

Biosensori e riparazione tessutale cutanea

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Professor and Chairman

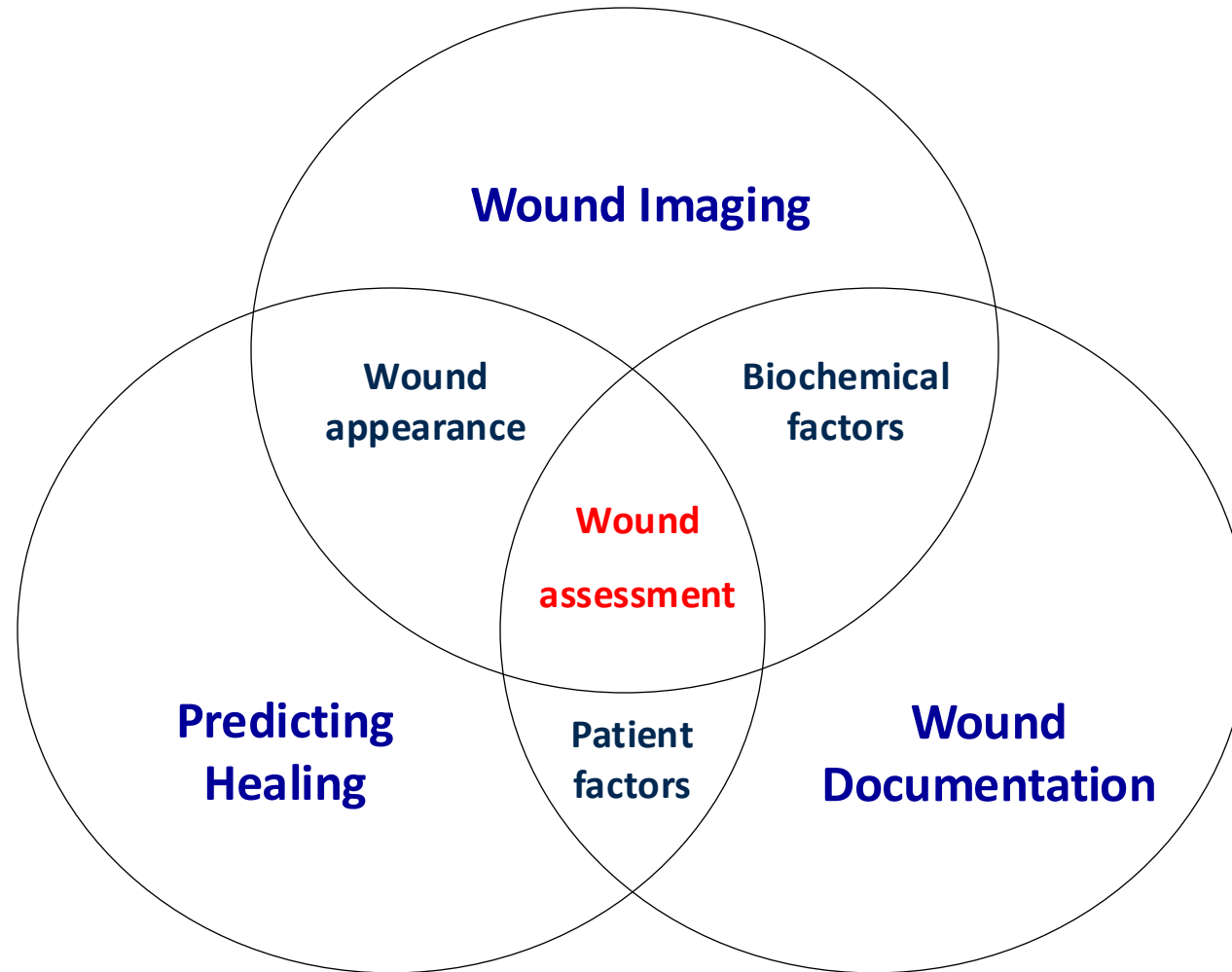
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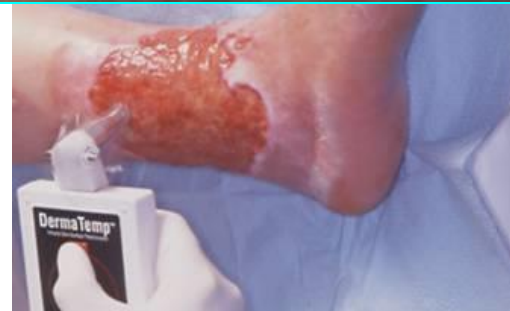
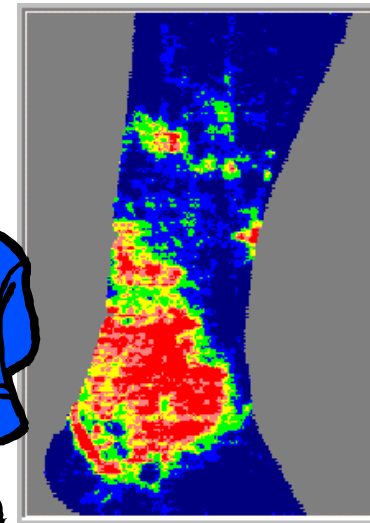
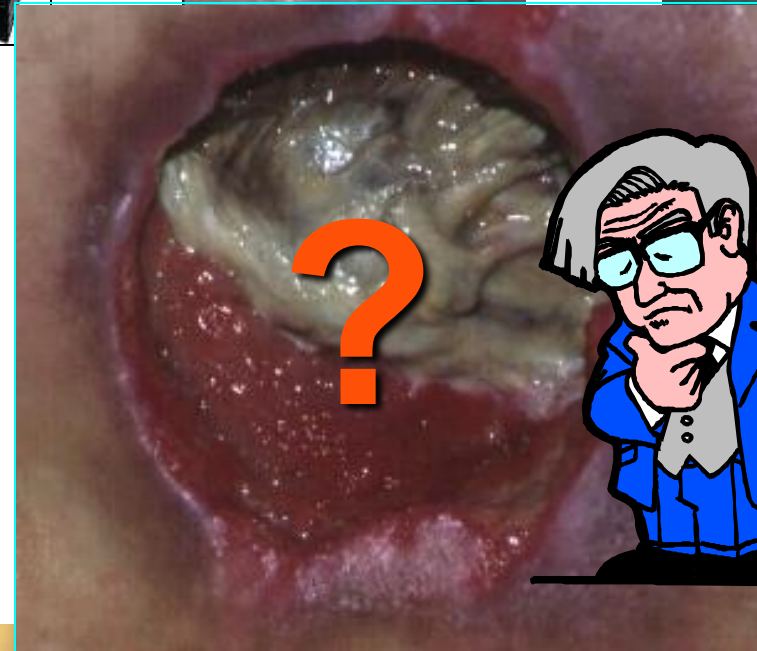


Disclosures

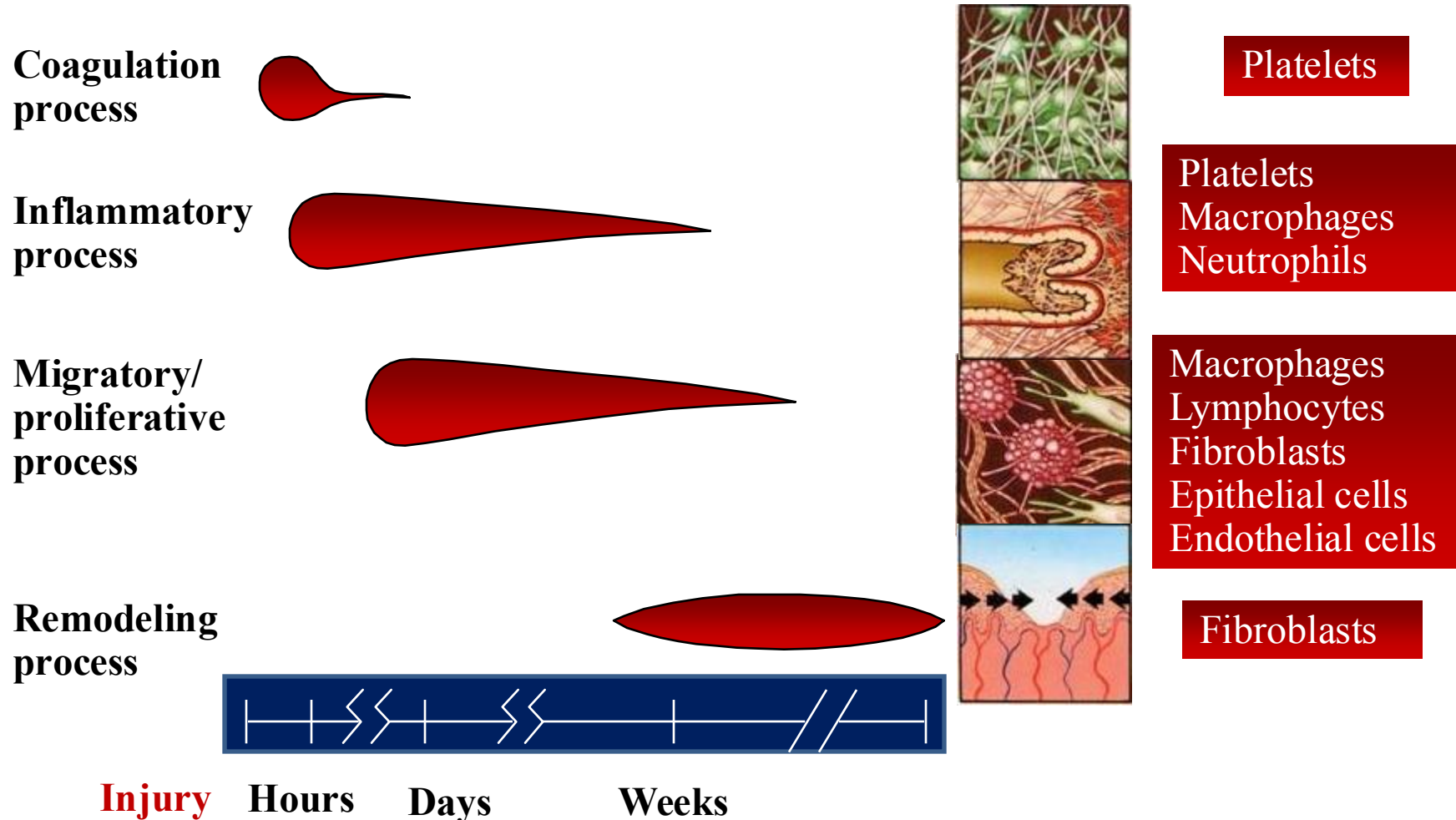
- Abbvie
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- Sanofi

Assessing Wound Healing

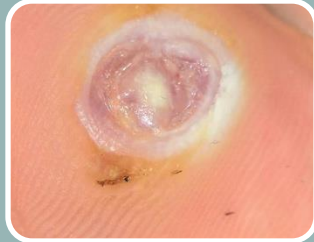




Wound Healing



Purpose of Wound Assessment



Better understanding of the processes which impair healing

- Diagnosis
- Clinical decision support



Outcomes

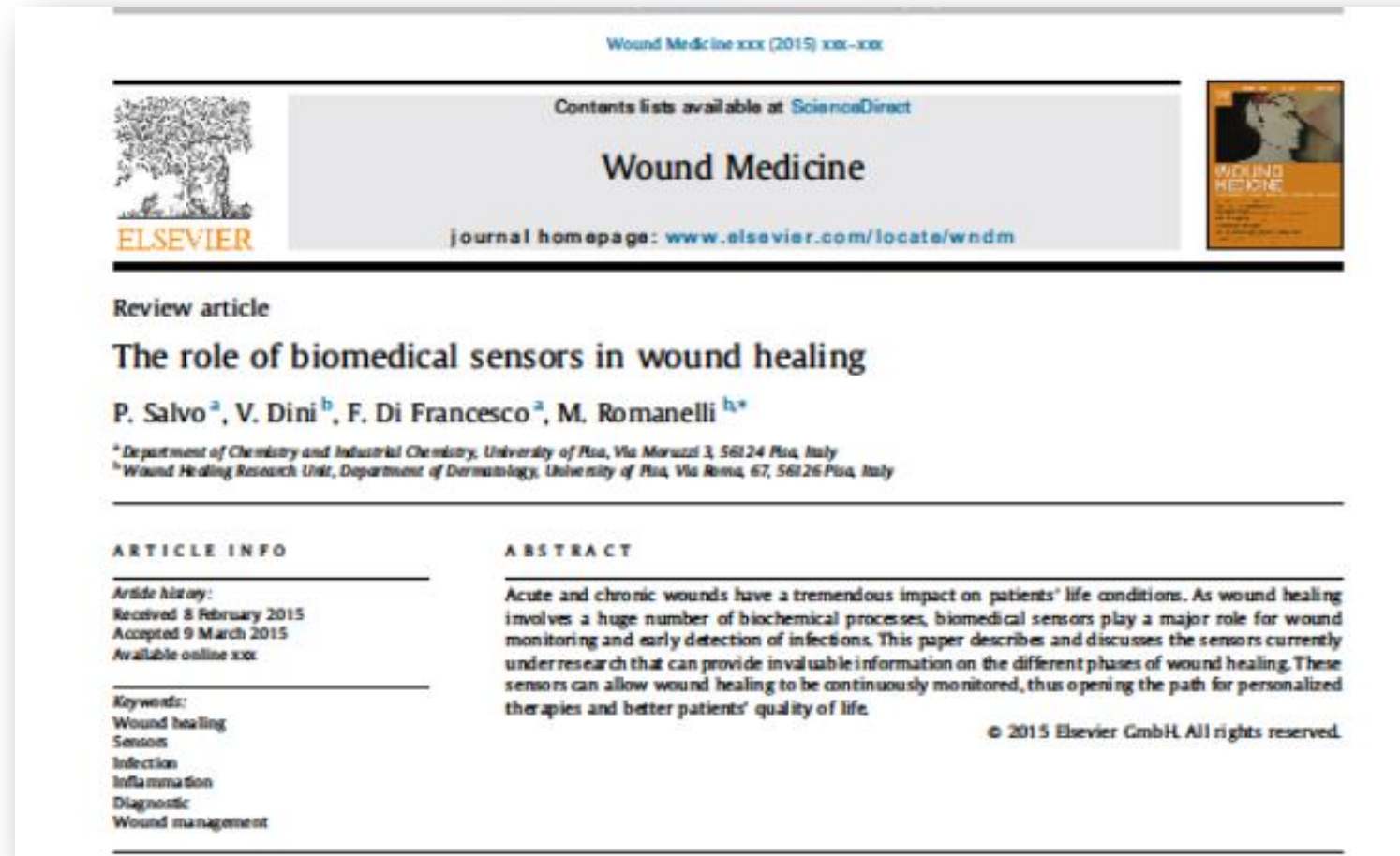
- Monitoring the effectiveness of treatment
- Prediction of outcome



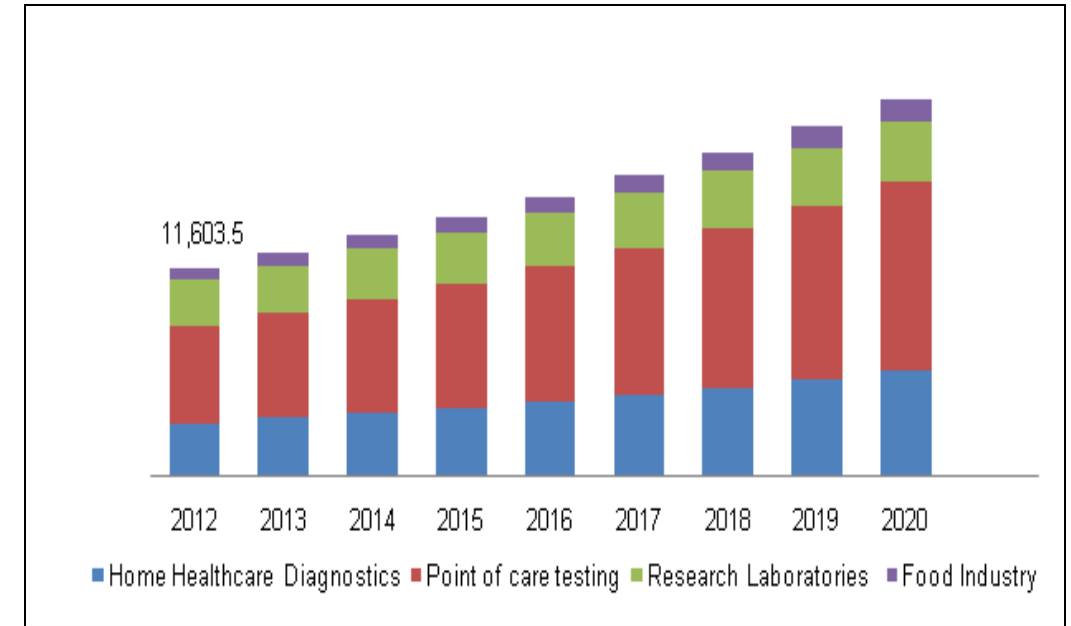
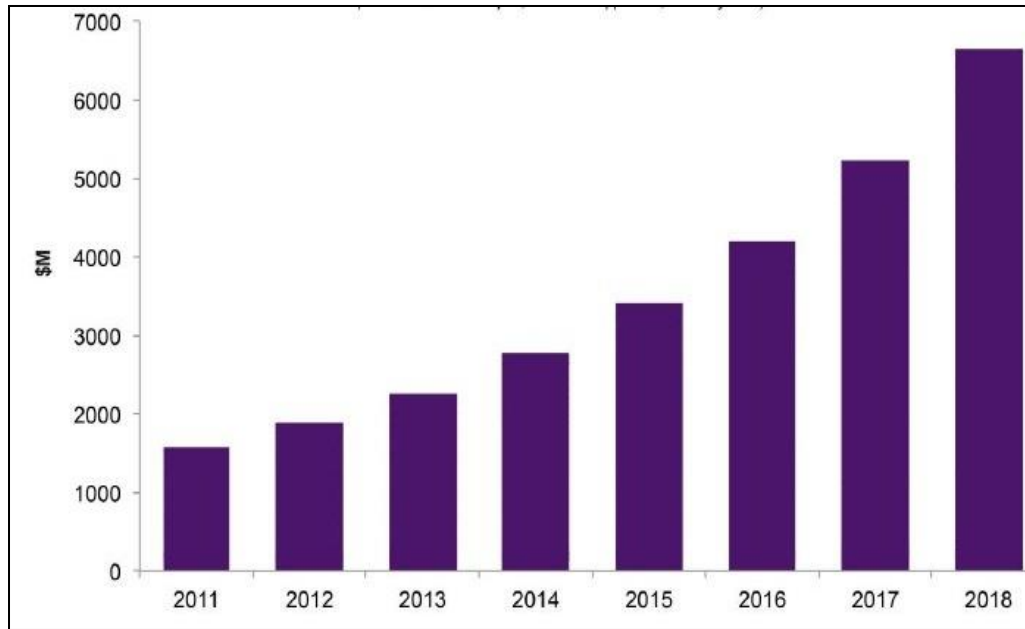
Prevention wound complications

- Amputation (DFU)
- Infection

The Role of Biomedical Sensors in Wound Healing



Sensors in the science market



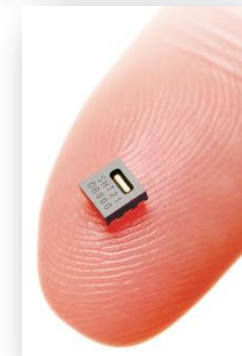
Sensors

A small device that detects and responds to some type of input from the physical environment and sends the information to other electronics.



Biosensors

Sensors which detect analytes thanks to a biological component (cells, protein, nucleic acid or biometric polymers).



Sensors must be :

- Thin and flexible
- Low cost, disposable
- Suitable for integration in wound dressing
- Biocompatible
- Able to resist in a chemically aggressive environment
- Able to monitor large areas



Flexible sensor chip

Wound Assessment Parameters



Temperature



pH



TEWL

International Journal of Nanomedicine

Dovepress

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ORIGINAL RESEARCH

Temperature- and pH-sensitive wearable materials for monitoring foot ulcers

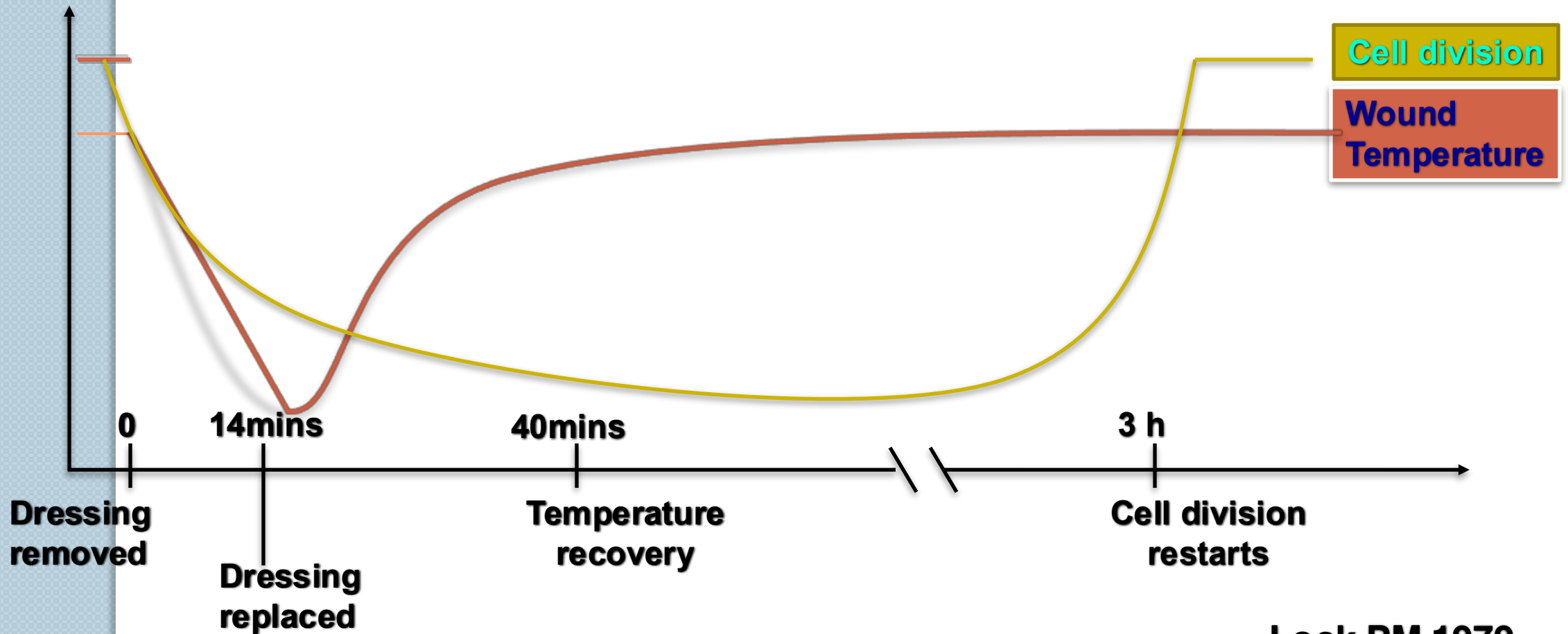
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Abstract: Foot ulcers account for 15% of comorbidities associated with diabetes. Presently, no device allows the status of foot ulcers to be continuously monitored when patients are not hospitalized. In this study, we describe a temperature and a pH sensor capable of monitoring diabetic foot and venous leg ulcers developed in the frame of the seventh framework program European Union project SWAN-iCare (smart wearable and autonomous negative pressure device for wound monitoring and therapy). Temperature is measured by exploiting the variations in the electrical resistance of a nanocomposite consisting of multiwalled carbon nanotubes and poly(styrene-*b*-(ethylene-*co*-butylene)-*b*-styrene). The pH sensor used a graphene oxide (GO) layer that changes its electrical potential when pH changes. The temperature sensor has a sensitivity of $\sim 85 \Omega/^{\circ}\text{C}$ in the range 25°C – 50°C and a high repeatability (maximum standard deviation of 0.1% over seven repeated measurements). For a GO concentration of 4 mg/mL,

The Role of Temperature in Chronic Wounds

Effect of Cooling on Wounds



The Role of Temperature in Chronic Wounds

- Increase in temperature is an established marker of infection
- Differences of $>4^{\circ}\text{F}$ were used as the trigger point of ulceration

Amstrong et al. 2007



Temperature sensors

- ✓ Resistance temperature detectors
- ✓ Thermocouples
- ✓ Thermistors
- ✓ Infra-red sensors
- ✓ Silicone-based sensors

Temperature Sensor

Skin temperature monitoring by a wireless sensor

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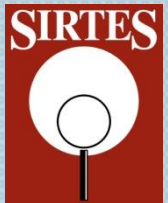
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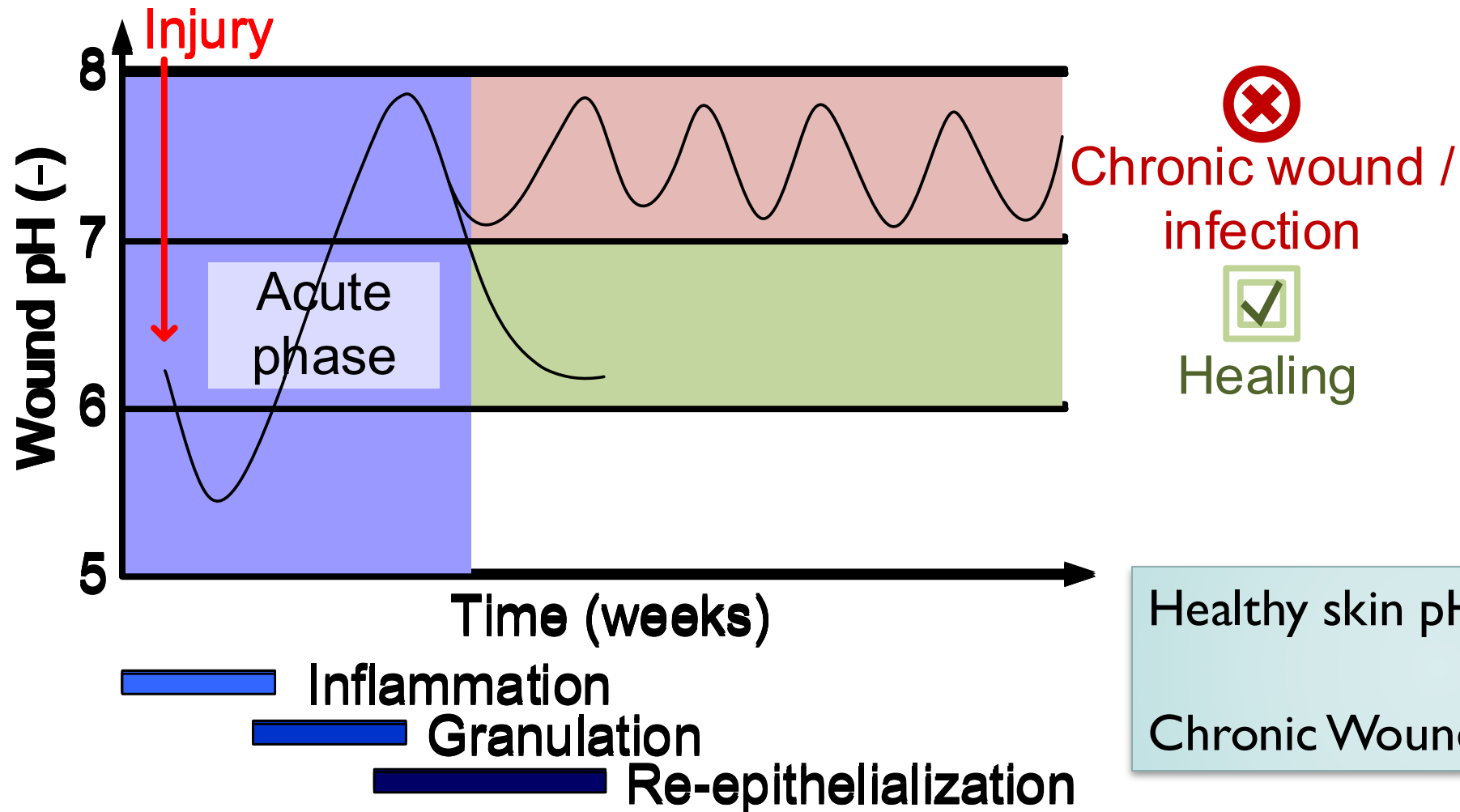
Abstract- The present paper reports the development of a temperature data logger based on a temperature sensitive resistive film and an RFID tag. Thanks to its capability of wireless communication, the device is going to be used for a minimally invasive remote monitoring of the skin temperature under a bandage or a wound dressing.

(SEBS, Europrene Sol TH 212) was supplied by Enichem Elastomeri and used as such. Toluene was supplied by Sigma Aldrich and used without any purification. A stock solution was prepared by dissolving SEBS (0.4 g) in toluene (100 mL). Aliquots of this solution (3 ml) were poured in vials containing 12 mg of MWCNTs, thus obtaining a 1:1 w/w

- Wireless
- A mixture of multi-walled carbon nanotubes and poly SEBS (styrene-b-ethylene-co-butylene-b-styrene) that coat a Kapton[®] substrate.
- The signal can be remotely transmitted by radio frequency identification transceiver.



The Role of pH in chronic wounds



The Role of pH in chronic wounds

1. pH changes is related to wound healing process



Stage I
 5.7 ± 0.5



Stage II
 6.9 ± 1.0



Stage III
 7.6 ± 0.2

pH value by stanging

Tsukada et al. WOUNDS 1992

2. In Chronic wound an increase in pH is a sign of infection if compared with the normal surrounding skin

pH measurement

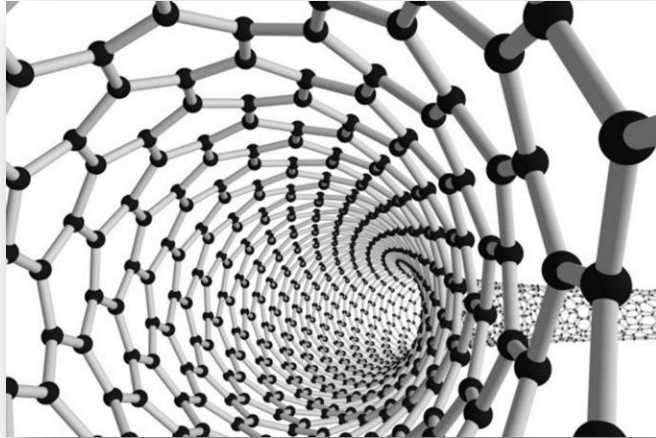
A flat glass electrode in contact with the skin and connected to a potentiometer

Disadvantage

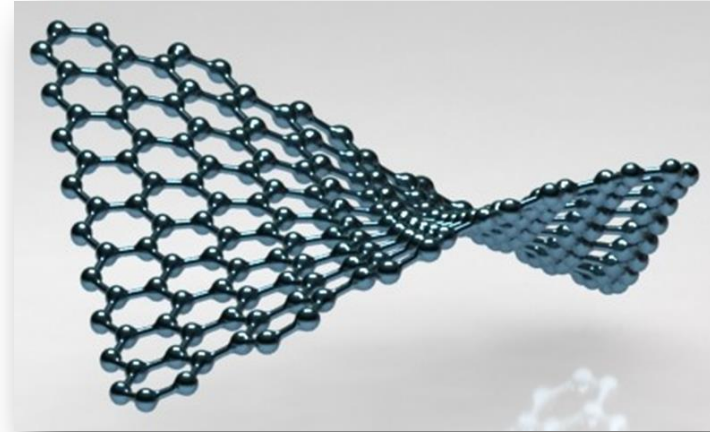
Impractical for implementation in wound dressing for continuous monitoring



Innovative Sensitive Materials



Carbon nanotube



Graphene



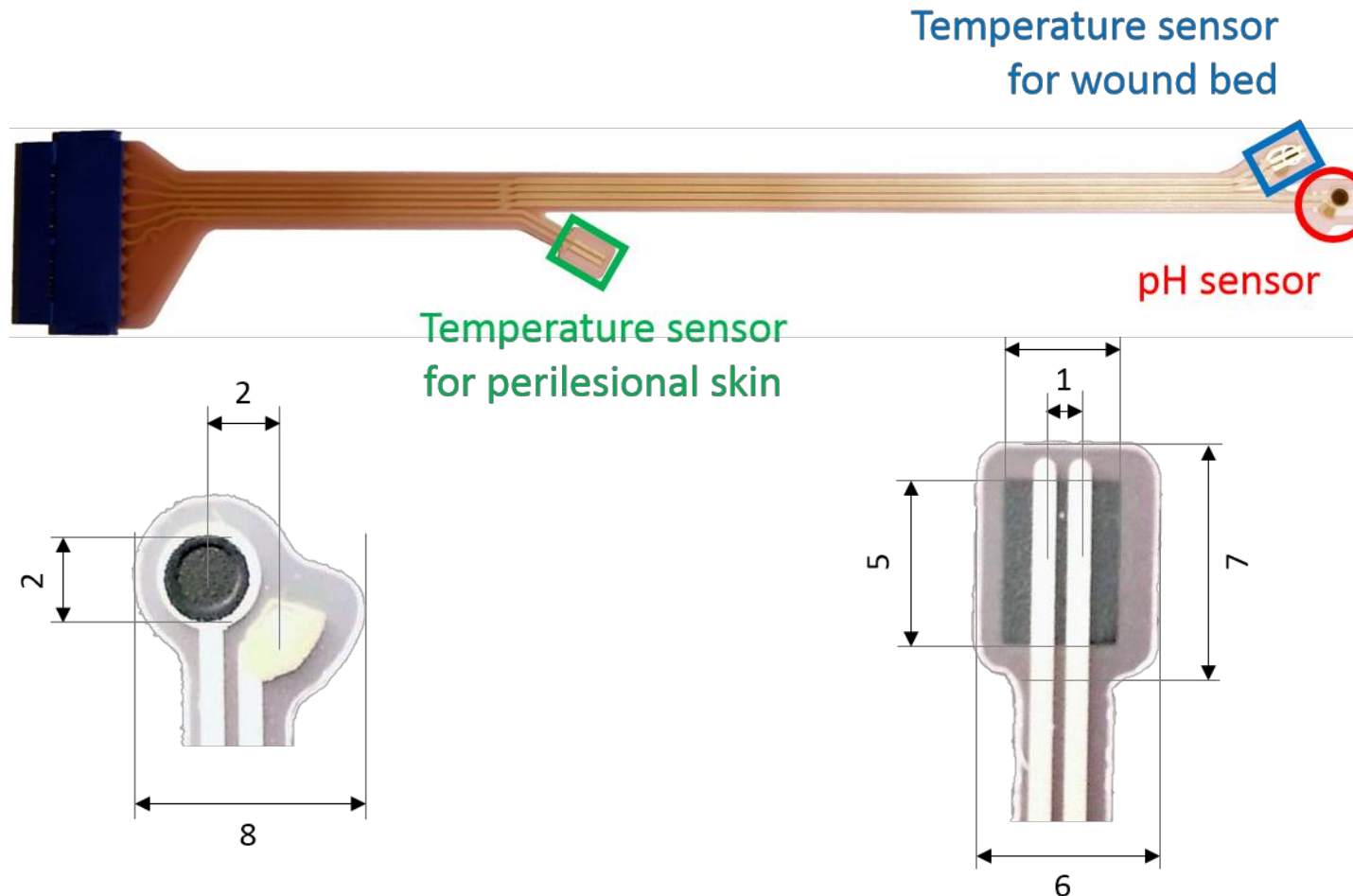
Photo: U. Montan
Andre Geim



Photo: U. Montan
Konstantin Novoselov

Sensors fabrication

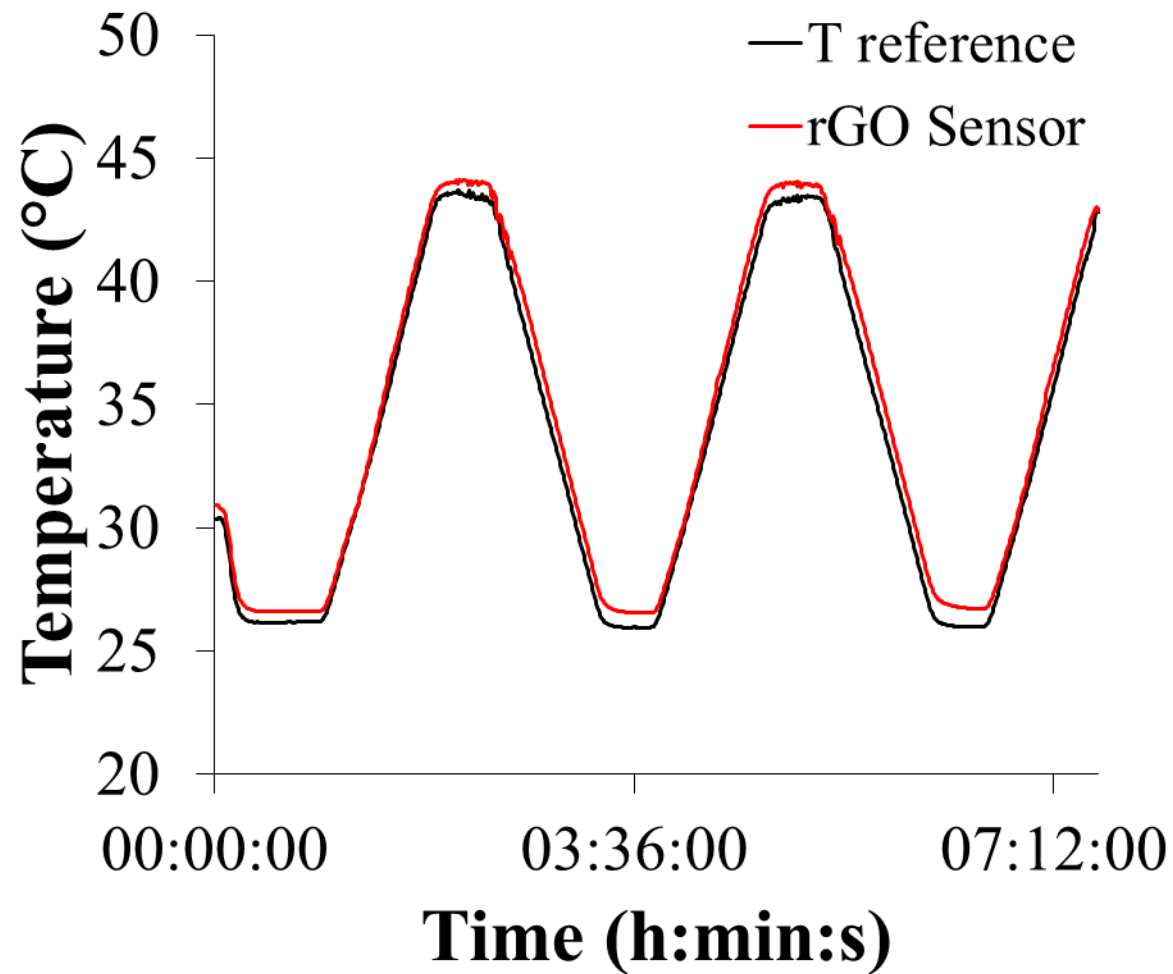
Graphene Oxide films were deposited by drop casting; for temperature sensing, GO films were reduced by a water solution of ascorbic acid (25 mg/L, 20 minutes at 80 °C)¹.



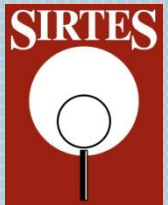
Reduced graphene oxide shows an almost linear dependence of electrical resistivity from temperature

Graphene oxide dispersions in water can be used to prepare pH sensitive films and disposable pH sensors

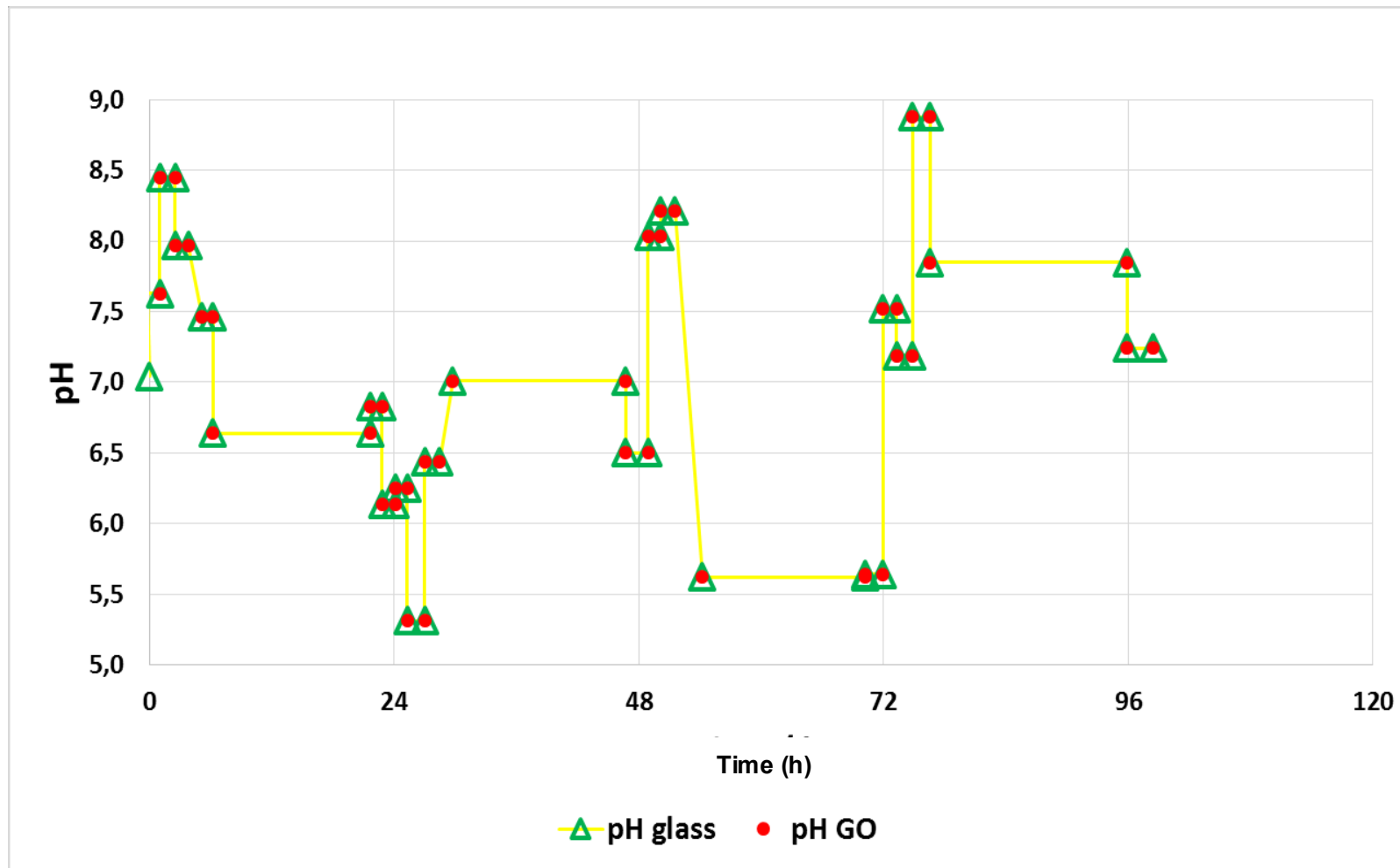
Test in human serum



Comparison between an rGO sensor and a thermistor in human serum at 30 and 45 °C

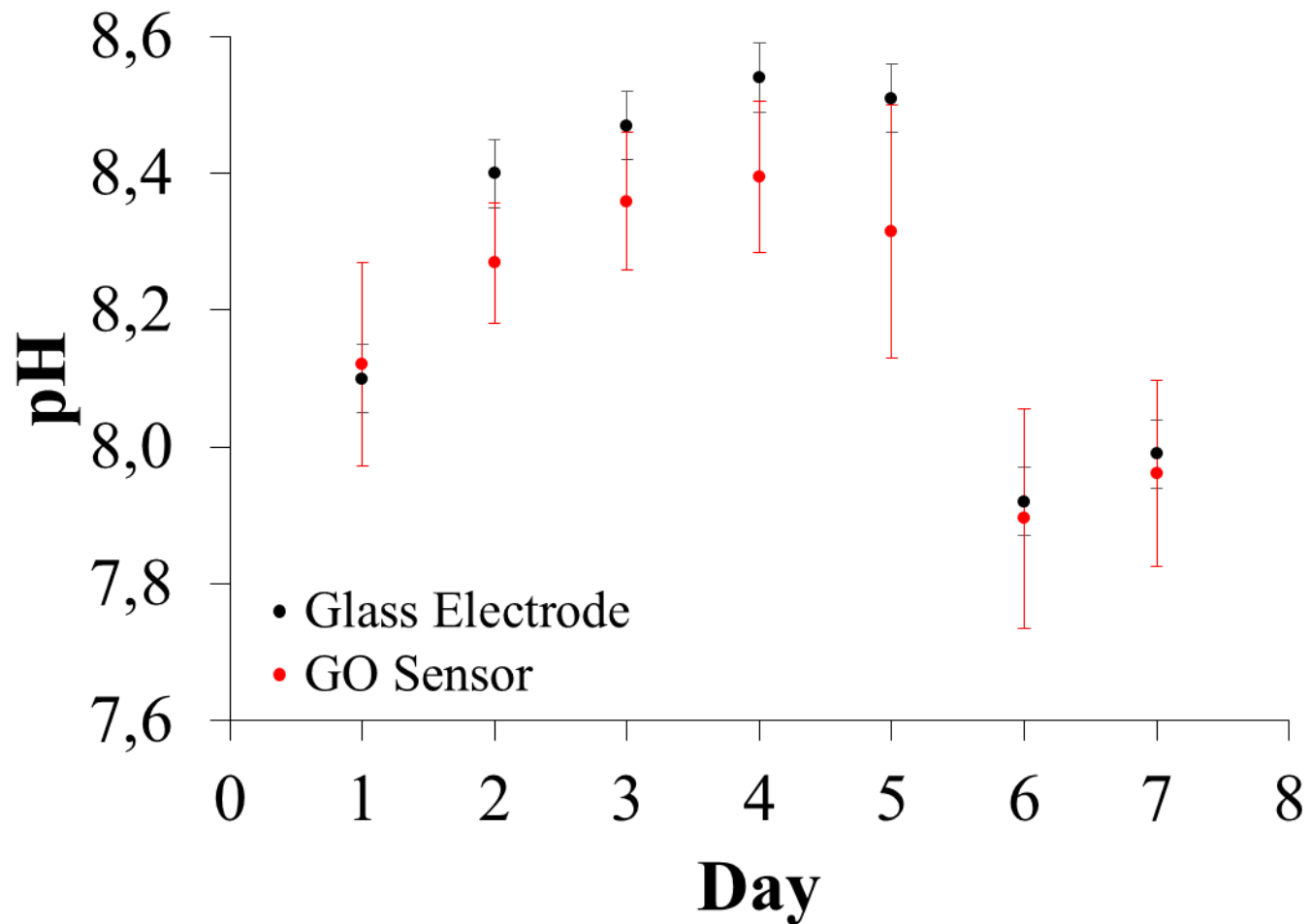


GO sensor vs glass electrode over time



Test in Hank's buffer salt solution

pH measurements in wound exudate

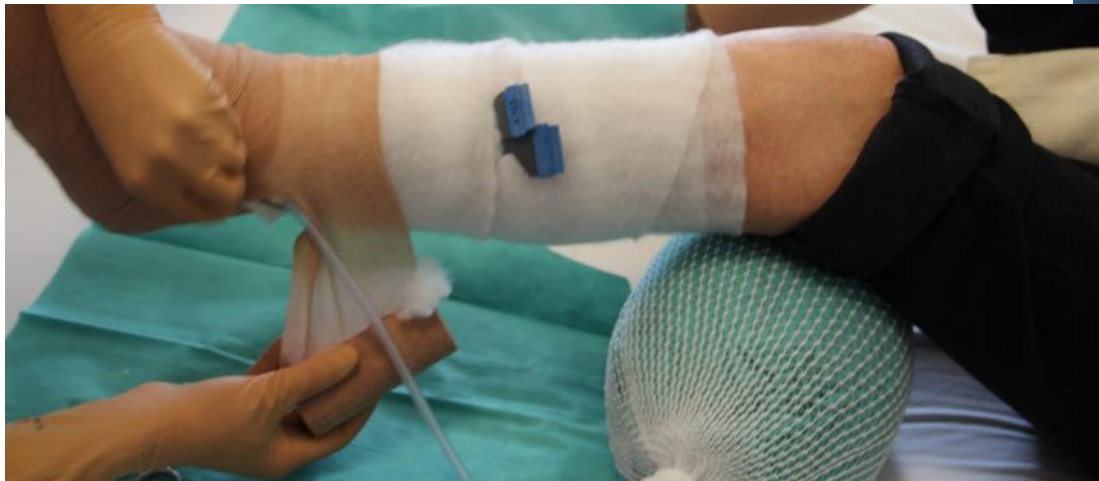


Minor deviations (0,2 pH units) from a reference glass electrode over one month

pH and Temperature evaluation



Application of the SWAN-iCare system



SWAN-ICARE SYSTEM

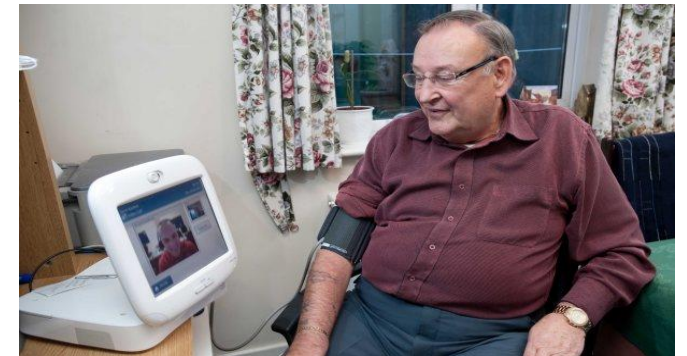
- Development of system focus:
 - 3 sensors
 - pH, temperature and MMP
 - 3 sub-systems :
 - In Wound Sensor Device (IWSD)
 - Smart Negative Pressure Device (SNPD)
 - Clinical back end



Expected Impact: the patient

Benefits for the patient

- Continuous home monitoring of a number of wound parameters
- Personalised therapy initiated by the physician remotely and adapted to the daily measurements
- Faster wound healing due to the early identification of potential problems
- Wound deterioration can be identified early and acted upon, therefore leading to reduced morbidity and amputation rates
- Reduced disturbance to patients life and possible need for hospitalisation
- Better quality of life with better mobility, more comfort ,less stress



Expected Impact: Society and Healthcare

Benefits for society and healthcare

- Reduced healthcare costs as a result of reduced need for hospitalization
- Reduced burden for the patients relatives due to faster wound healing and remote monitoring
- Reduced social costs and improved productivity as the patient returns to work earlier
- Increased access to best practice wound care for patients living in remote geographical locations
- Reduced daily nursing visits allows for more new patients' to be added to the case load

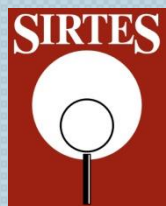


Conclusions

- Disposable temperature and pH sensors are required to monitor chronic wounds
- The SWAN-iCare project aims at coupling negative pressure therapy and monitoring of wound conditions
- Graphene oxide dispersions in water can be used to prepare pH sensitive films and disposable pH sensors
- Reduced graphene oxide shows an almost linear dependence of electrical resistivity from temperature
- **Sensor materials are biocompatible and can be used in contact with the wound bed**
- These sensors show a good stability over time

Acknowledgements

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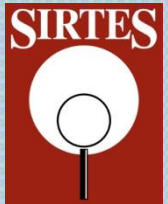


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If you want to run fast ,run alone

If you want to run far, run together....



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