

Low temperature assembly method of microfluidic bio-molecules detection device

Stéphanie van Loo^a, S. Stoukatch^a, N. van Overstraeten-Schlögel^b, O. Lefèvre^c, F. Axisa^a, J. Destiné^a, D. Flandre^b, and P. Mertens^c

^a EMMI/Microsys lab, Université de Liège, Belgium

^b ICTEAM, Université catholique de Louvain-la-Neuve, Belgium

^c Coris BioConcept, Gembloux, Belgium

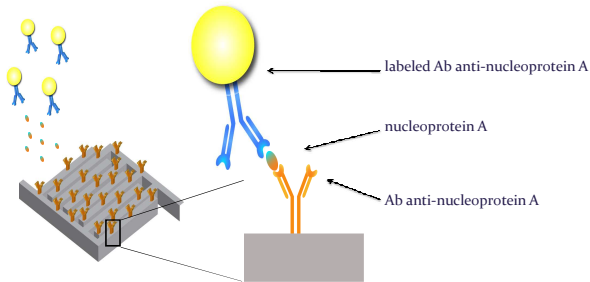
Detection principle of the microfluidic device

Detection target: nucleoprotein A (Influenza A virus proteins)

Sensor: interdigitated capacitive electrodes

Biofunctionalization: grafting of anti-nucleoprotein A antibodies on the electrodes

→ **Measurement:** the capacitance variation induced by the fixation of gold - labeled Ab anti-Influenza A on the sensing area



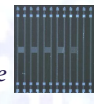
Assembly of the bio-sensor in DIL package

☹️ The bio-sensor is sensitive to contaminants, temperature above 37°C and UV exposure → *Need for alternative packaging method*

Die attach:

Standard: high T cure adhesive

→ *Alternative: RT cure adhesive*

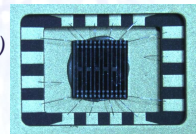


Interdigitated array micro-electrodes (IDAM): 4 sensors and 2 calibration structures, on a 3 x 3 mm² Si die

Wire bonding:

Standard: Au wire bonding (200°C)

→ *Alternative: RT Al wire bonding*



Wire bonded die in DIL package

Encapsulation:

Standard: high T curable encapsulant

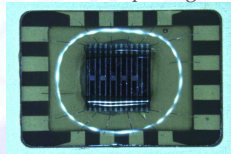
→ *Alternative: UV curable encapsulant*

• higher viscosity to define a sensing area

• lower viscosity to encapsulate the Al wires

UV intensity: 18.5 W/cm², wavelength:

320 - 500 nm, duration: max 20 s

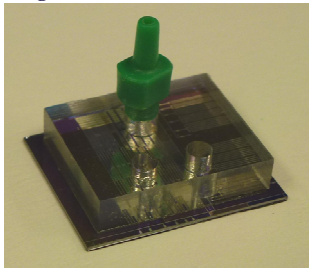


Encapsulated die in DIL package

Low temperature bonding of PMMA cover

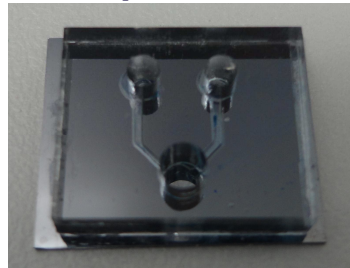
First step:

- dispensing of epoxy adhesive on the KMPR on the Si die at RT
- Deposition of the PMMA cover on the chip with visual alignment



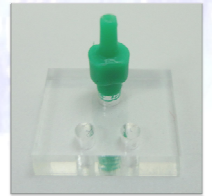
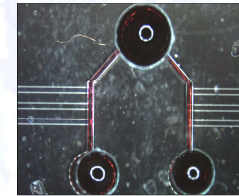
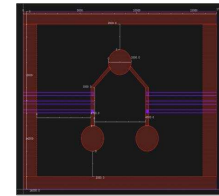
Second step:

- dispensing of epoxy adhesive on the PMMA, half cure (30 min at 60°C)
- PDMS stamp
- Deposition of the PMMA cover on the chip with visual



Integrated microfluidic chip

Integrated microfluidic chip with two sensing areas inside microfluidic channels for differential detection on silicon die with PMMA cover.



Description of the integrated chip:

- Silicon die: 17 x 14 mm²
- KMPR photoresist: 120 μm thick, 14.6 x 12 mm²
- 2 channels: 300 μm wide
- Transparent PMMA cover: 14.6 x 12 mm²
- 2 inlets, 1 outlet: 2 mm diameter

Bonding of the PMMA cover:

☹️ The conventional methods (thermal, plasma, solvent bonding etc) are not suitable with the bio-layer grafted on the sensor

→ Biocompatible low temperature adhesive

Assembly of the microfluidic chip on PCB

☹️ The bio-sensor in microfluidic chip is sensitive to temperature above 37°C and UV exposure → *Need for alternative packaging method*

Die attach:

Standard: high T cure adhesive

→ *Alternative: RT cure adhesive*

Wire bonding:

Standard: Au wire bonding (200°C)

→ *Alternative: RT Al wire bonding*

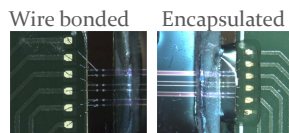
Encapsulation:

Standard: high T curable encapsulant

→ *Alternative: UV curable encapsulant*

UV intensity: 18.5 W/cm², wavelength:

320 - 500 nm, duration: max 20 s

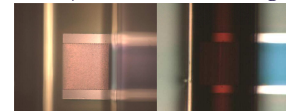


Mounted on PCB



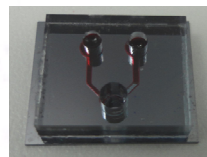
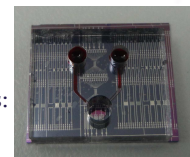
Results and conclusion

We injected food coloring to investigate the hermeticity :



Zoom on sensing area inside the channel. Without cover (left), with cover and coloring agent (right)

Food coloring in the channel confirmed gross-leak tightness:



We developed low temperature assembly methods which cause no damage to the biological layer grafted on the bio-sensor

Ref.:

N. Van Overstraeten-Schlögel et al., *Bio-Sensing Technology conf.*, 2011.

S. Stoukatch et al., *Plastic Electronics conf.*, 2011.