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ARTICLE TYPE

# One-Touch-Activated Blood Multidiagnostic System using a Minimally Invasive Hollow Microneedle Integrated with a Paper-Based Sensor

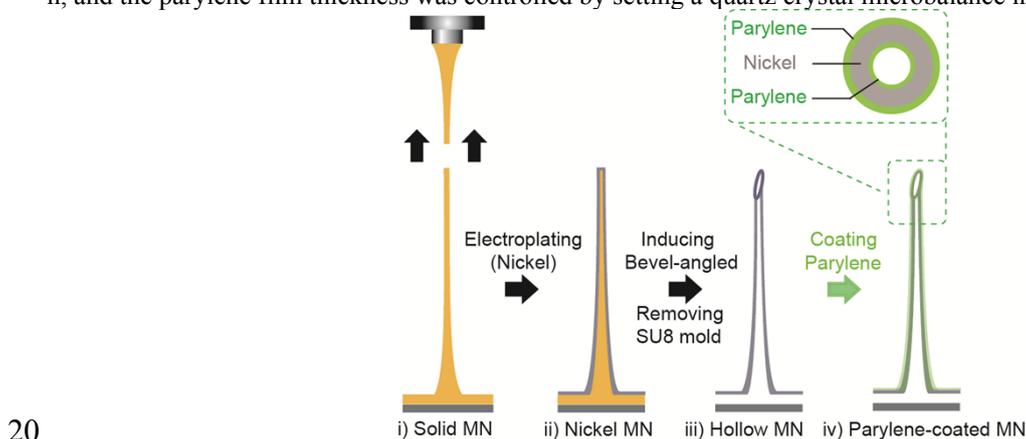
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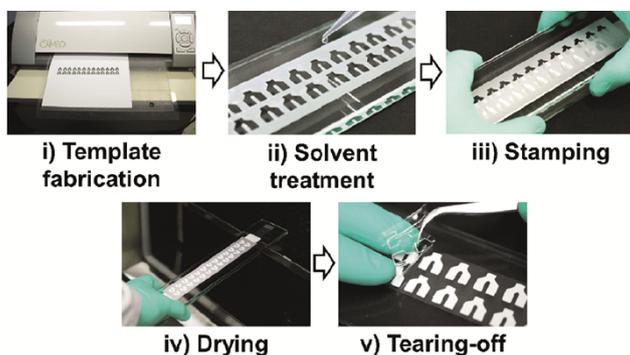
## Supplementary Information

*Fabrication of the biocompatible hollow microneedle:* Negative photoresist SU-8 2050 (MicroChem Corp., Westborough, MA., USA) was spin-coated onto a 120  $\mu\text{m}$ -thick flat glass panel. After placing on a 120  $^{\circ}\text{C}$  hot plate for 5 min and cooling to 60  $^{\circ}\text{C}$ , the pillar with a diameter of 300  $\mu\text{m}$  was used to draw the SU-8 at a rate of 10  $\mu\text{m s}^{-1}$ , producing a microstructure with a height of 3,600  $\mu\text{m}$ . This SU-8 mold was cured for 30 min at room temperature to solidify the polymeric bridge, and then separated from the frame at a drawing speed of 700  $\mu\text{m s}^{-1}$ . A SU-8 solid microneedle (MN) was fabricated with a height of 1,800  $\mu\text{m}$  and tip-diameter of 60  $\mu\text{m}$ . After a silver layer was deposited onto the surface of the solid microneedle by Tollen's test, nickel electroplating was performed to fabricate a nickel microneedle with a wall thickness of 30  $\mu\text{m}$ . The 15 $^{\circ}$  bevel angle was induced at the tip of the nickel microneedle by laser cutting (K2 Laser System, Inc, Gyeonggi-Do, Korea), and then the solid mold was removed by SU-8 Remover (MicroChem Corp.) to complete the hollow microneedle. Finally, a 1  $\mu\text{m}$ -thick parylene film was thermally deposited on the surface of the microneedle by a parylene coating system (Femto Science Inc., Gyeonggi-Do, Korea). All parylene coating processes were performed under vacuum conditions in 1.5 h, and the parylene film thickness was controlled by setting a quartz crystal microbalance in the deposition chamber.



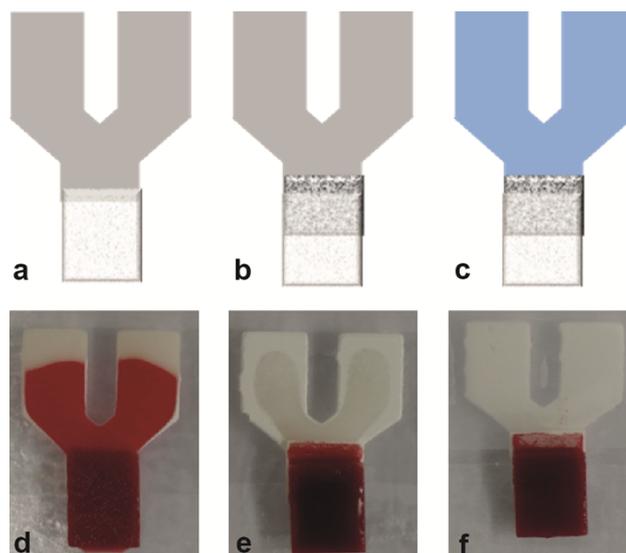
**Fig. S1** Schematic diagram of the fabrication process of biocompatible hollow microneedle (MN). i) a three-dimensional SU-8 solid mold was obtained by a drawing lithography method; ii) nickel was electroplated onto an SU-8 solid mold; iii) a bevel angle was introduced and the solid SU-8 mold was removed; iv) a biocompatible parylene film was coated onto the inner and outer surface of the hollow microneedle.

*Fabrication of the patterned nitrocellulose (NC) membrane:* The Y-shape NC membrane pattern was fabricated by “tear-off patterning”. Briefly, the Y-shape pattern was designed by Silhouette Studio software (version 3.3.451), and the paper template was fabricated using a craft cutter. After, treatment of the paper template with dimethyl sulfoxide (DMSO) was conducted by pipetting, and the NC membrane and the template were placed on a glass support facing each other. After 25 min, the template and membrane were incubated in a dry oven heated to 37 °C, for 15 min. Finally, the stamped area was removed with forceps.



**Fig. S2** Schematic of the fabrication process of Y-shaped nitrocellulose (NC) membrane by the tear-off patterning method. i) Paper template fabrication by craft cutter; ii) solvent treatment on template; iii) stamping the solvent treated template to the NC membrane; iv) drying, and v) tearing off the stamping region.

**Figure S3**



**Fig. S3** Schematic diagrams and photograph images for optimization of sensor structure. The paper-based sensor contained (a and d) only sample pad (fusion 5 8151-6621); (b, c, e, and f) sample pad and asymmetric polysulfone membrane; and (c and f) nitrocellulose membrane treated with 0.5% surfactant 10G.

**Supplementary Table S1. *In vitro* whole-blood sample analysis**

Rabbit #	Glucose concentration (mg dL <sup>-1</sup> )		Cholesterol concentration (mg dL <sup>-1</sup> )	
	Lab Analyzer	OBMS	Lab Analyzer	OBMS
# 1	138	145.5 ± 2.43	11	11.4 ± 2.07
# 2	153	153.9 ± 6.54	20	20.2 ± 4.93
# 3	128	137.8 ± 4.87	31	26.3 ± 5.66

10 Comparison of the one-touch-activated blood multidagnostic system (OBMS) and lab scale analyzer (Lab Analyzer; FUJI DRI-CHEM 4000 Chemistry Analyzer). Values are expressed as mean ± SD (n = 3).

### Supporting Videos:

15 Supplementary Video 1: *In vitro* one-touch-activated blood diagnosis (Video clip: 2.36 MB)

Supplementary Video 2: *In vivo* one-touch-activated blood diagnosis (Video clip: 6.95 MB)