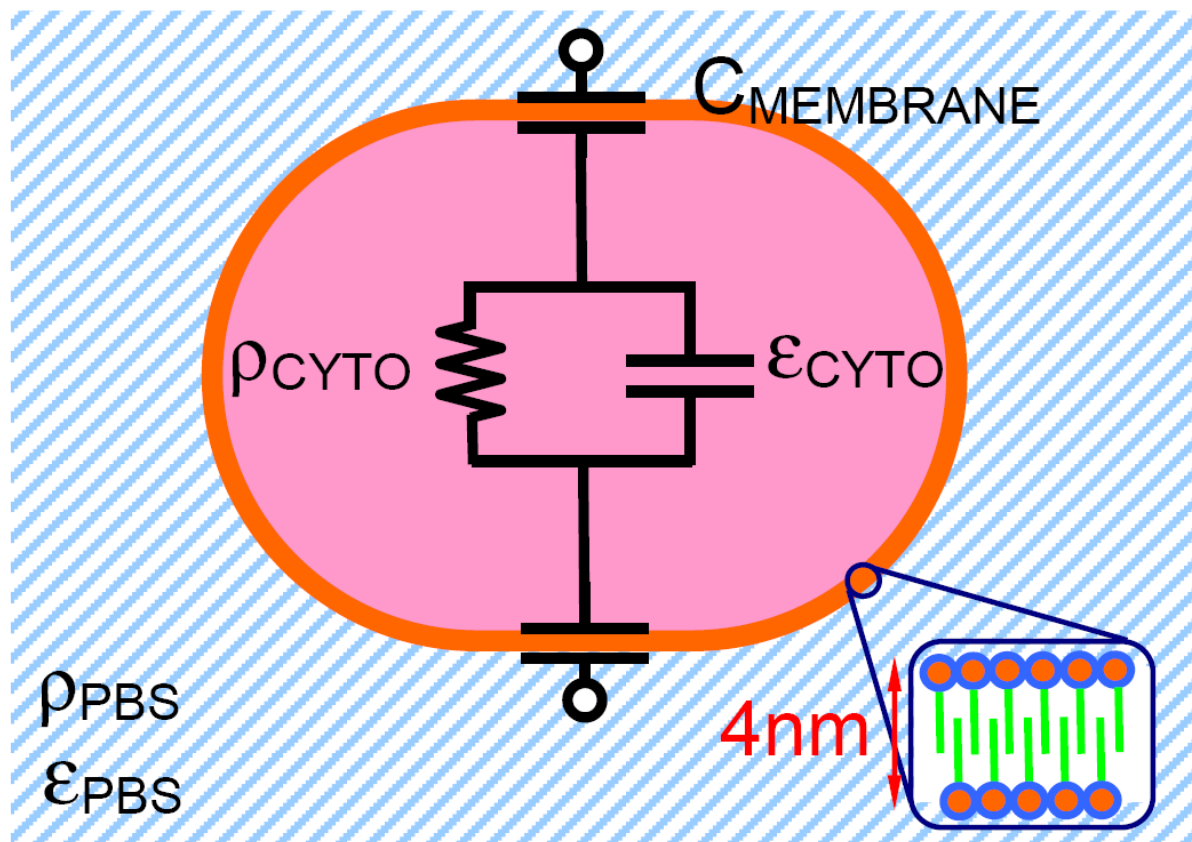


1. Biological Cells (*resistance*)
2. Airborne Dust (*capacitance*)
3. Silicon Photonics (*resistance in AC*)



Applications to Cell Biology

Small signal equivalent of a passive cell: the **single shell model**



- $\rho_{PBS} = 66\Omega\text{cm}$
- $\epsilon_{PBS} = 78$
- $C_{MEM} = 0.01\text{pF}/\mu\text{m}^2$
- $\rho_{CYTO} \sim \rho_{PBS}$
- $\epsilon_{CYTO} = 60$

At low frequency ($<1\text{MHz}$), the cell can be treated as an insulating sphere (resistivity contrast)



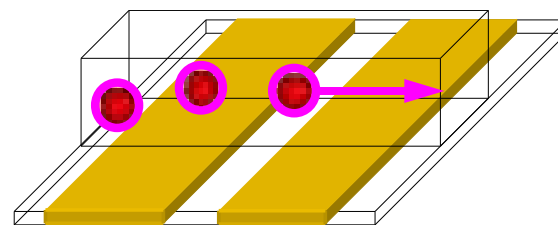
Cell Counting

The presence of cells can be sensed by impedance as a **perturbation** of the electrical field between two electrodes

Two approaches:

“Static”

“Dynamic”

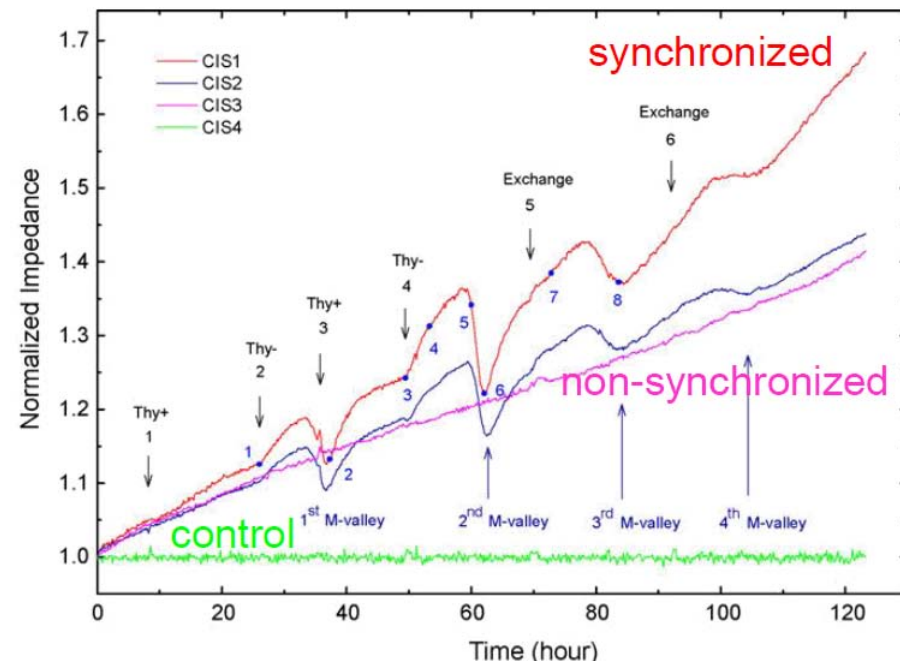
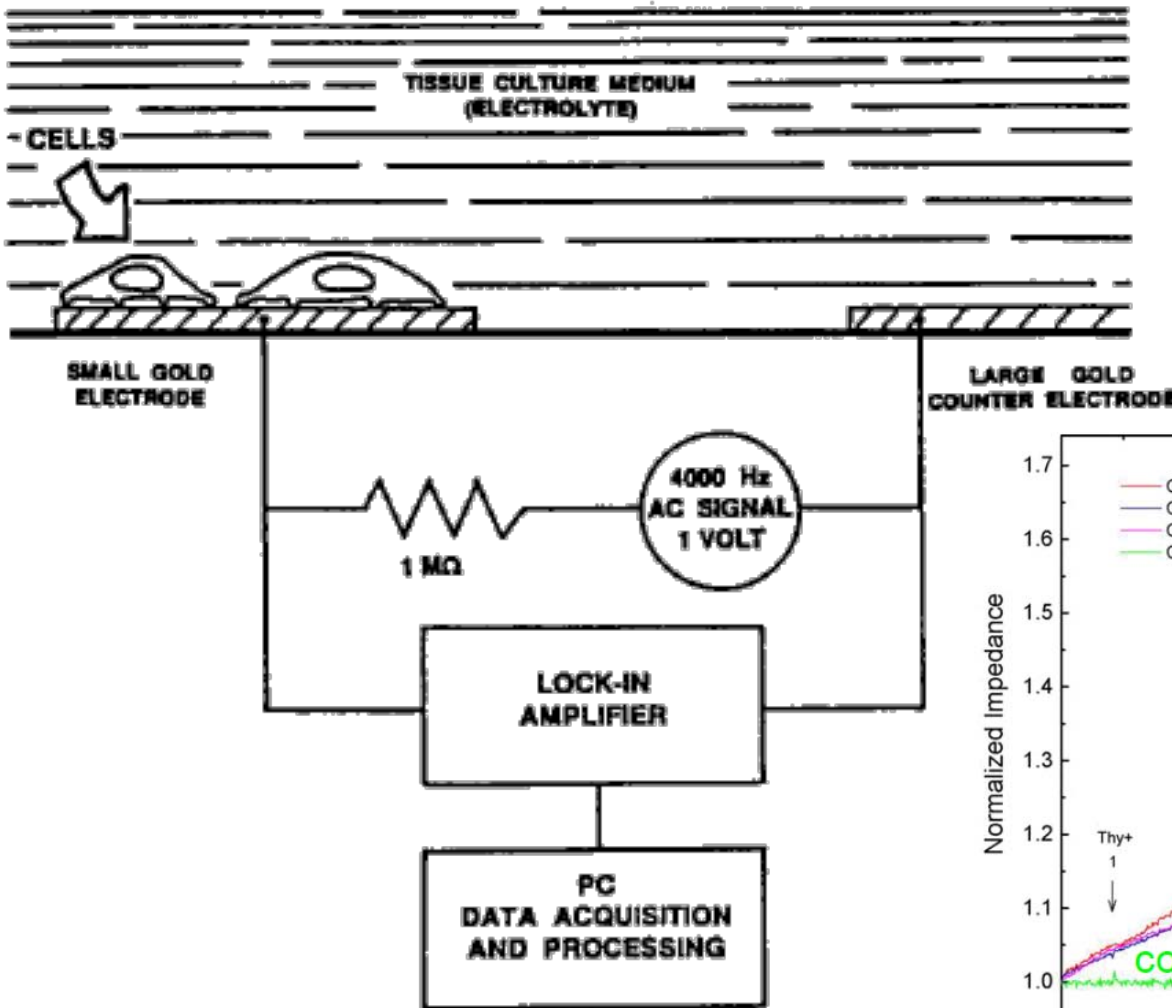




Electric Cell-substrate Impedance Sensing

ECIS Technique invented by Giaever and Keese

Cell adhesion, spreading and growth can be monitored

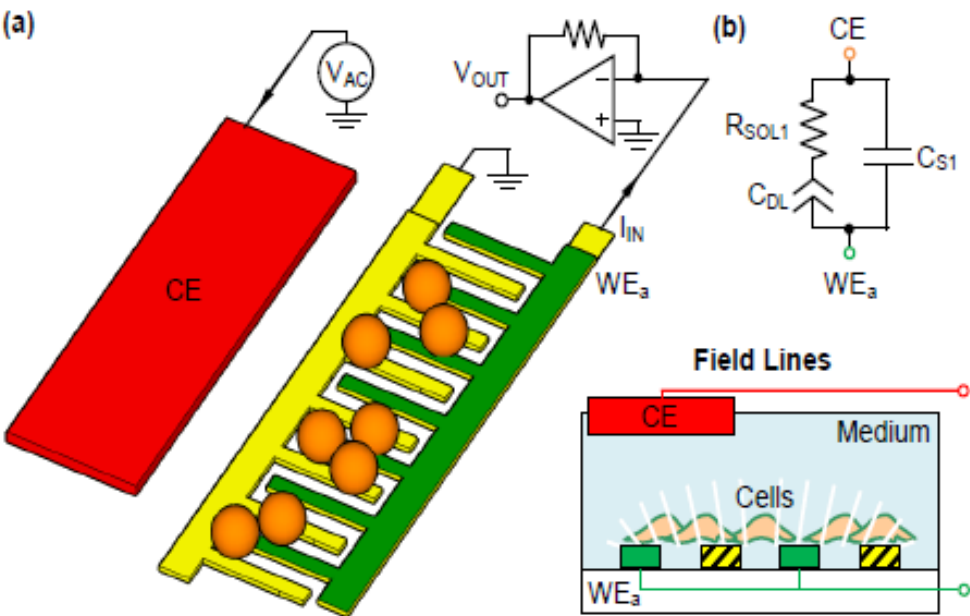


Lo, Keese, Giaever, *Biophys J.* **69** 1995

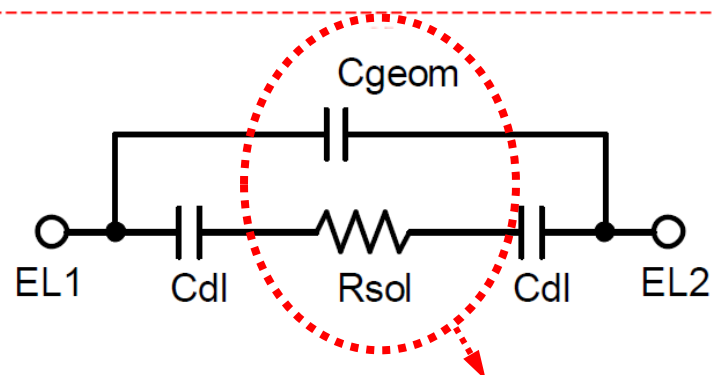
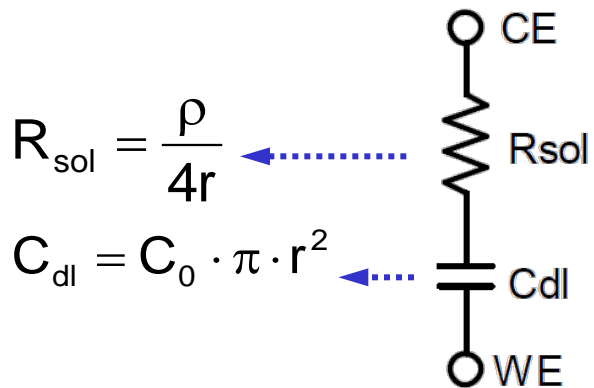
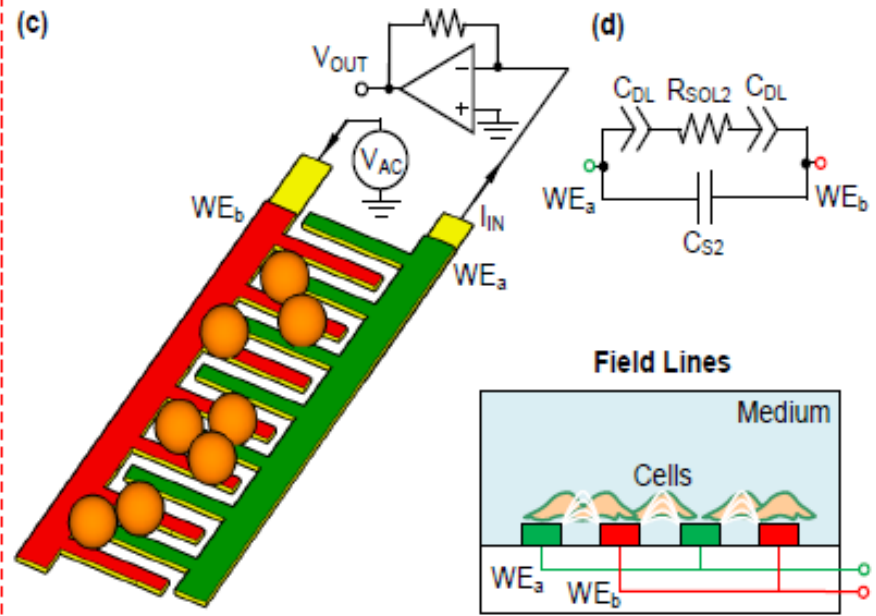


Comparison of Sensing Geometries

Vertical Sensing Configuration and Equivalent Impedance Model



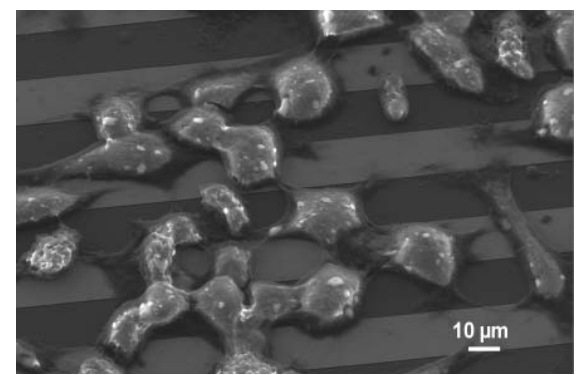
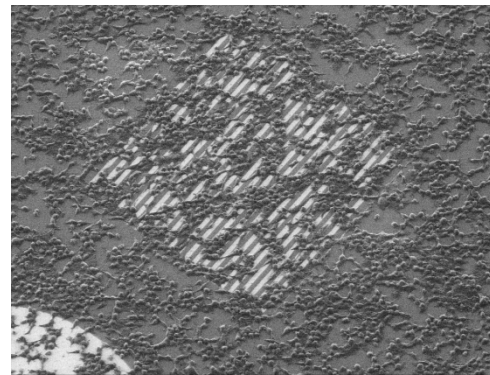
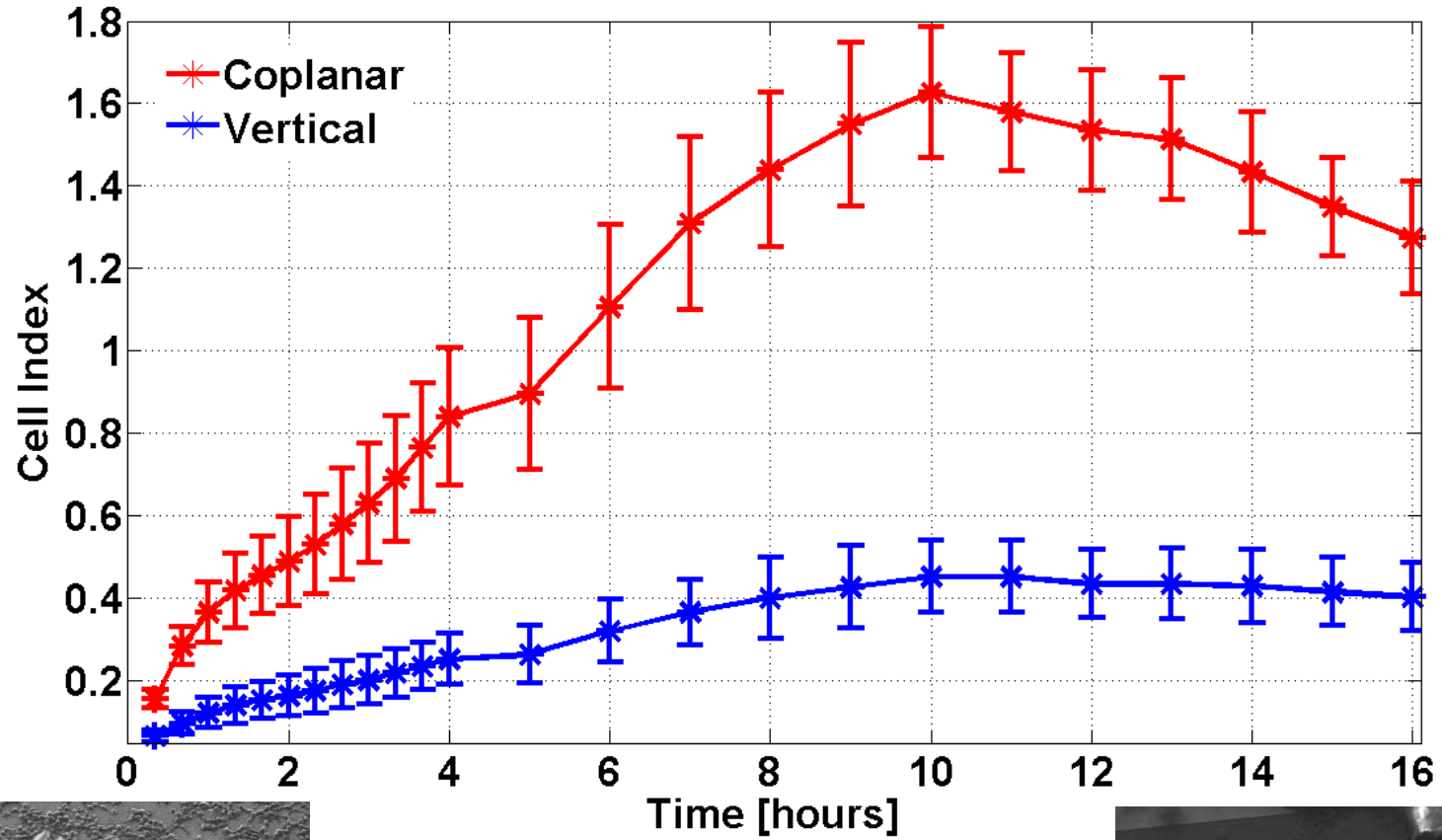
Coplanar Sensing Configuration and Equivalent Impedance Model



Conformal Mapping

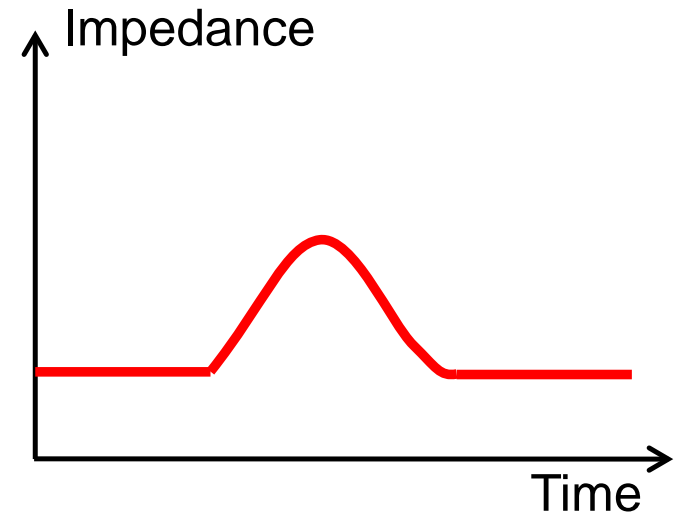
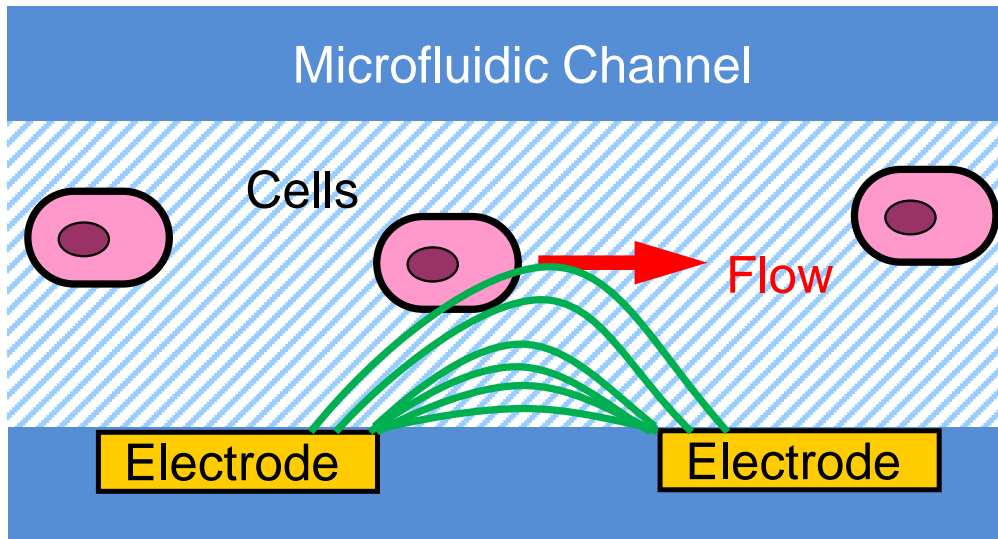


Comparison with HeLa Cells





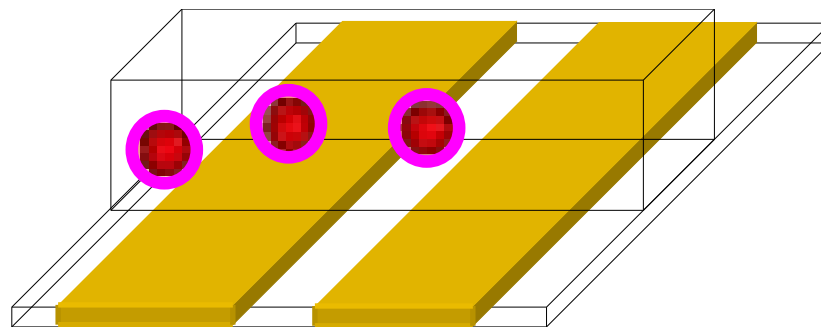
Impedance Flow Cytometry



- A novel approach, pioneered by Morgan and Renaud (~2000)
- Tracking impedance between close electrodes over time
- High-throughput single-cell analysis (beyond sizing)

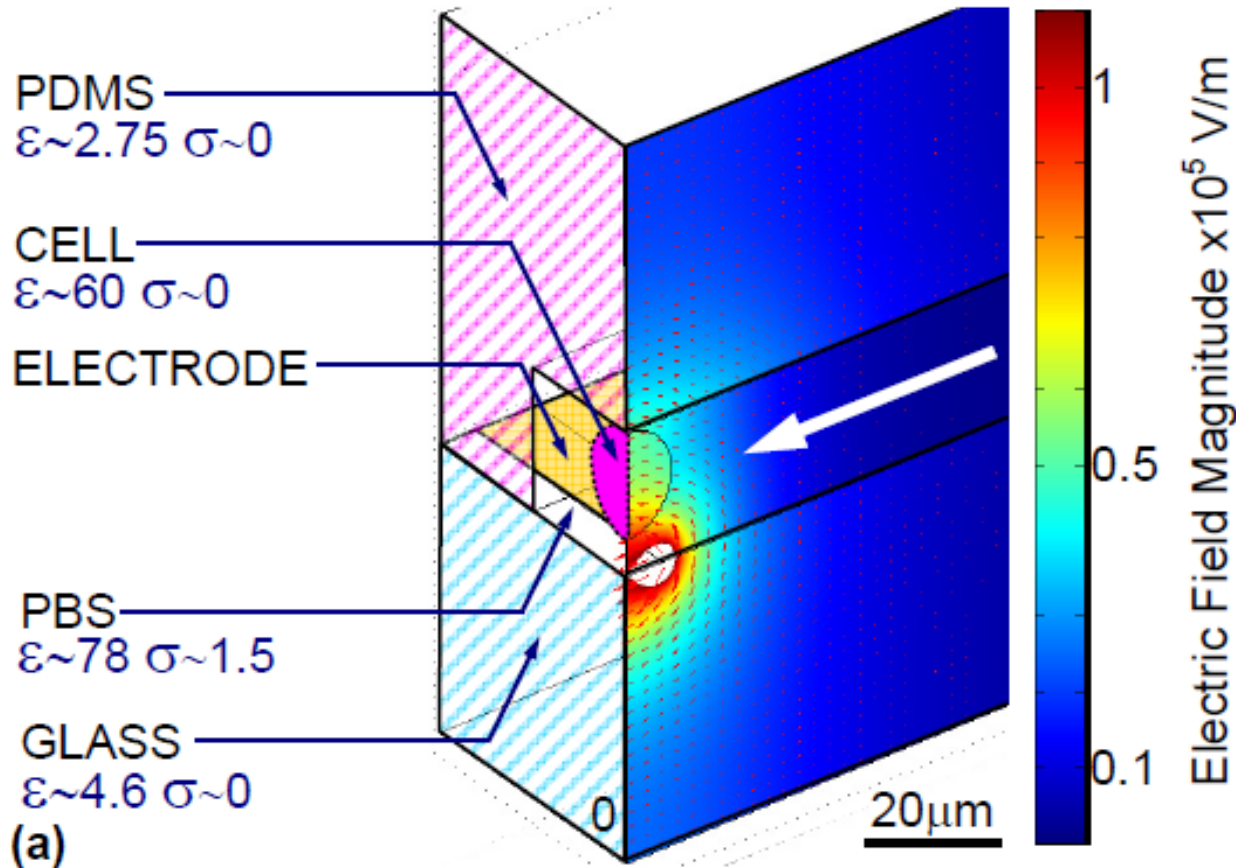
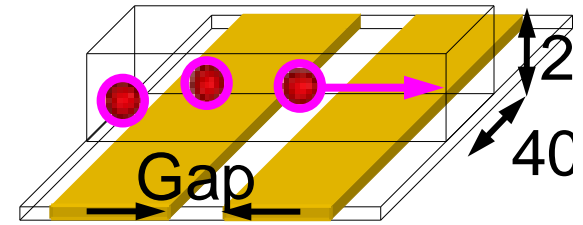


Electrodes Configurations



Design of Sensing Electrodes

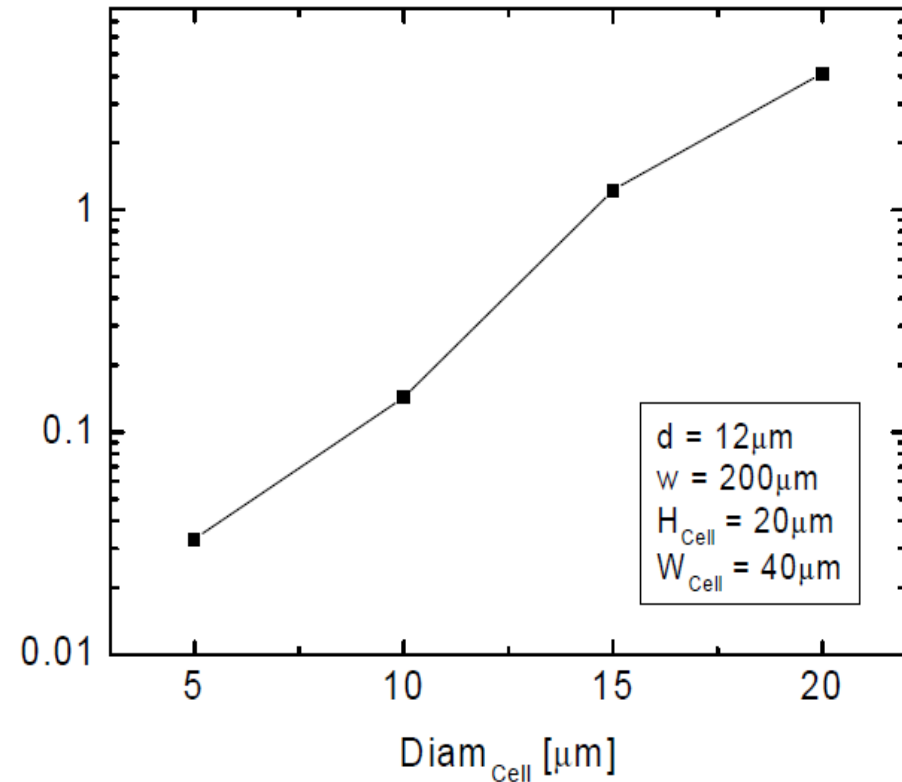
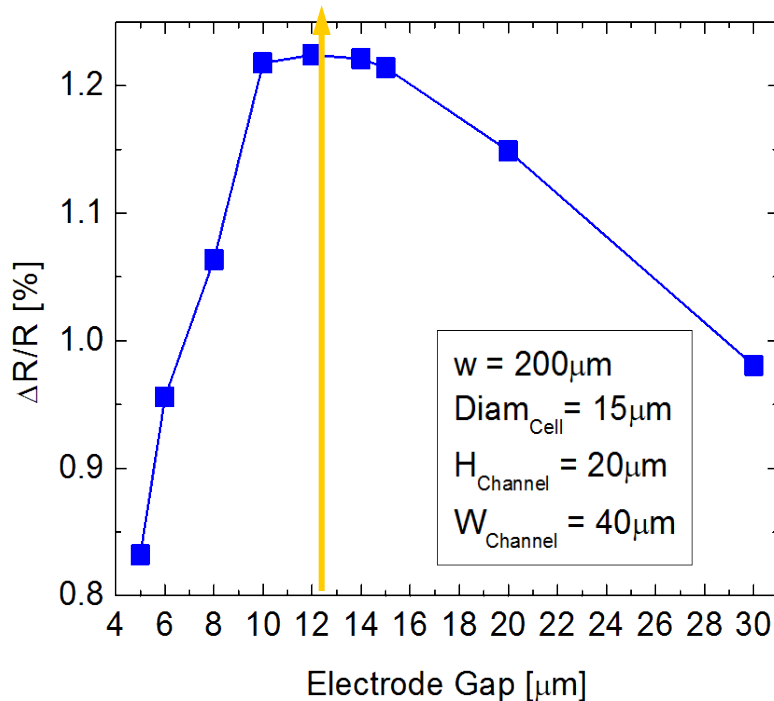
- Simple technology → coplanar
- Ease of alignment → transverse
- Target cells 5-15 μm → 20 x 40 μm^2





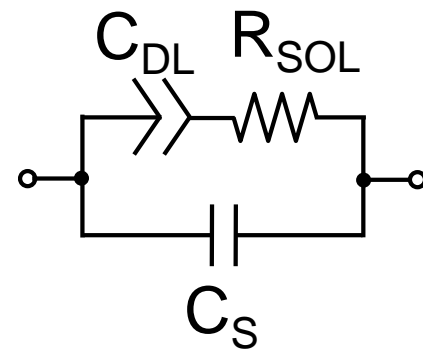
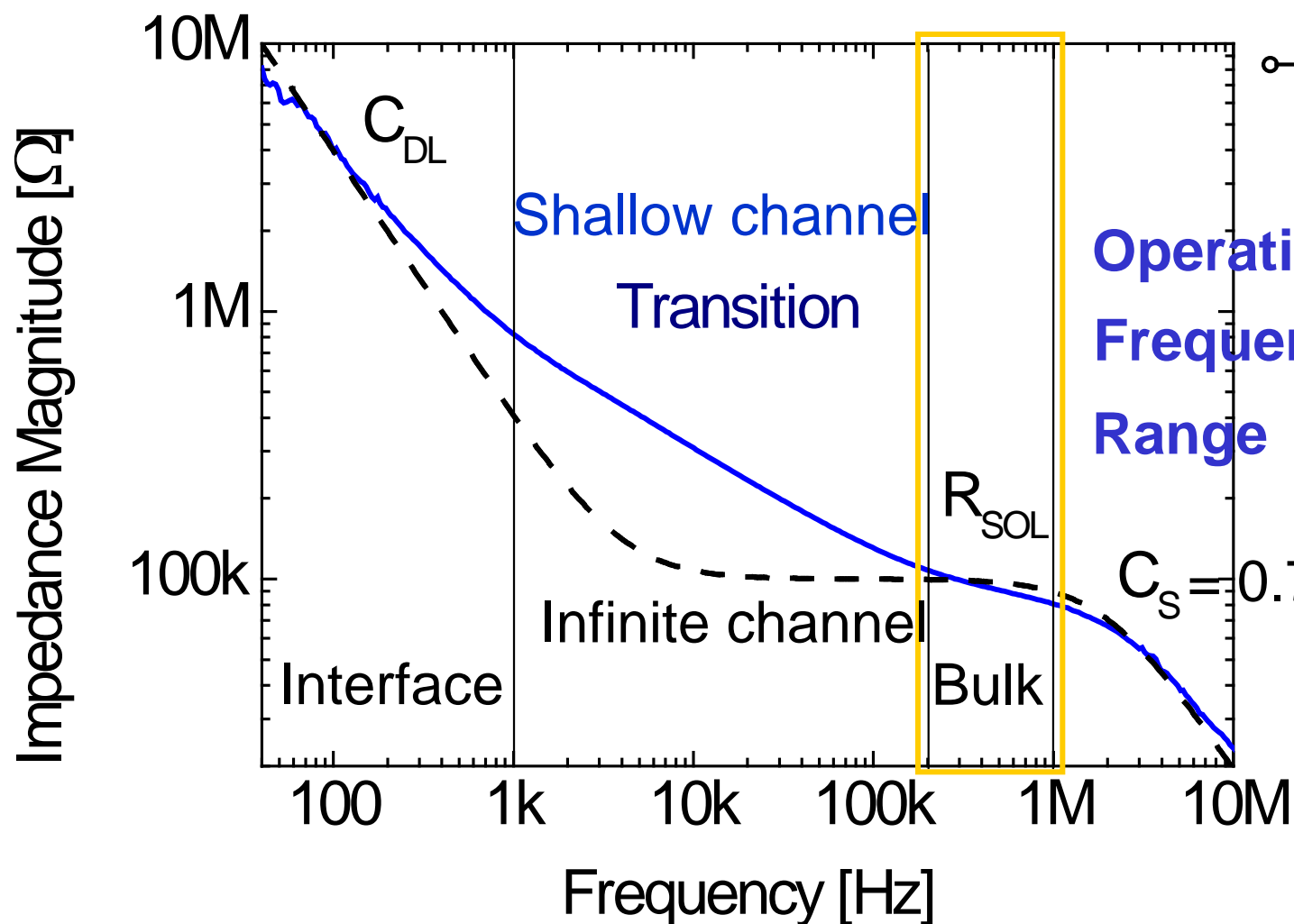
Optimal Electrode Design

- The gap sets the vertical extension of the sensing volume
- An optimal gap exists for a given cell diameter
- Sensitivity scales with cell volume and vertical height



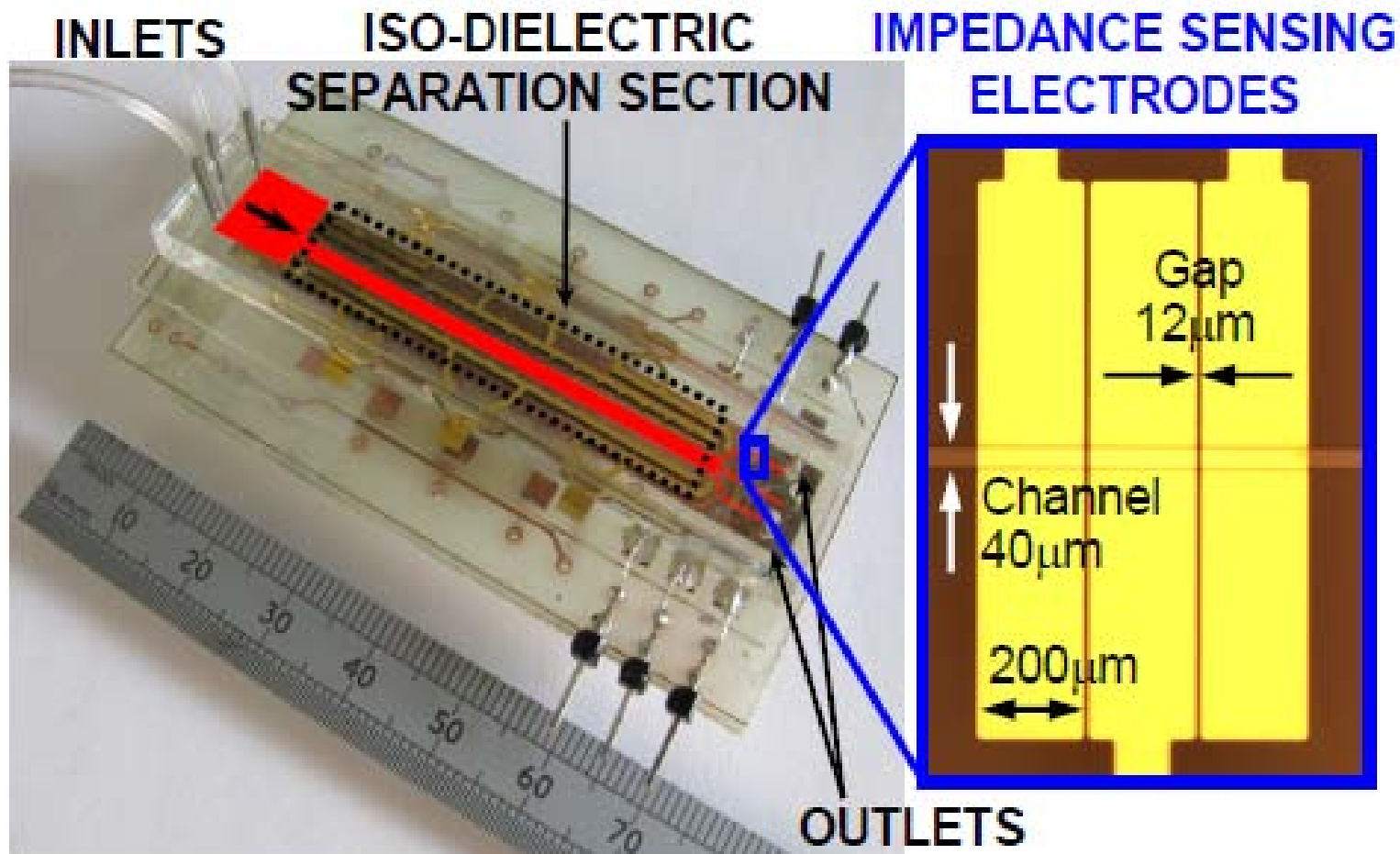
Operating Frequency

Four regions in the impedance spectrum:



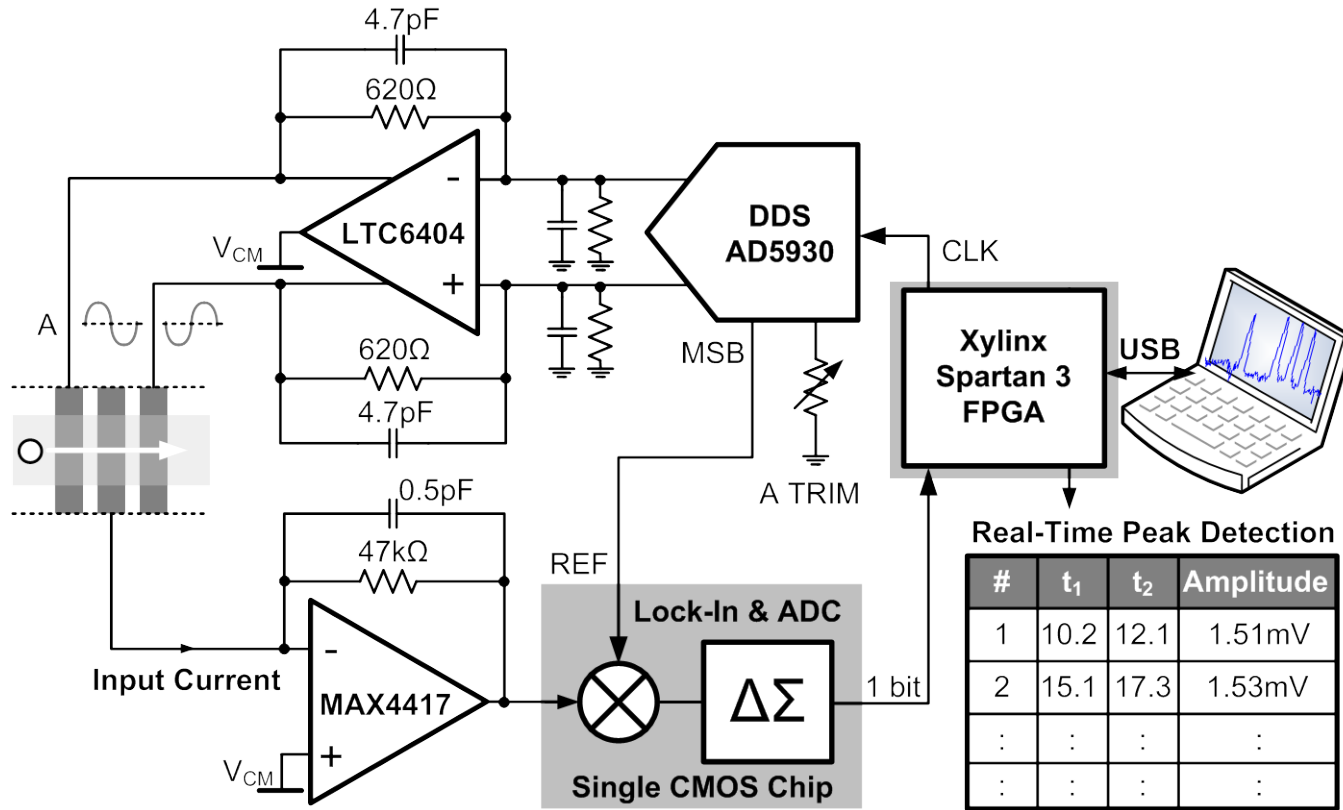
stray to be minimized

Enhanced Device Architecture



- Integrated into the device with no size increase
- Third electrode available for differential sensing (drift)

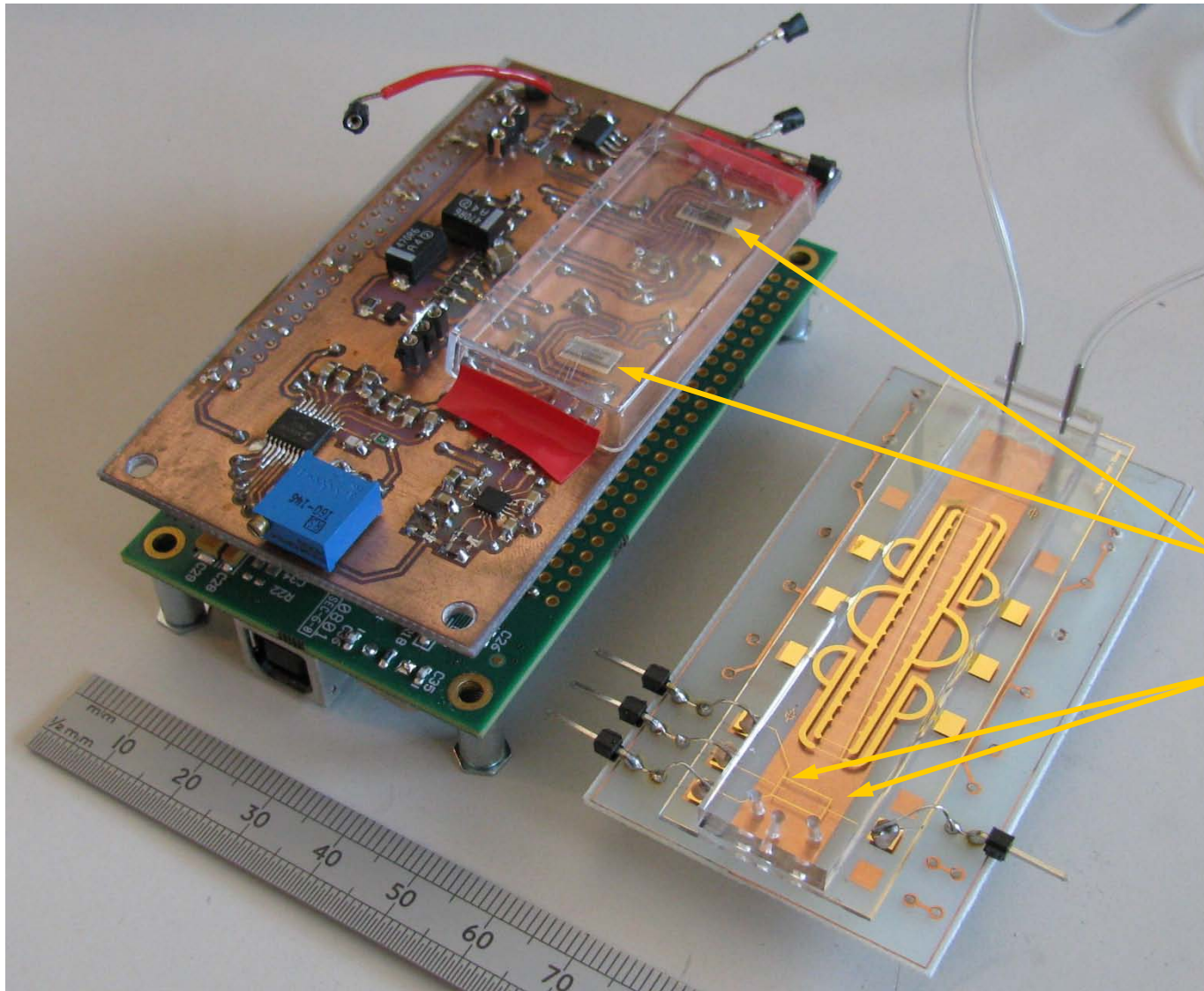
Miniaturized Impedance Detector



- On-board signal generator
- ASIC 1MHz CMOS lock-in and 20bit $\Sigma\Delta$ converter
- Real-time FPGA peak detection algorithm
- Digital USB data acquisition



Credit Card Sized Implementation



Ultra compact system

3.3V USB Power Supply

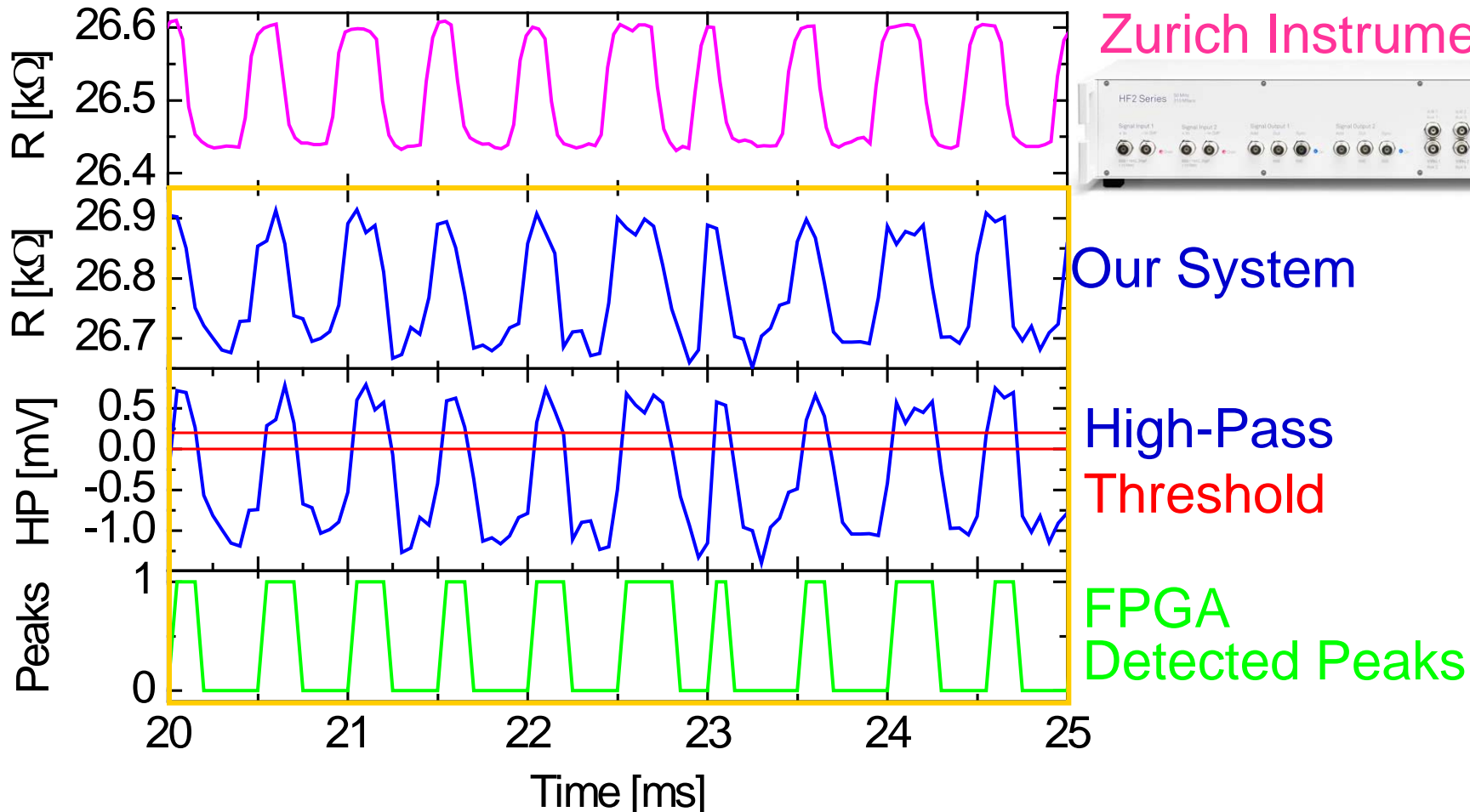
2 channels:

- 2 outlets
- Complex impedance



Performance Assessment

- Impedance tracking comparable with state-of-art instrument
- Error-free **real-time FPGA peak counting** at 2000 events/s



Zurich Instruments



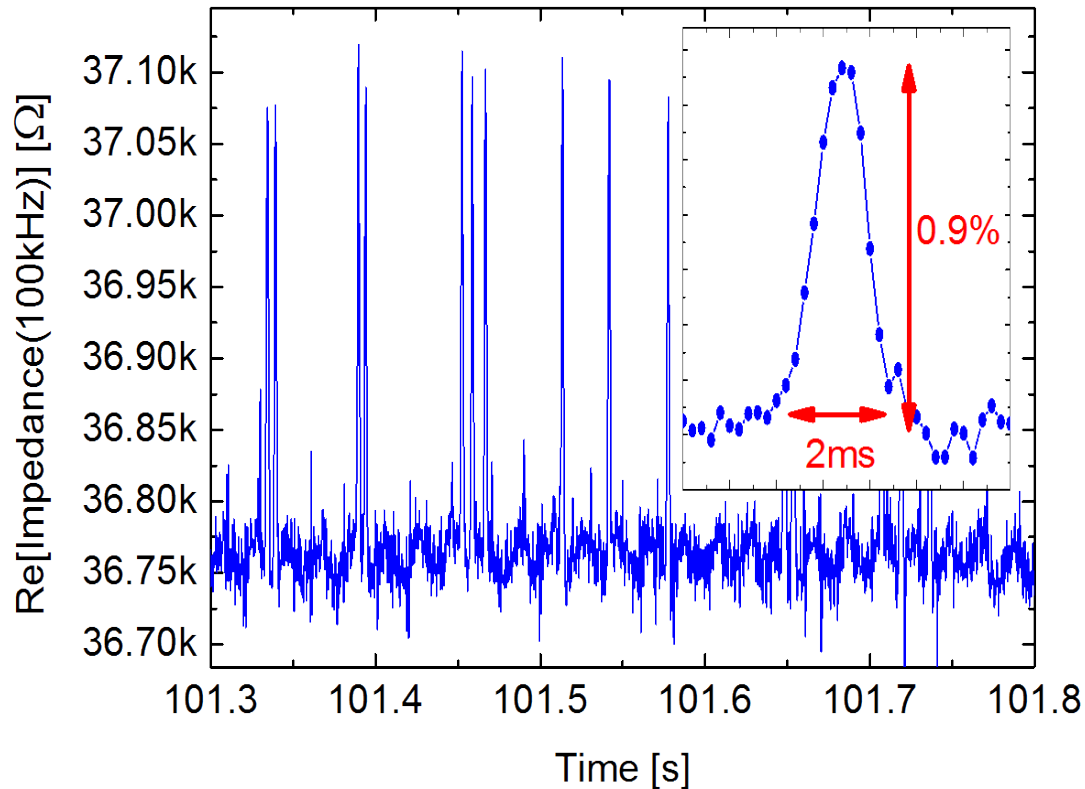
Our System

High-Pass
Threshold

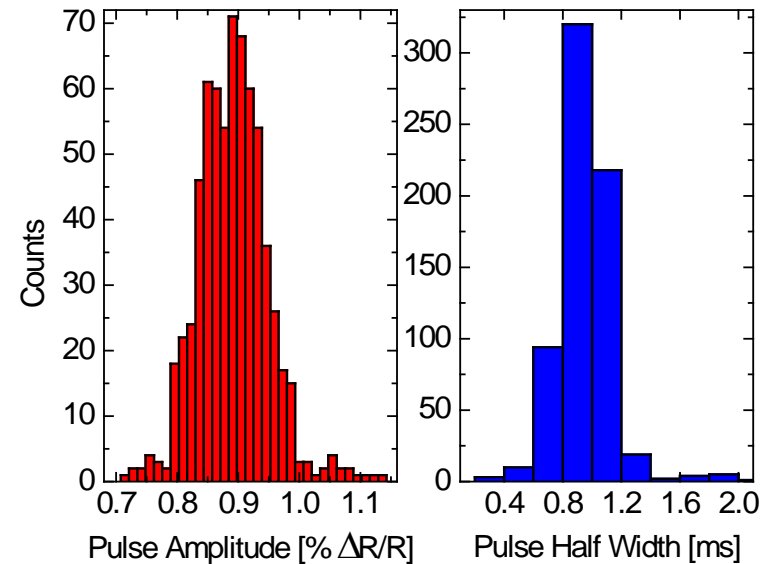
FPGA
Detected Peaks

System Validation with Beads

- 10 μm polystyrene beads in PBS
- 50 beads/s (1 $\mu\text{l}/\text{min}$)
- $V_{AC} = 50\text{mV}$, $f_0 = 100\text{kHz}$, 5kSa/s



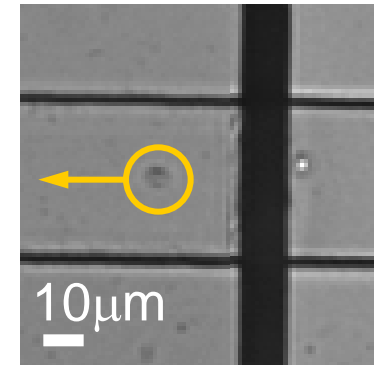
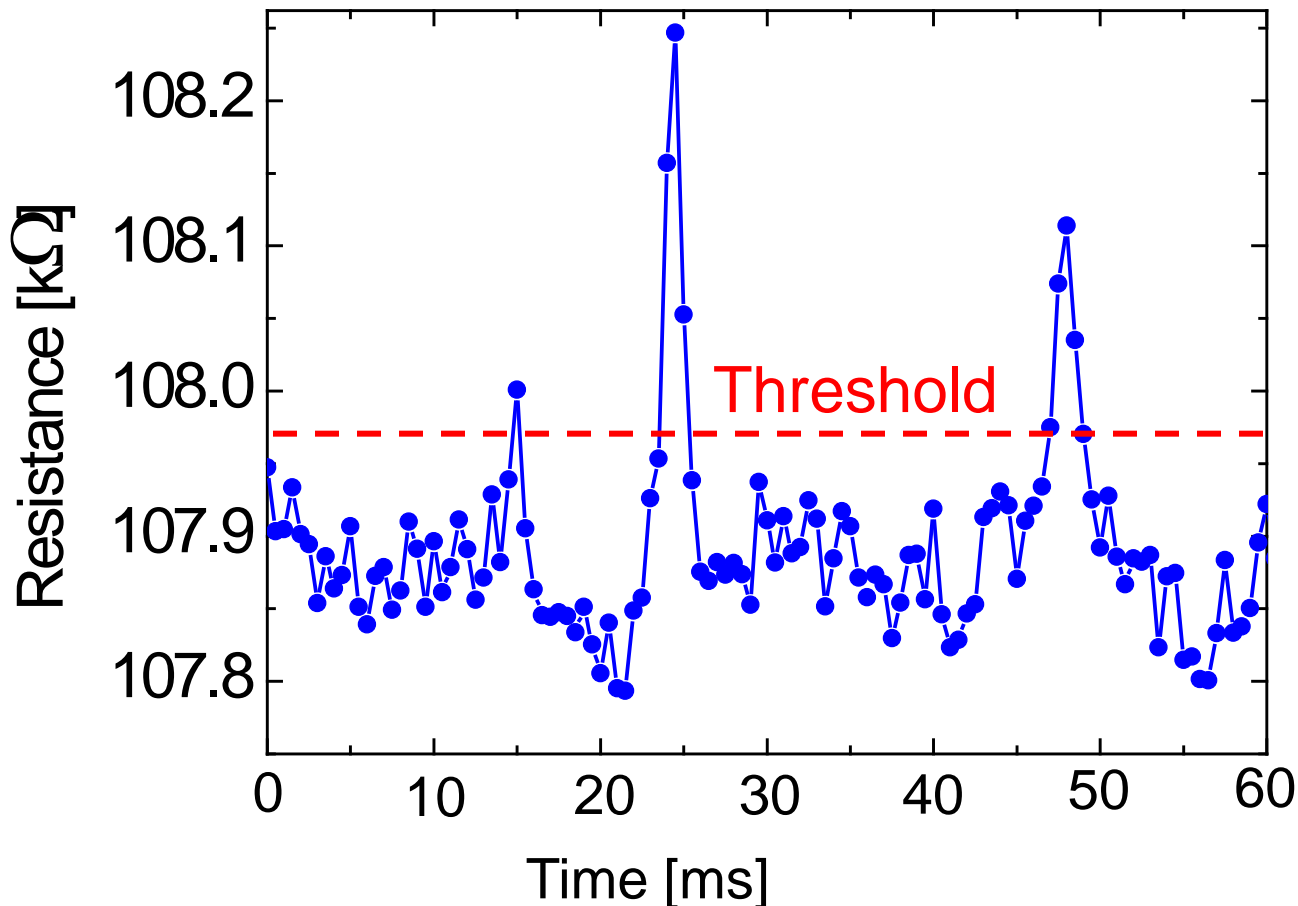
As expected:





Counting Yeast

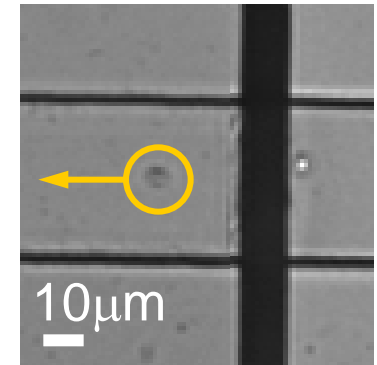
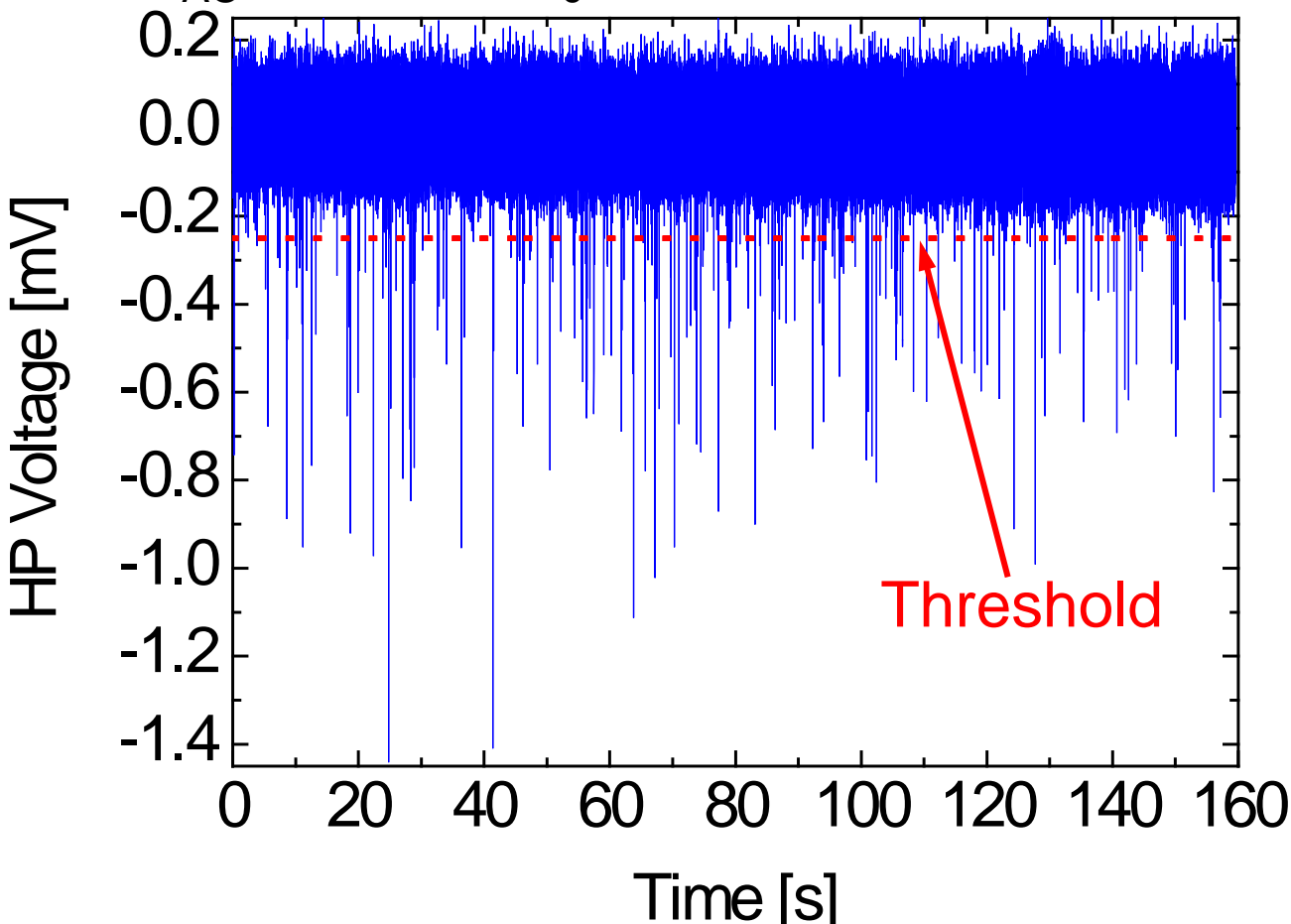
- *Sacc. Cerevisiae* ($5\mu\text{m}$)
- 10^5 cells/ml in 0.35S/m medium (with BSA)
- $V_{AC} = 650\text{mV}$, $f_0 = 100\text{kHz}$, 2kSa/s





Counting Yeast

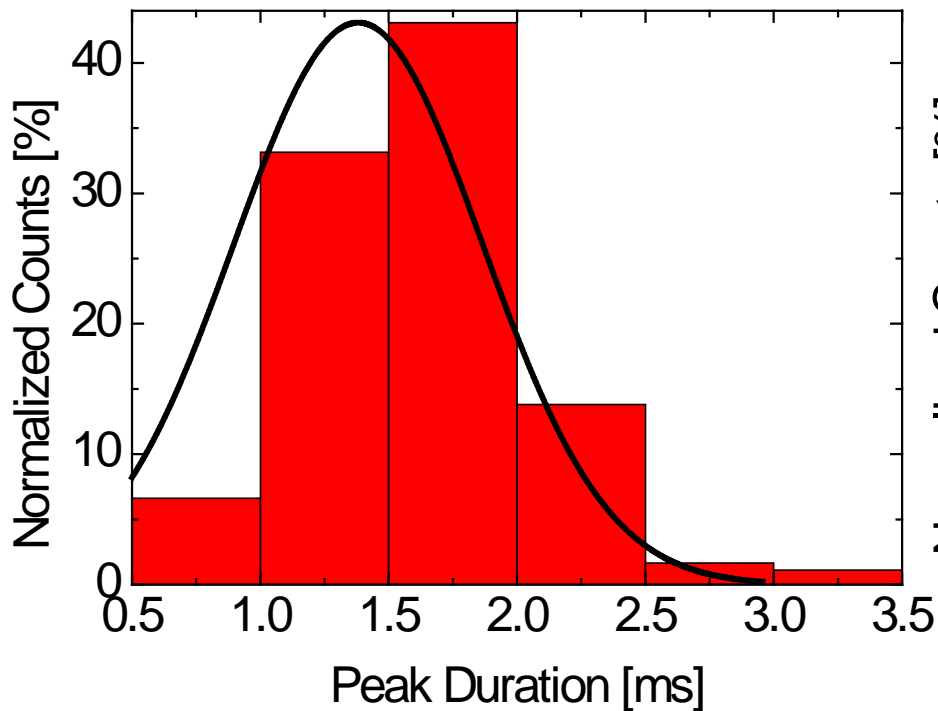
- *Sacc. Cerevisiae* ($5\mu\text{m}$)
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- $V_{AC} = 650\text{mV}$, $f_0 = 100\text{kHz}$, 2kSa/s



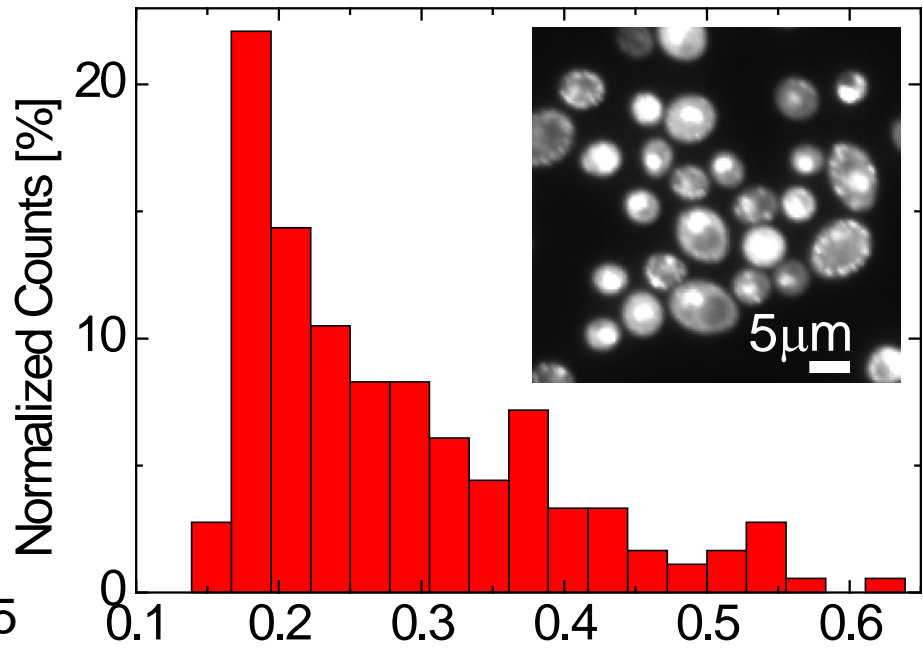


Yeast: Peak Analysis

Pulse Width:

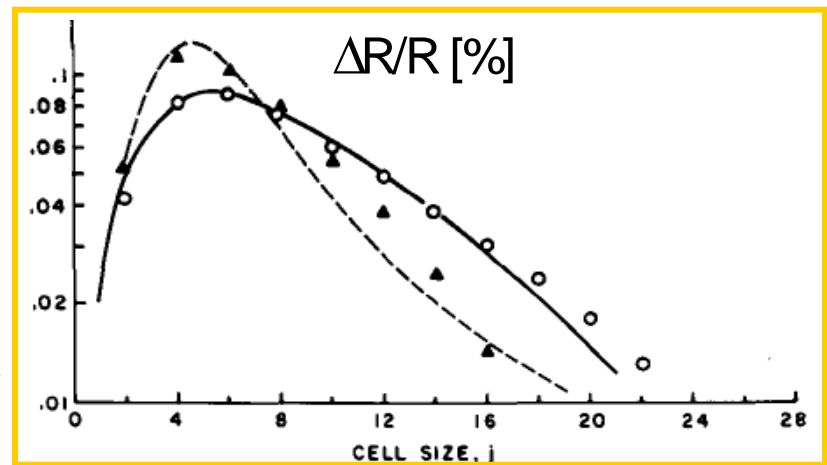


Amplitude:



Consistent with:

- cell velocity (3cm/s)
- cell volume (10 μm beads/8)
- cell size distribution

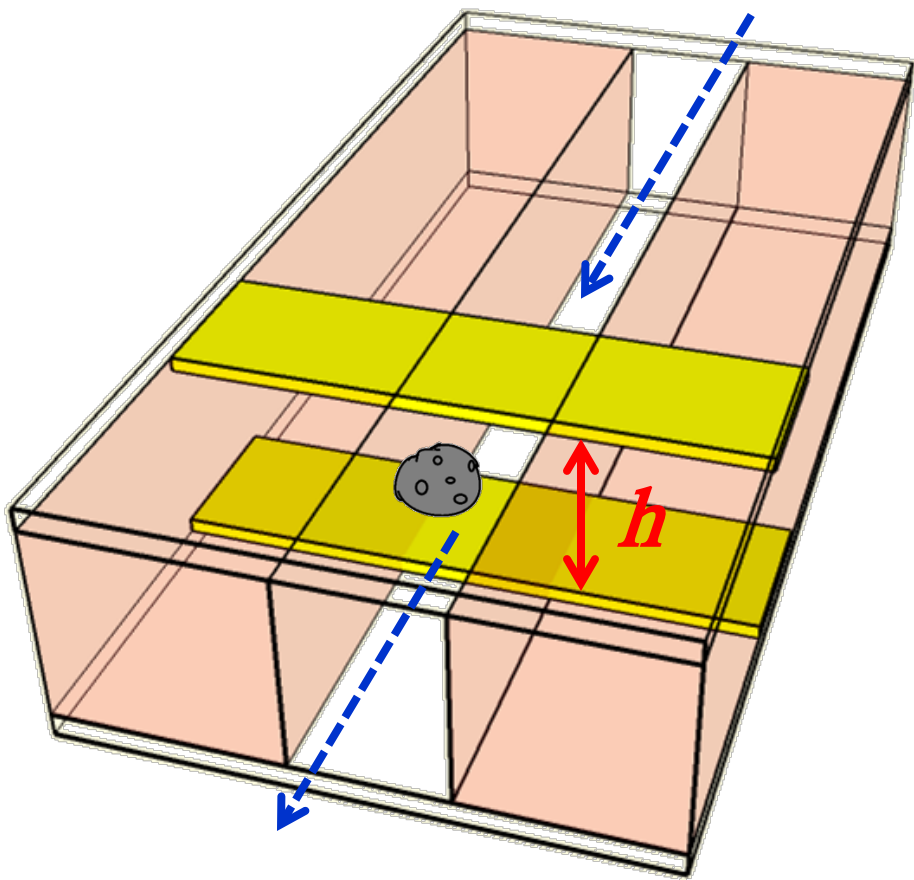


P.Palatt and G. Saidel, *Ann. Biomed. Eng.* 7 1979



A Novel Technique: Electrical Detection

Real-time capacitive detection of single PM particles



$$\frac{\Delta C}{C} = \frac{Volume_{particle}}{Volume_{total}} \cdot \Delta \epsilon$$

$$\Delta C = \frac{4}{3} \cdot \pi \cdot \frac{R^3}{h^2} \cdot \Delta \epsilon \cdot \epsilon_0$$

$$R = \sqrt[3]{\frac{\Delta C \cdot h^2}{\frac{4}{3} \cdot \pi \cdot \Delta \epsilon \cdot \epsilon_0}}$$

M. Carminati, Capacitive detection of micrometric airborne particulate matter for solid-state personal air quality monitors, *Sensors Actuators A* **219**, 2014.

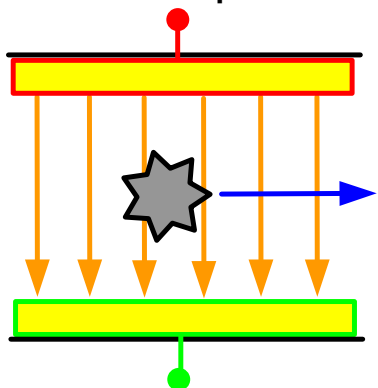


Choice of the Sensing Configuration

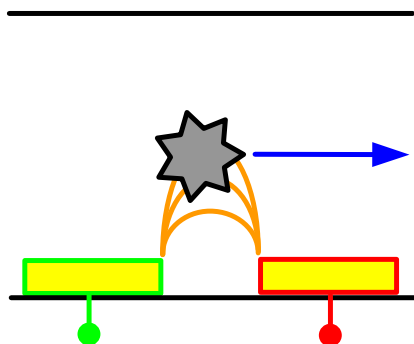
Key design choices:

- **Air** vs liquid
- Electrode configuration:

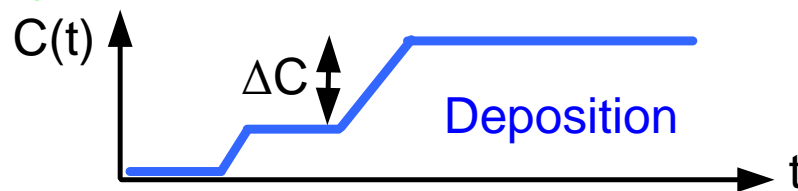
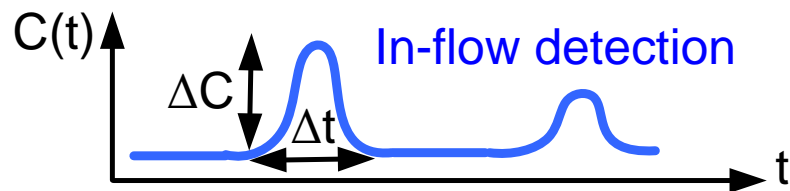
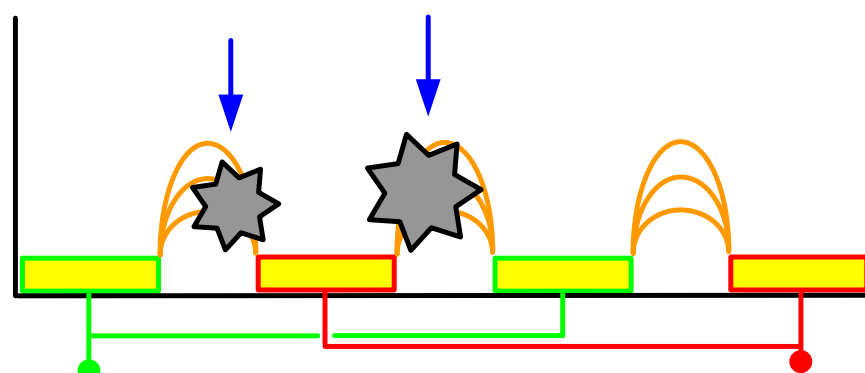
Parallel plates



Coplanar pair



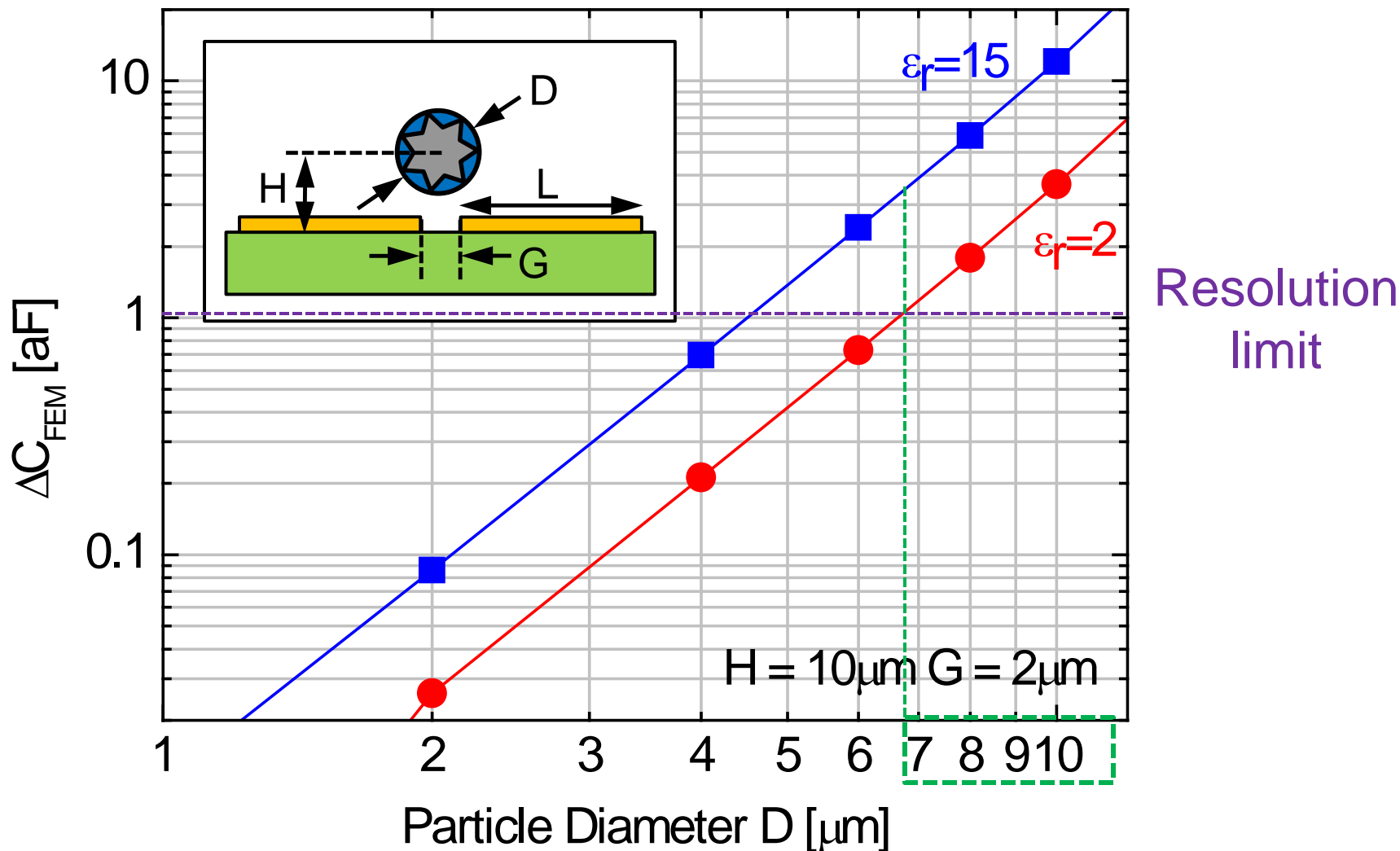
Interdigitated combs



M. Carminati, Capacitive detection of micrometric airborne particulate matter for solid-state personal air quality monitors, *Sensors Actuators A* **219**, 2014.

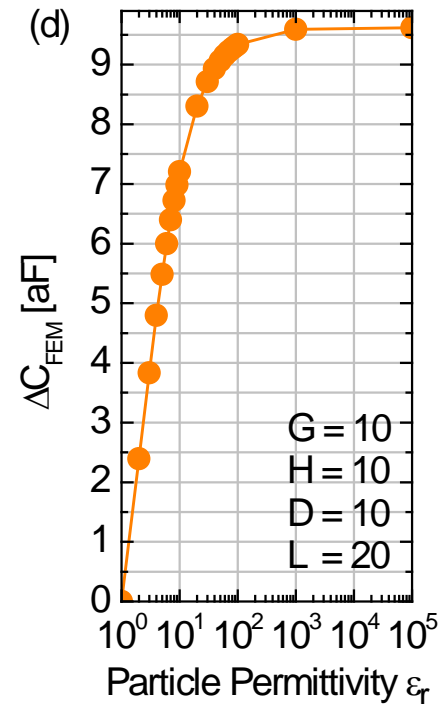
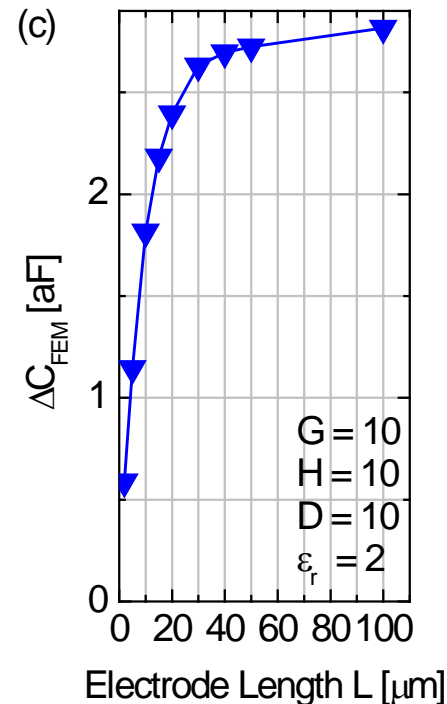
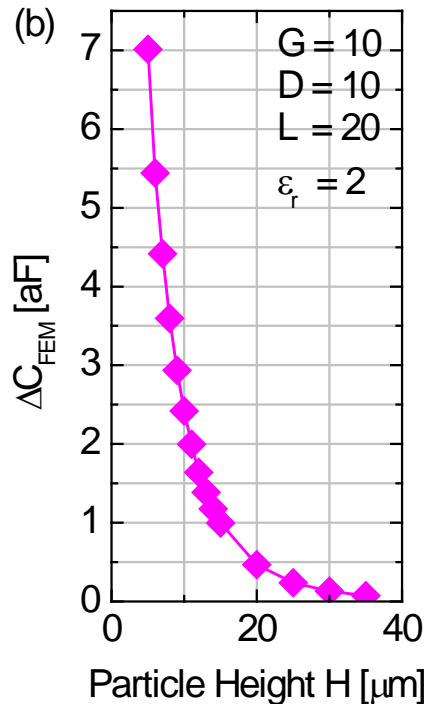
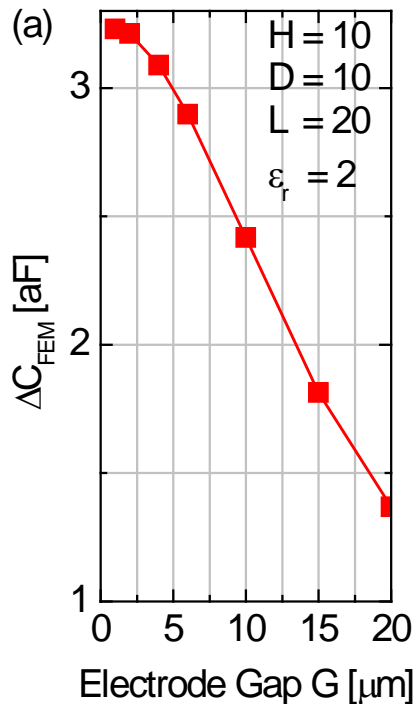
Preliminary FEM Simulations

Also in the coplanar geometry, ΔC has a volume dependence



Electrode Geometry Optimization

Fine tuning of the electrode parameters with FEM simulations



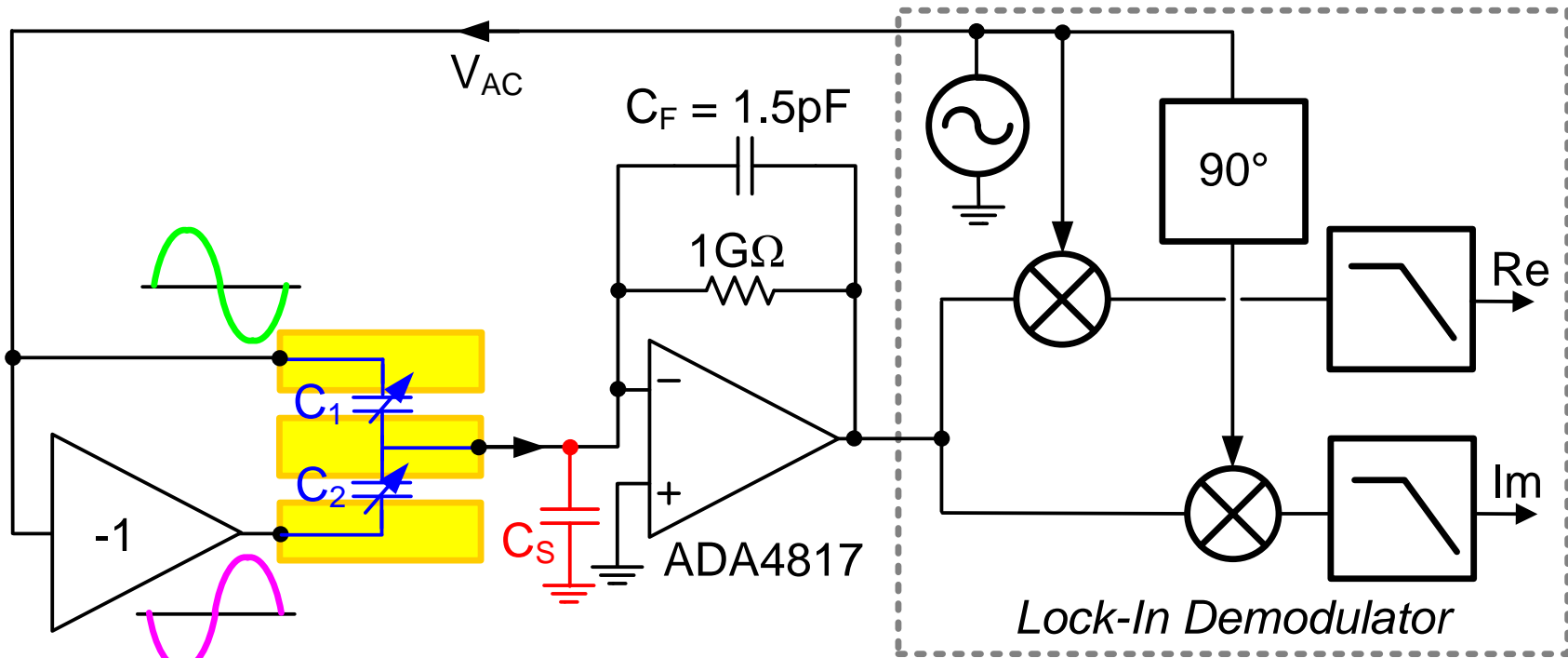
Gap: min.

Height: min.

Length: 30 μm

M. Carminati, Capacitive detection of micrometric airborne particulate matter for solid-state personal air quality monitors, *Sensors Actuators A* **219**, 2014.

Low-Noise Detection Electronics

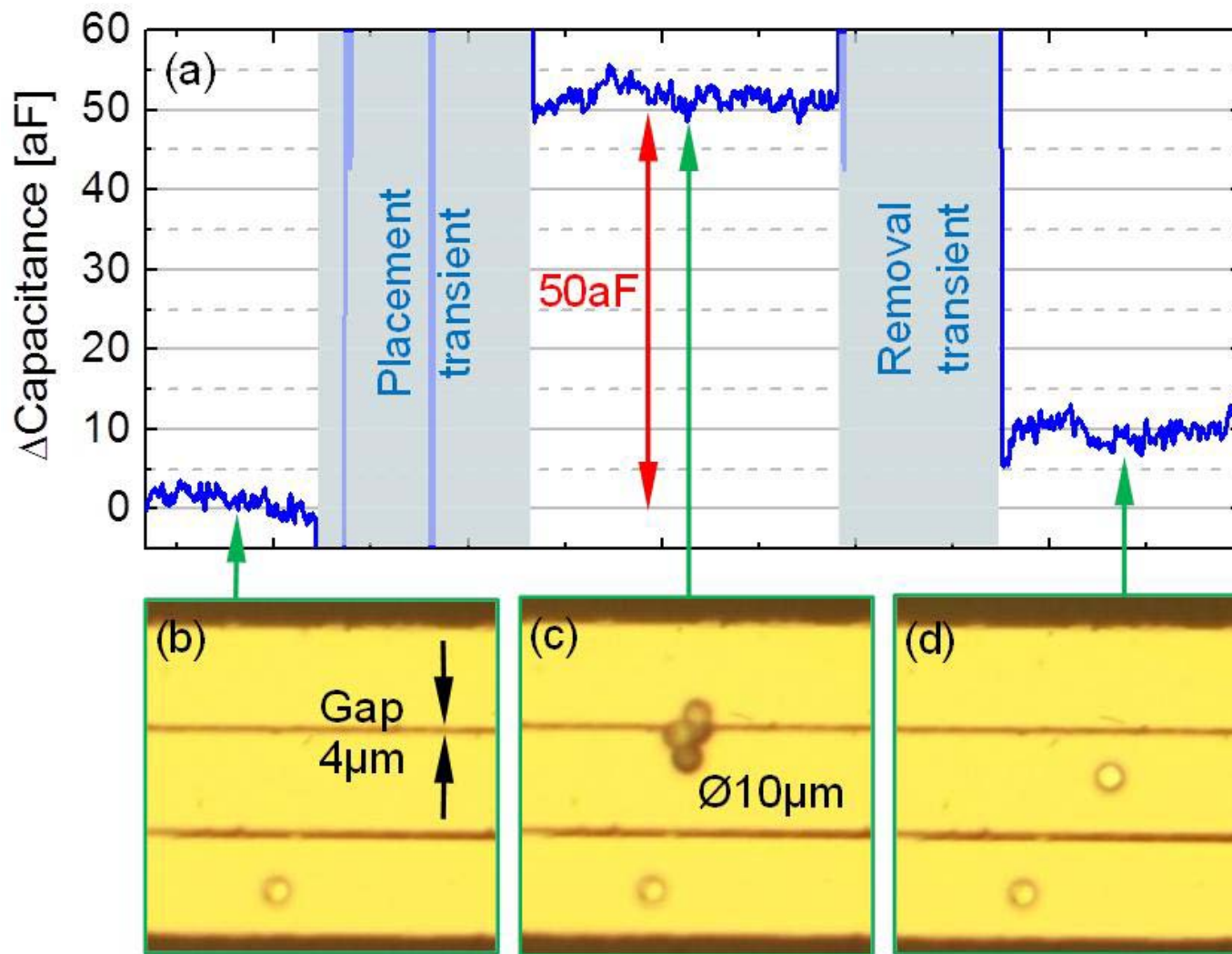


- Low-noise custom front-end (integrator)
- Differential sensing configuration to reduce generator noise
- 1.1 aF resolution with 1 sec response time (5 aF with 1 ms)



Static Characterization: PM_{10} Detection

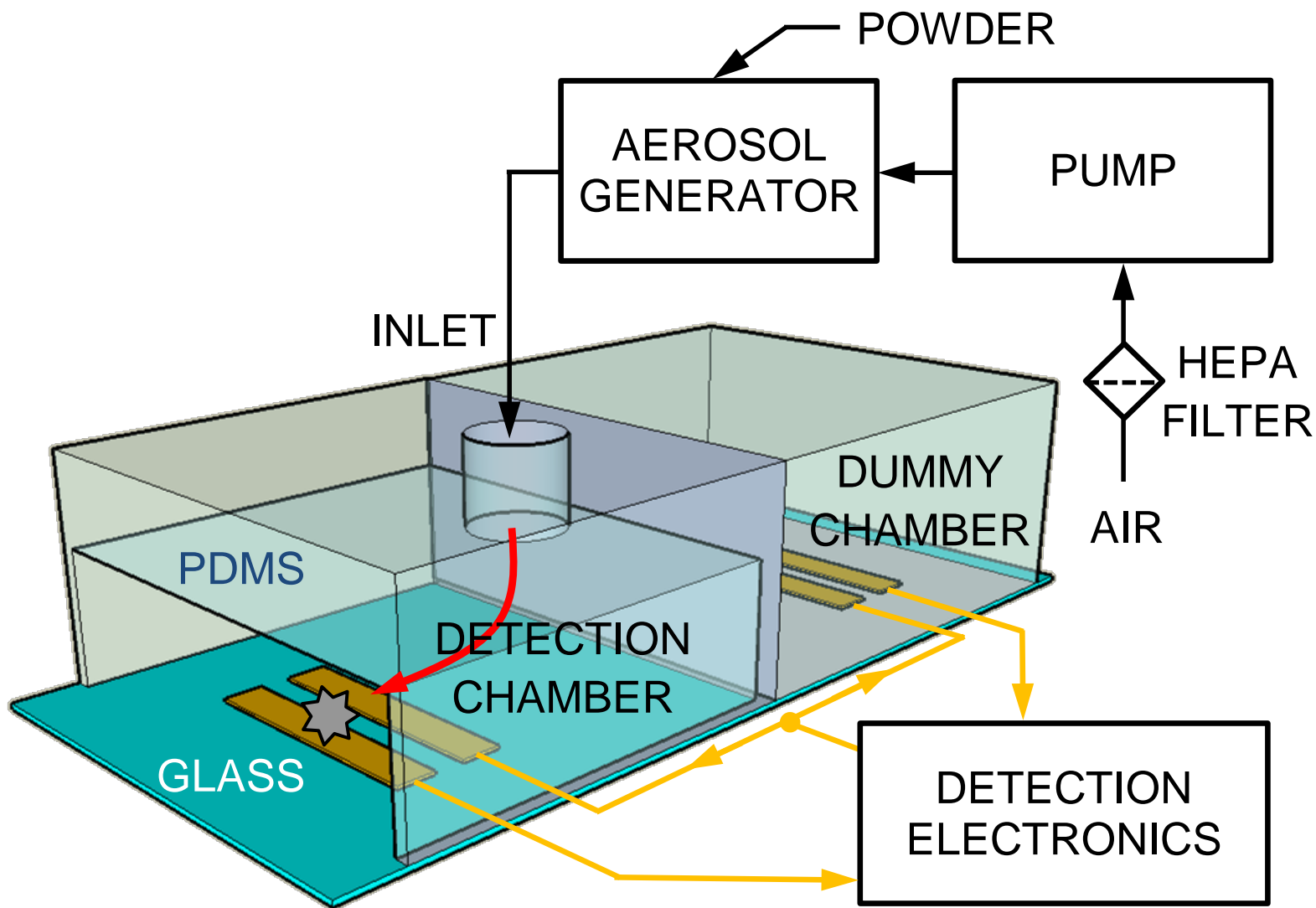
Validation with calibrated polystyrene $10\mu\text{m}$ beads



ΔC consistent with simulations

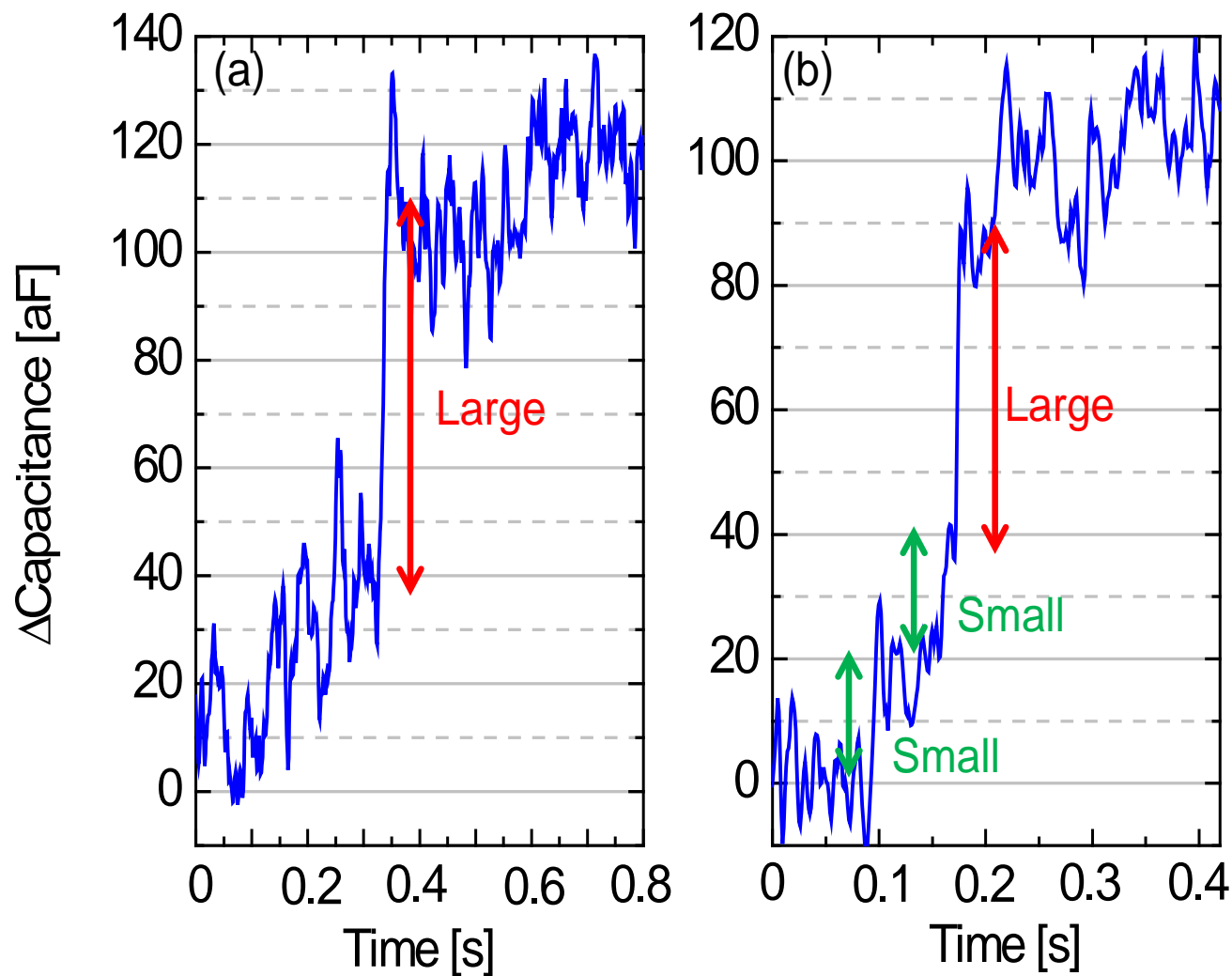


Full Detection System





PM₁₀: Talc Detection

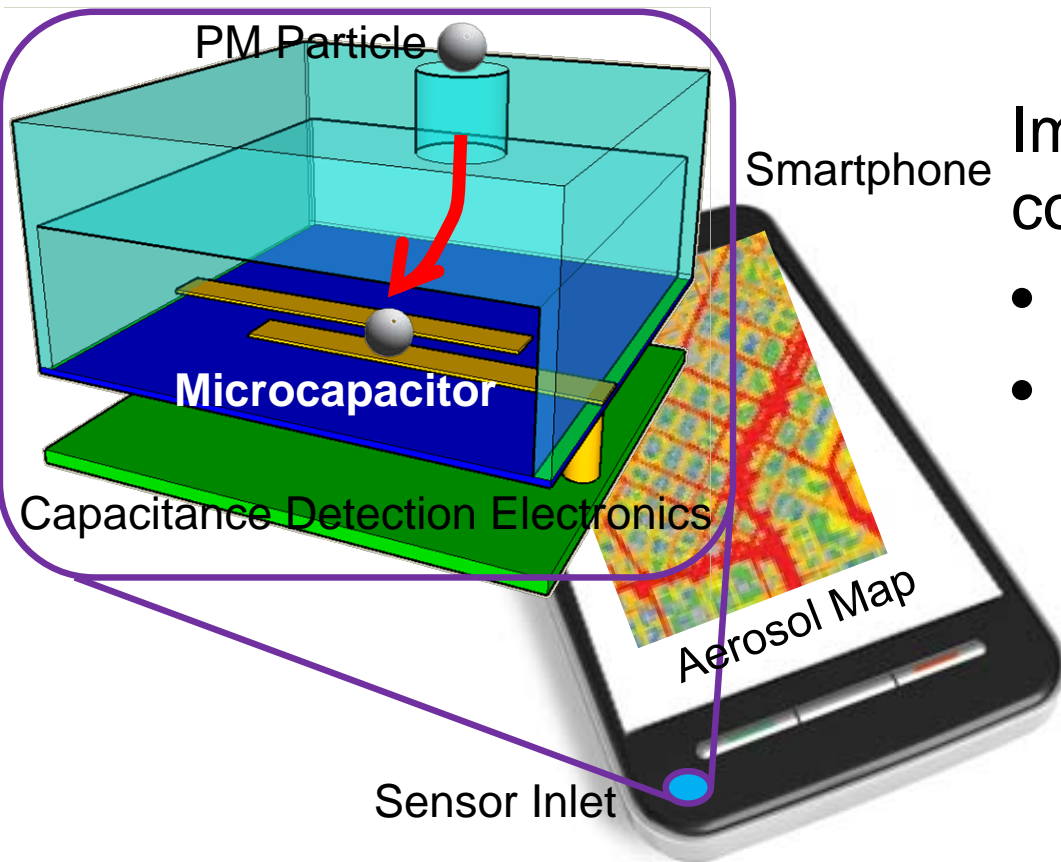


Real-time deposition tracking ($\Delta t = 10\text{ms}$) $\epsilon_r = 2.2$





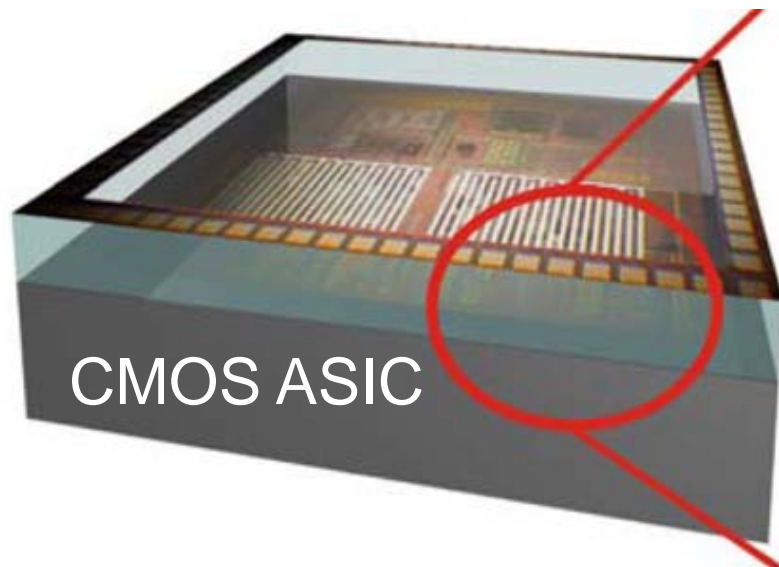
Single-Chip Detector: Towards Smartphones



Improving the resolution by combining **on the same chip**:

- Scaled lithography
- Integrated electronics with ZeptoFarad resolution

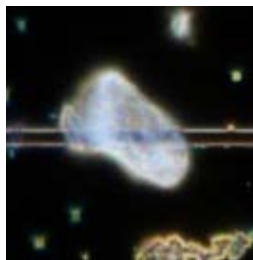
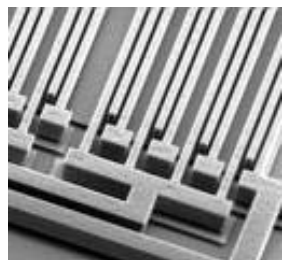
Targeting 1 μm PM



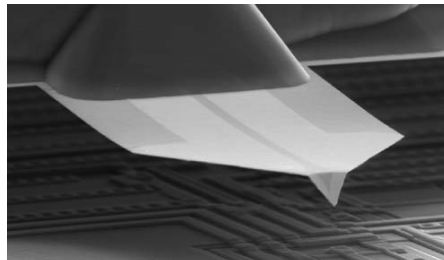


Cap. Resolution: Spanning Orders of Magnitude

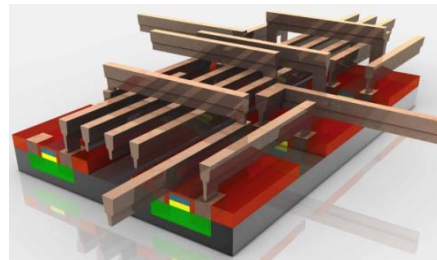
MEMS Microsensors



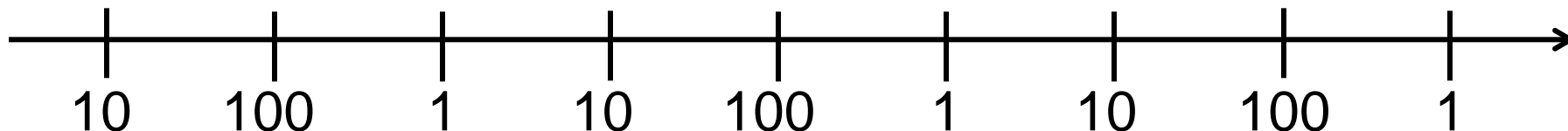
AFM



CMOS Parasitics



COTS



zepto

atto

femto

pico

Dedicated ASIC

Dedicated discrete-components

General purpose instrumentation

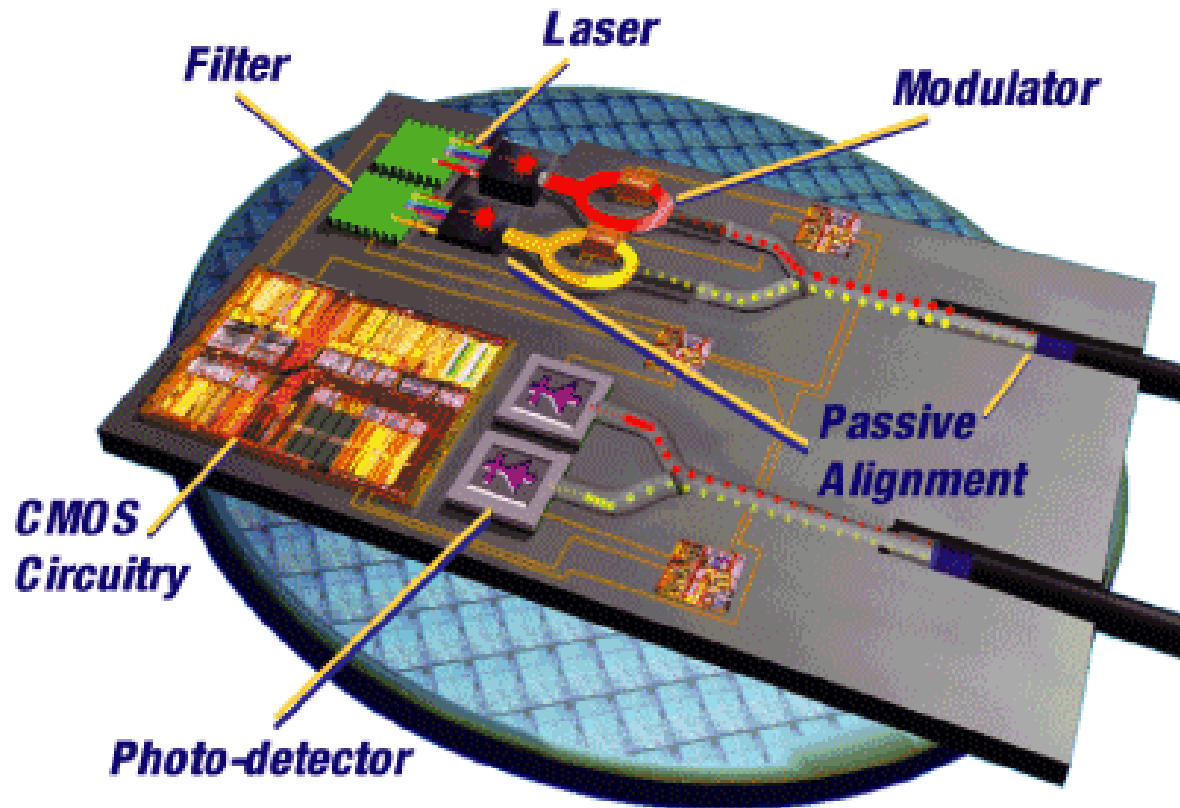
	σ [zF]	BW [Hz]	V_{AC}
Analog Devices	12	0.1	10.5
M.Carminati <i>et al.</i>	5	1.6	6

M. Carminati *et al.* "ZeptoFarad capacitance detection with a miniaturized CMOS current front-end for nanoscale sensors", *Sens. Act. A*, **172** 2011.





Integrated Photonics: Context and Motivation



Silicon Photonics:
great promise for
communications and
sensing

but

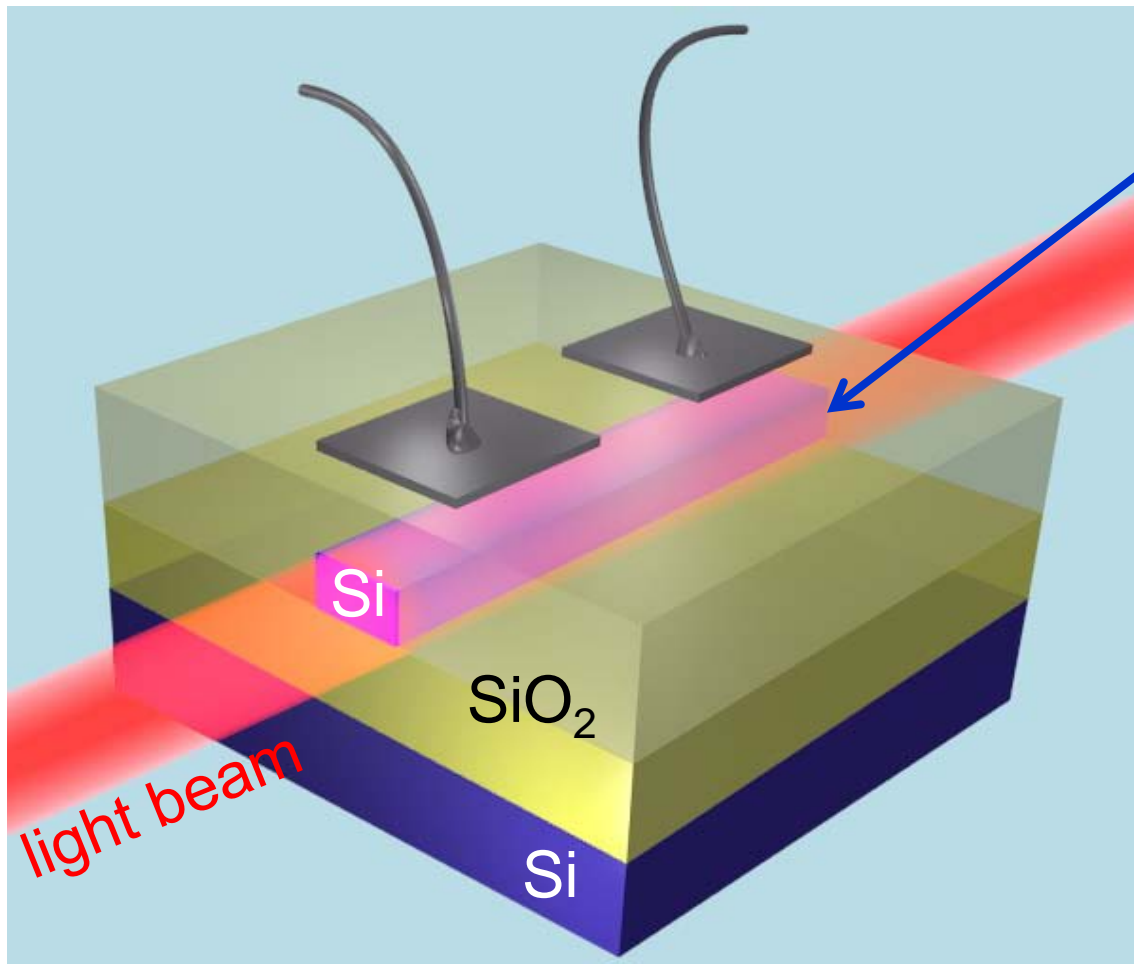
Large-scale integration of several components is limited by:

- lack of a consolidated/standard fabrication approach
- lack of **local feedback** to counterbalance drifts, variations
- limited integration with CMOS circuits

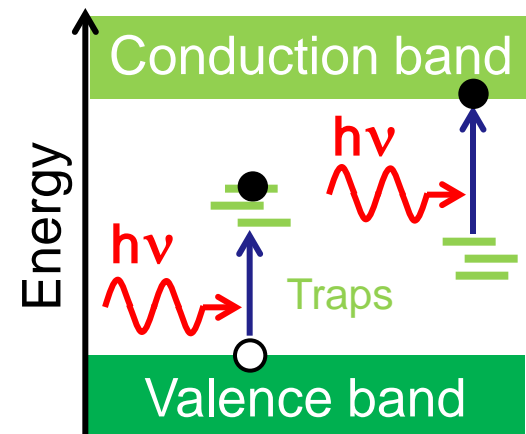


A Novel Transparent Light Monitoring Technique

At $\lambda = 1550\text{nm}$ the Si waveguide is **transparent** but



defects at the Si/SiO₂ interface produce surface states



Photogenerated carriers

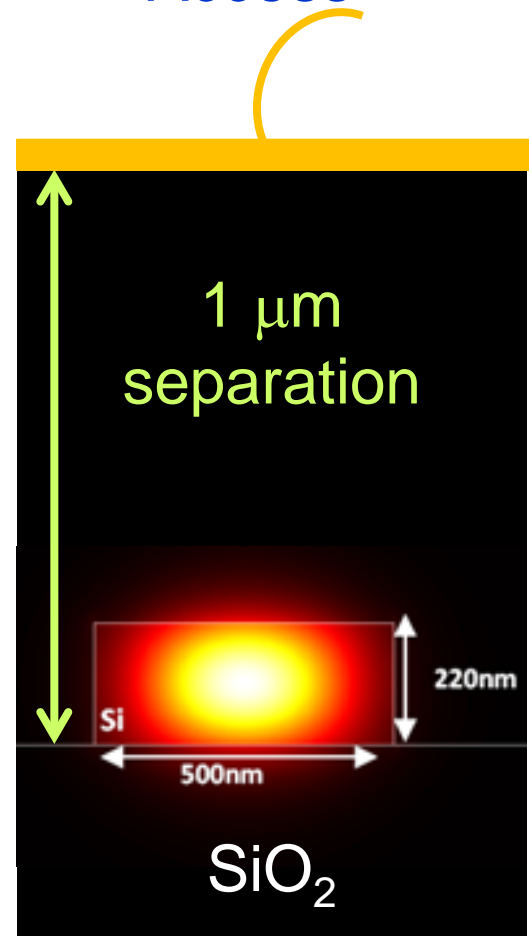
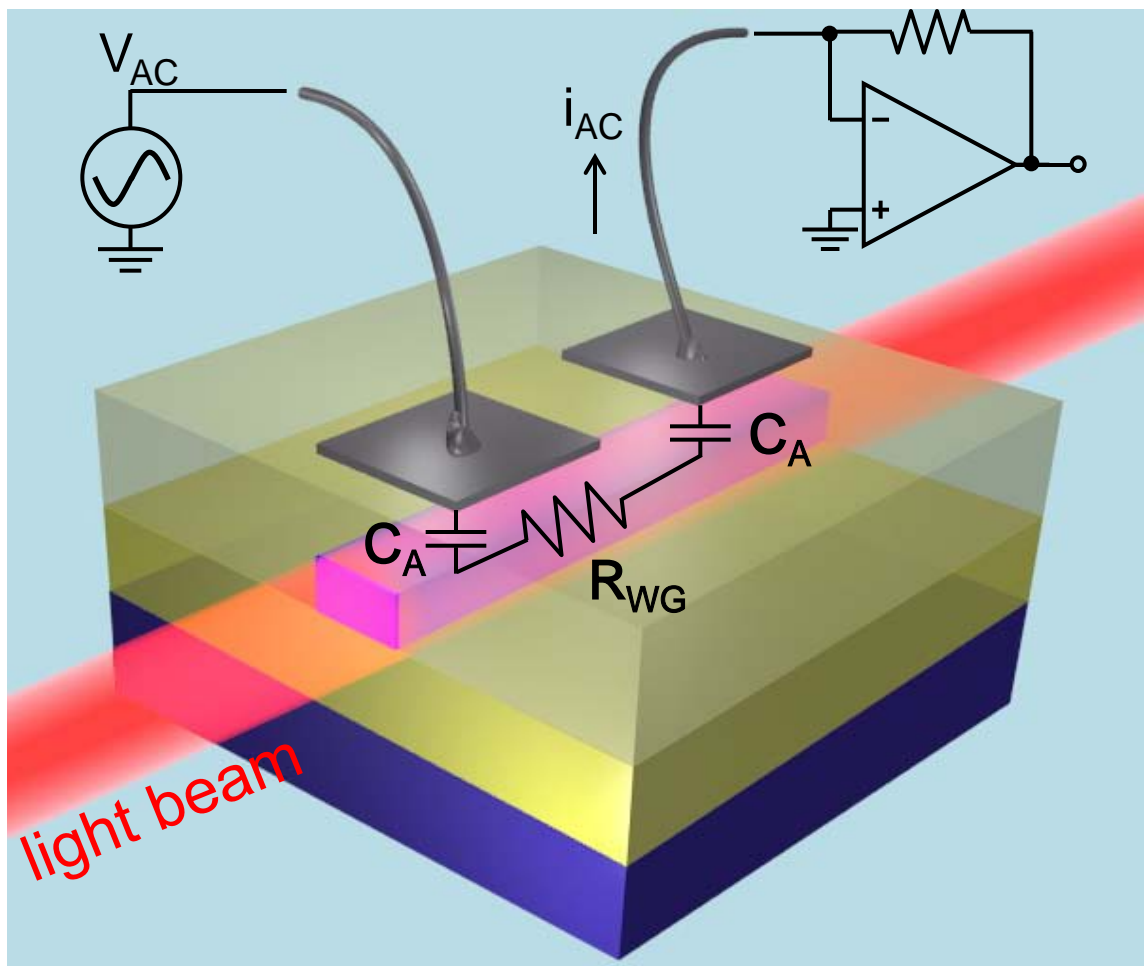
F. Morichetti et al., *IEEE J. Sel. Top. Quant. Electron.*, **20** (2014) .



A Novel Transparent Light Monitoring Technique

To avoid perturbation of the radiation

ContactLess Capacitive Access



F. Morichetti et al., *IEEE J. Sel. Top. Quant. Electron.*, **20** (2014) .

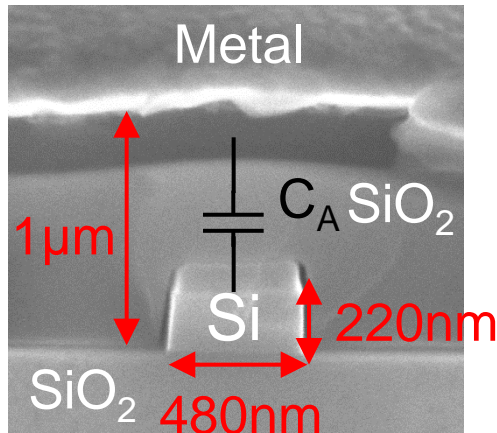
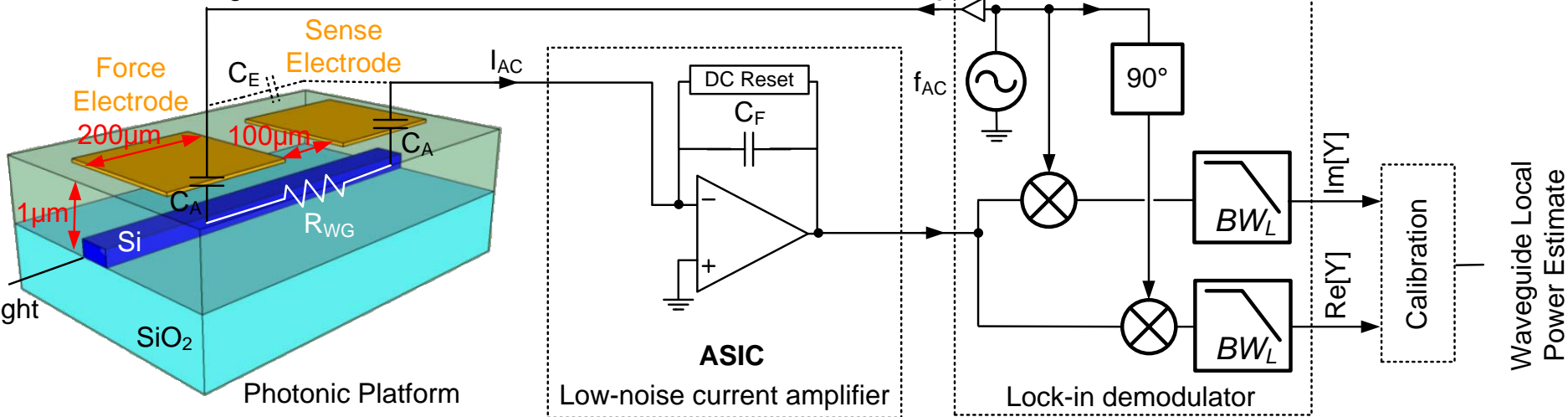
PATENTED!



Detection Architecture

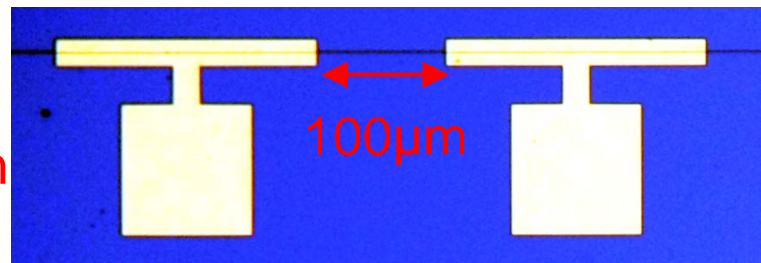
Low-noise impedance tracking platform

ContactLess Integrated Photonic Probe



Cross section

Optimized electrode layout



Top view

- $R_{WG} \sim 150M\Omega$
- $\Delta G \sim nS$
- $C_A = 5fF$
- $C_E = 500fF$



Non-Invasivity Enables Multipoint Monitoring

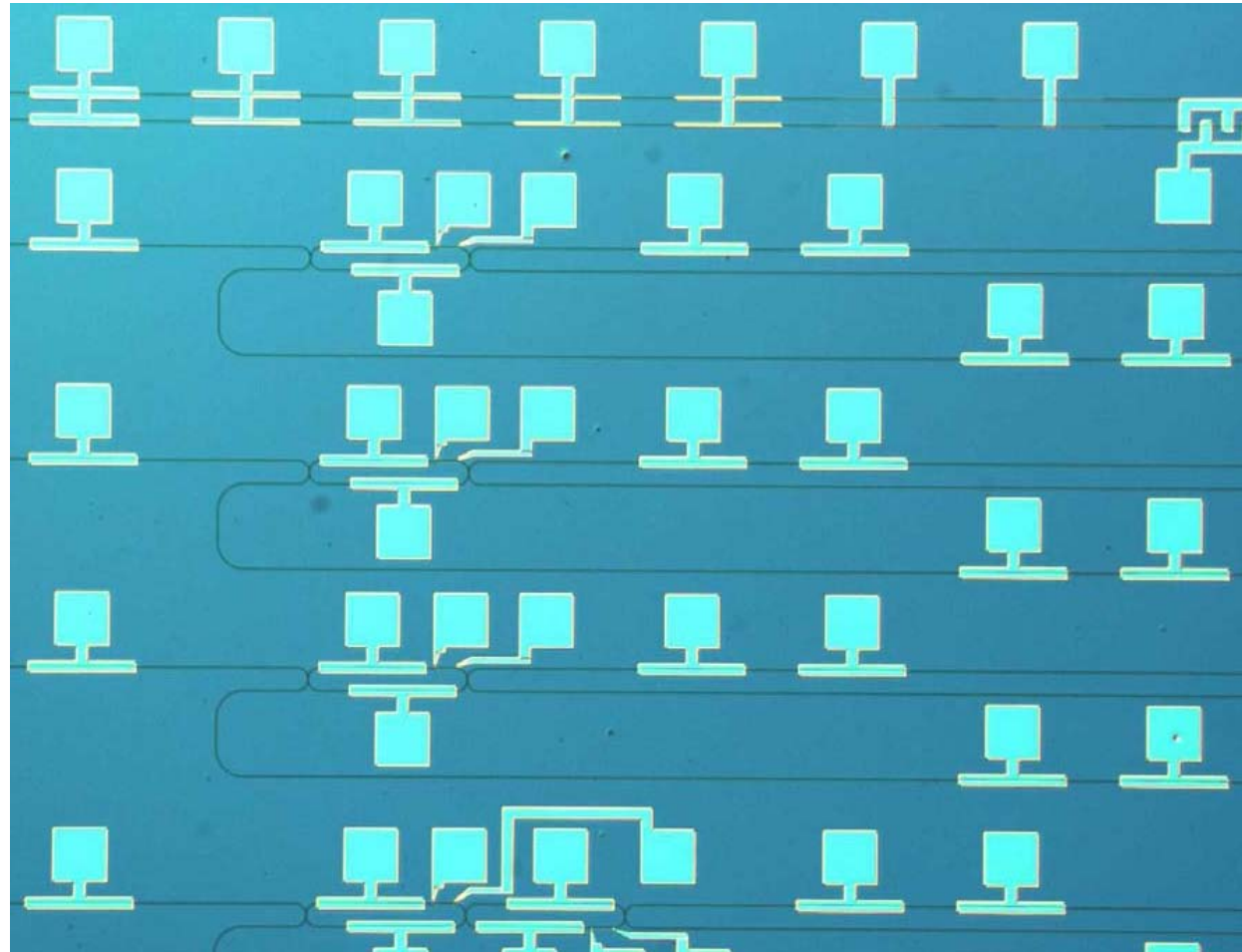
No extra light absorption



Multisite monitoring in complex circuits

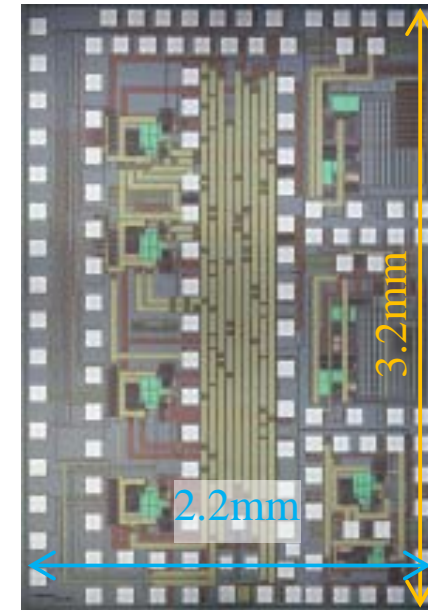
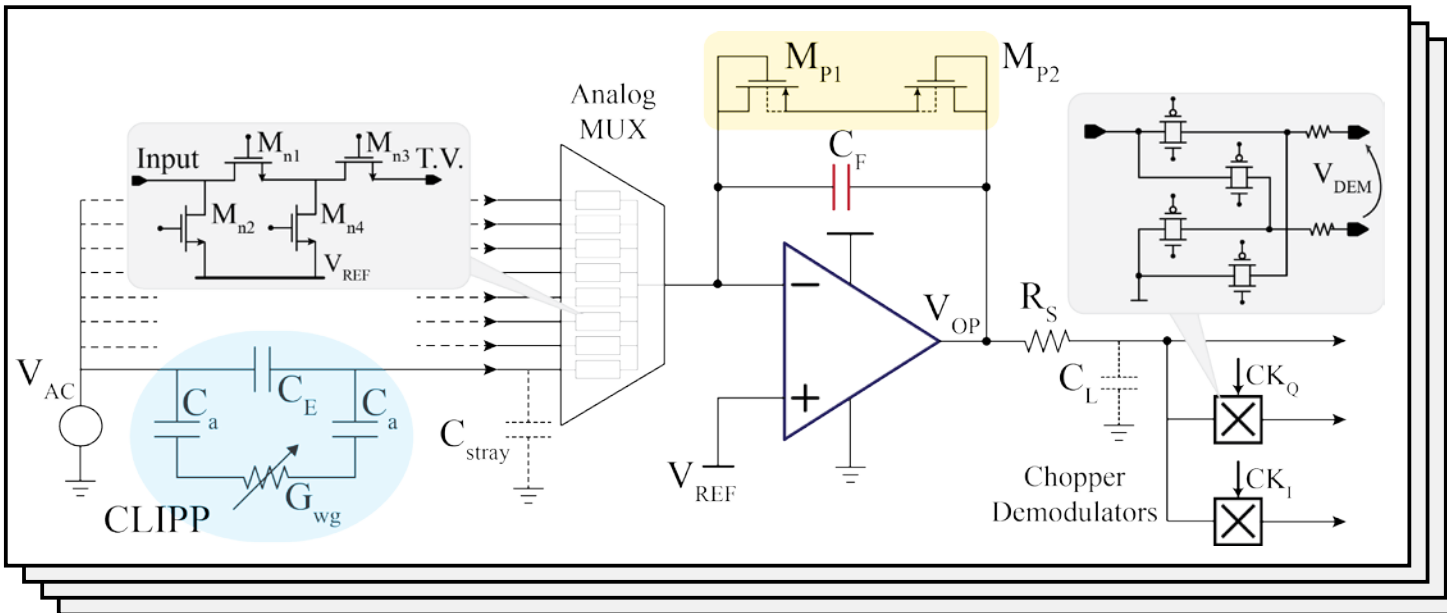


Multichannel ASIC is required





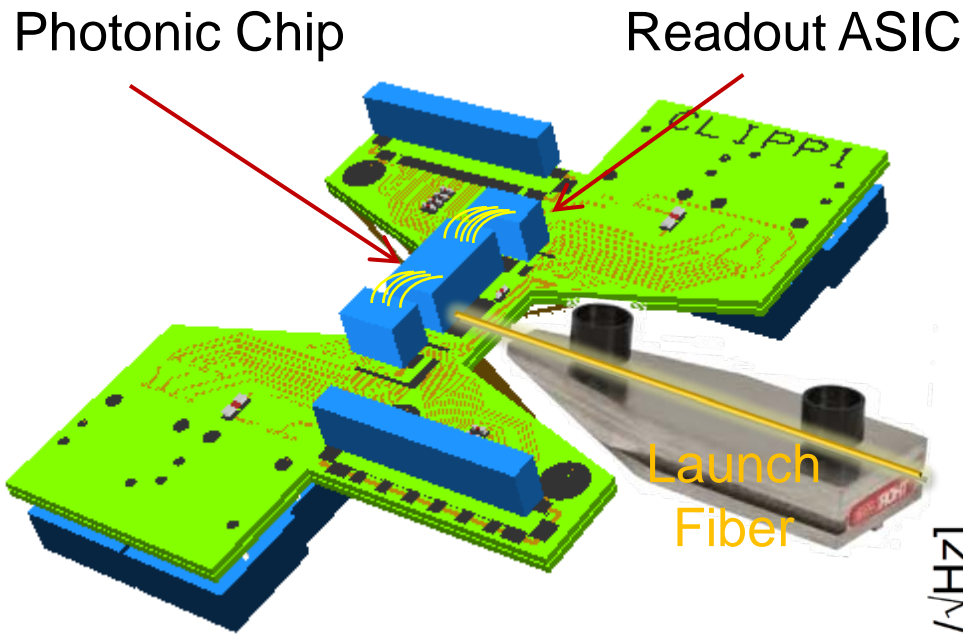
ASIC Design



- Integrator front-end ($C_F = 1\text{pF}$)
- DC handling network with sub-threshold transistors
- 8-channel low-parasitics MUX
- 2 square-wave multipliers
- 32 channels (4 parallel acquisition chains)
- 3.3V, 8mA current consumption, AMS 0.35 μm CMOS



Performances of the Integrated Setup

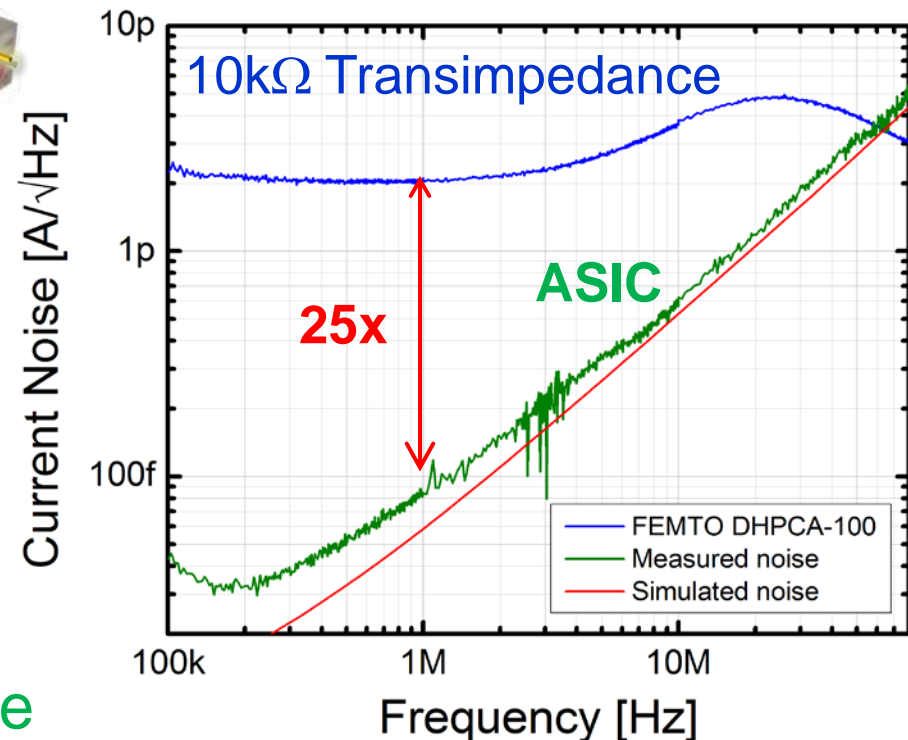


Radical **reduction of parasitic capacitance**

C_E of connections
from 500fF to 2fF

Advantages of integration:

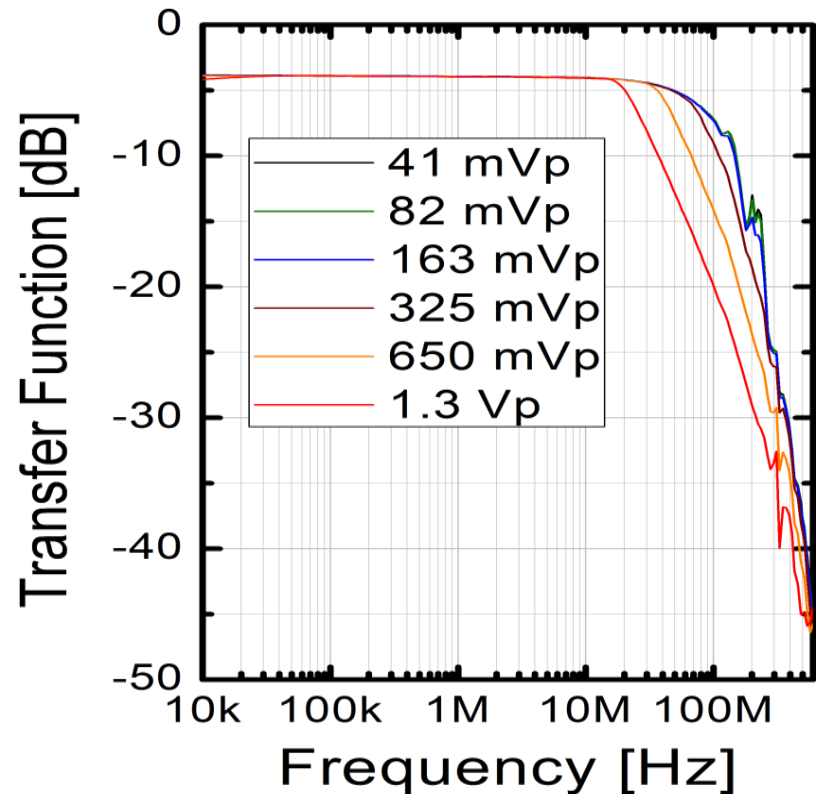
- Multipoint monitoring
- Miniaturization
- CMOS compatibility
- Reduction of $C_E \rightarrow$ **lower noise**



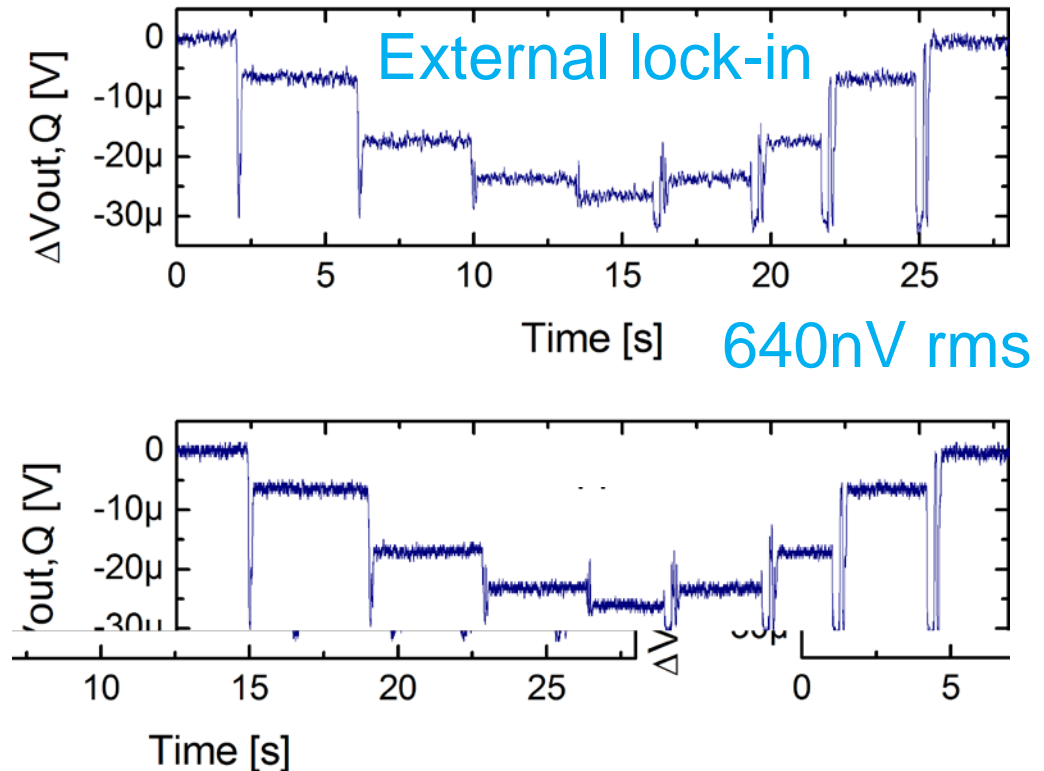


ASIC Characterization

100MHz bandwidth



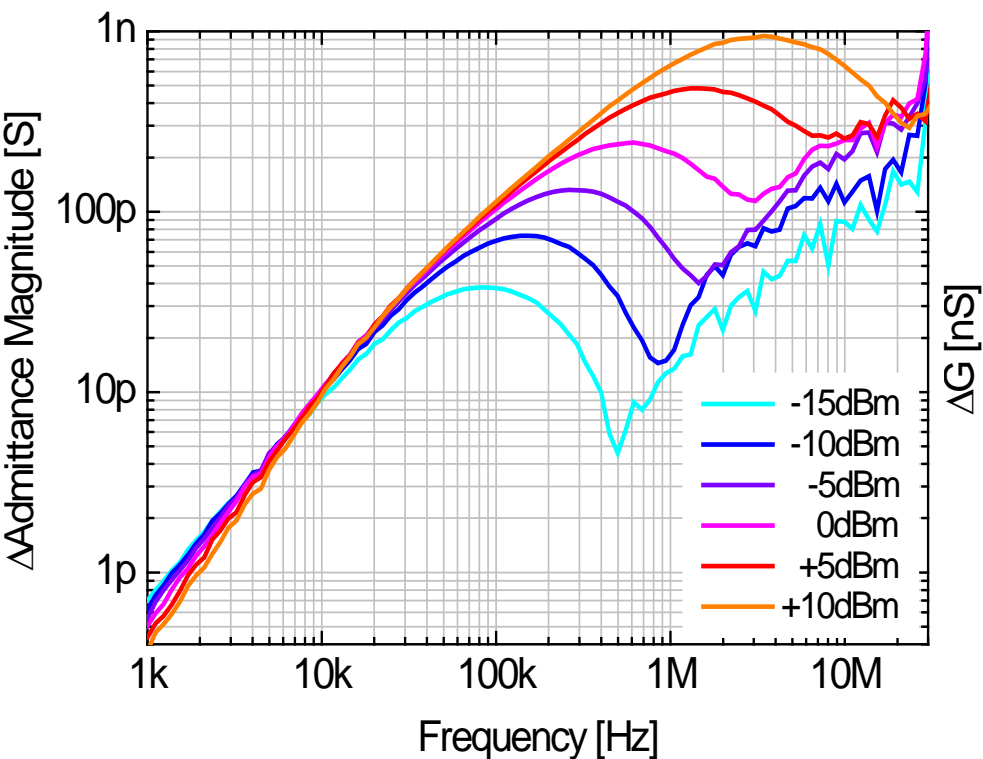
Same SNR of external lock-in





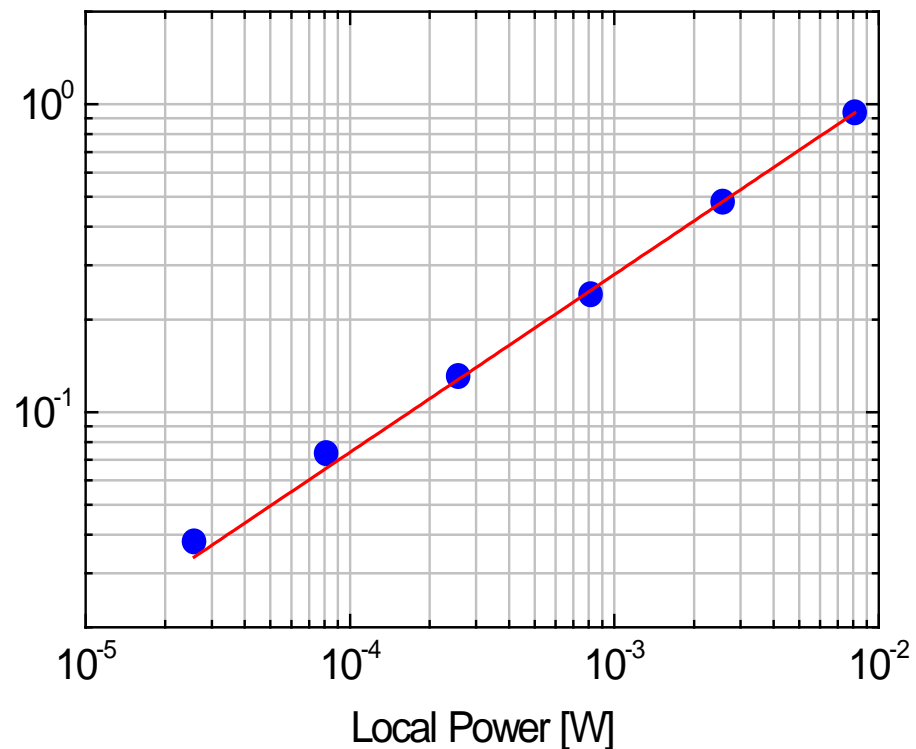
CLIPP Characterization: Power Monitor

Admittance spectroscopy for a wide range of optical power levels



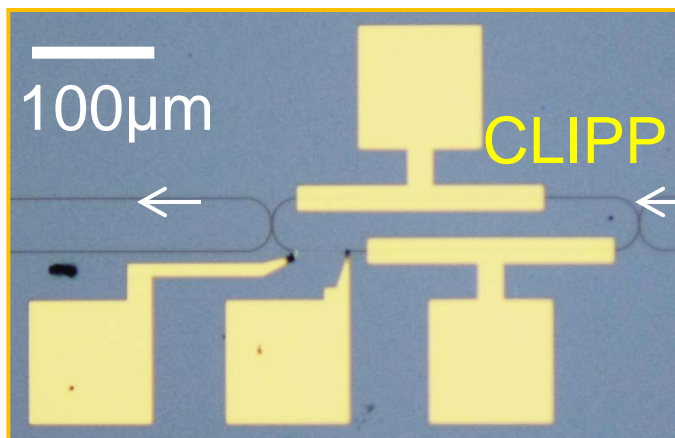
Monitor response curve

$$\Delta G \propto P^{0.6}$$



Min power: -30dBm, 40dB dynamic range

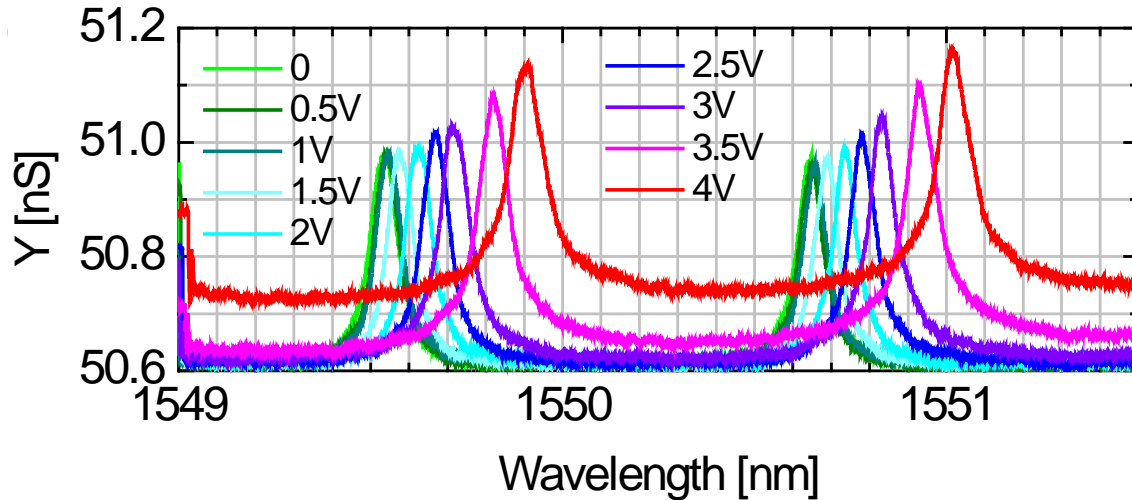
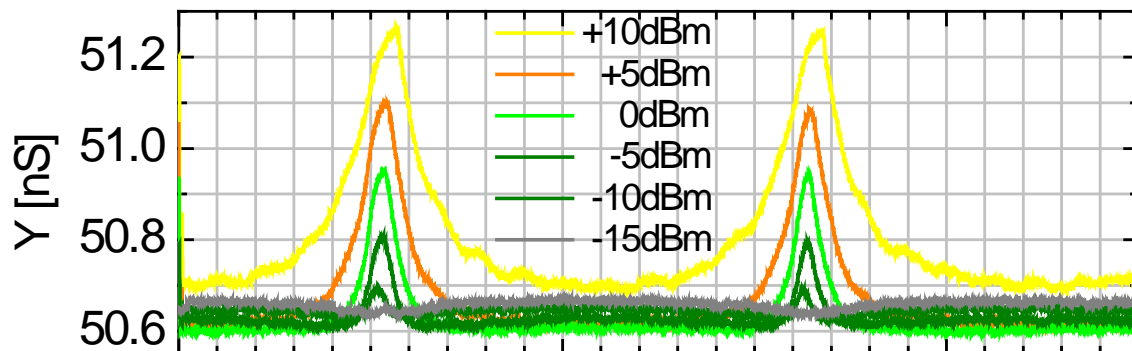
Looking Inside a Ring Resonator



Thermo-optic
Actuator

No perturbative effect
of V_{AC} on the
resonance!

Tracking input power

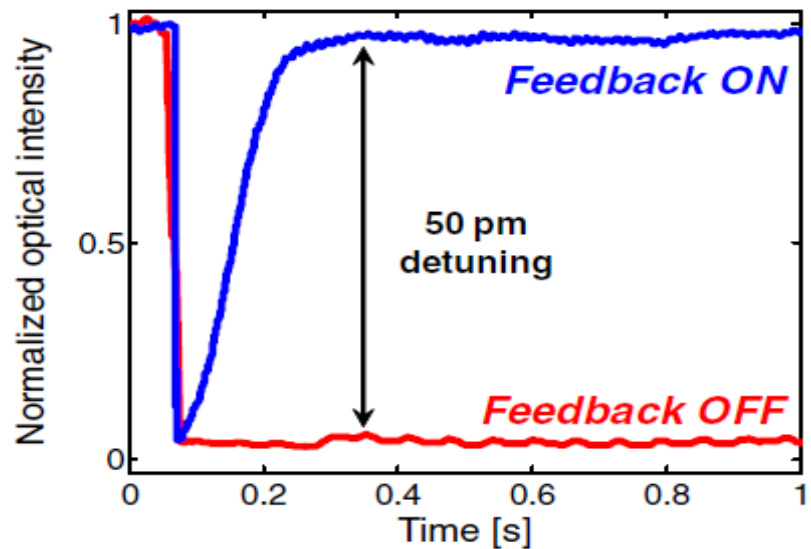
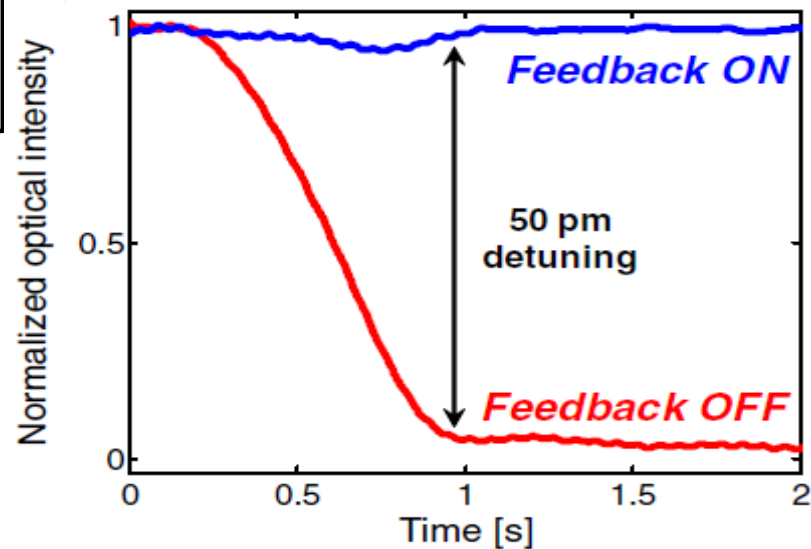
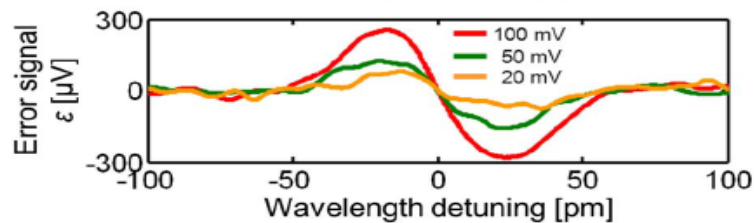
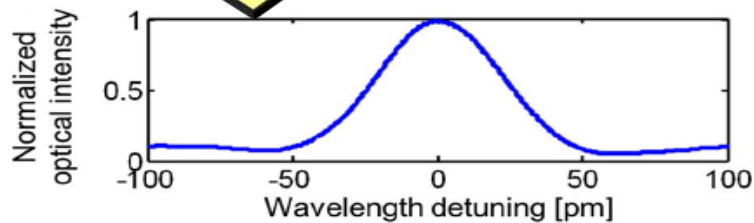
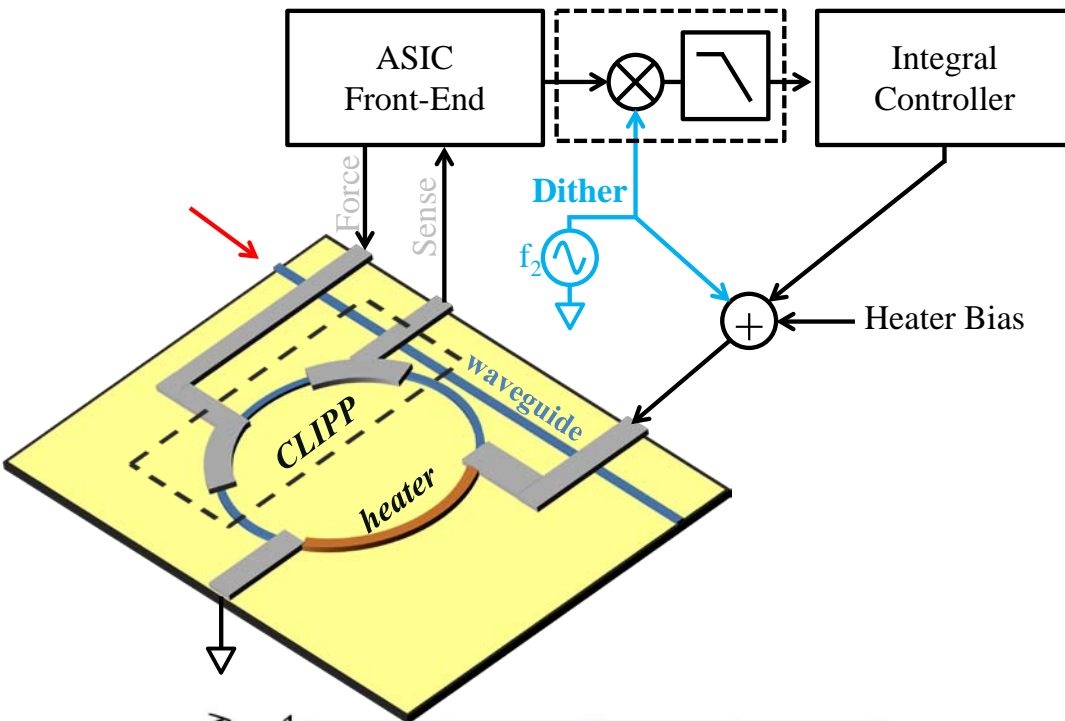


Tracking actuator voltage



Closing the Loop

S. Grillanda et al., *Optica.*, 1 3 (2014)





Summary

- Not all the impedance measuring techniques are equivalent in terms of noise and suitability for impedance spectroscopy.
- When measuring impedance at the **microscale**, the signals decrease and **noise** (unavoidable property of the electronic components) becomes relevant.
- The **current sensing** front-end coupled with a **lock-in** demodulator represents the best solution.
- The input capacitance must be **minimized**, paying attention to short coax cables and preferably insulating substrates.
- The **transimpedance amplifier** allows neutralization of the input stray impedance, but is prone to a noise/bandwidth trade-off, that can be relaxed by means of advanced topologies such as the integrator/differentiator cascade.



References

- M. Carminati, G. Ferrari, D. Bianchi, M. Sampietro, **Impedance spectroscopy for biosensing: circuits and applications**, in *Handbook of Biochips* (Editor M. Sawan), Springer *in press*.
- M. Carminati *et. al.* **“Theoretical and Experimental Comparison of Microelectrode Sensing Configurations for Impedimetric Cell Monitoring”** pp. 75-82, in *Lecture Notes on Impedance Spectroscopy: Vol. 4*, CRC Press, 2013.
- M. Crescentini, M. Bennati, M. Carminati, M. Tartagni, **“Noise Limits of CMOS Current Interfaces for Biosensors: a Review”**, *IEEE Trans. Biomed. Circuits and Systems*, vol. 8, no. 2, pp. 278-292, 2014.
- M. Carminati, G. Ferrari, D. Bianchi, M. Sampietro, **“Femtoampere integrated current preamplifier for low noise and wide bandwidth electrochemistry with nanoelectrodes”** *Electrochimica Acta*, vol. 112, pp. 950–956, 2013.
- M. Carminati, M. Vergani, G. Ferrari, L. Caranzi, M. Caironi, M. Sampietro **“Accuracy and resolution limits in quartz and silicon substrates with microelectrodes for electrochemical biosensors”**, *Sens. Act. B*, vol. 174, pp. 168-175, 2012.
- M. Carminati, G. Ferrari, F. Guagliardo, M. Sampietro, **“ZeptoFarad capacitance detection with a miniaturized CMOS current front-end for nanoscale sensors”**, *Sens. Act. A*, vol. 172, pp.117-123, 2011.
- M. Carminati, G. Ferrari and M. Sampietro, **“attoFarad resolution potentiostat for electrochemical measurements on nanoscale biomolecular interfacial systems”**, *Rev. Sci. Instrum.*, vol. 80, no. 12, pp. 124701/1-10, 2009.



Acknowledgments

Co-workers:

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S. Grillanda, F. Morichetti, A. Melloni

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Progetto **MINUTE**

and by EU under FP7 project «BBOI»

Breaking the Barriers of Optical Integration



fondazione
cariplo

Minute

