

Skin application of tape and MN patches Twenty consecutive D-Squame tape strips (22 mm diameter) were collected from the lesional skin of the patients with AD. Hollow MNs, blank patches, and HA-MNs were also applied to the lesional skin of these patients for 10 min. Upon removal from the skin, HA-MNs were dissolved, and proteins in the skin metabolites were collected for experimental analysis (Figure 1B2,B3).

**2.4 Protein extraction** To extract proteins from the skin metabolites of the tape, hollow MNs, blank patches, and HA-MNs, each sample was scraped into 1% sodium dodecyl sulfate (SDS) buffer<sup>34</sup>, 35 (Thermo Fisher Scientific).

**2.5 Total protein quantification** To quantify the concentration of total proteins in the skin metabolite extracts, a bicinchoninic acid (BCA) protein assay kit (Thermo Fisher Scientific) was used according to the manufacturer's protocol.

**3 RESULTS**

**3.1 MN patch design** Biocompatible HA-MNs (each 160 array, 1 mm base width, 650  $\mu$ m height) were fabricated using the droplet extension method<sup>32</sup> by drying a pharmaceutical-grade HA solution on top of a hydrocolloid patch (Figure 1A). Statistical significance was set at  $p < 0.05$ .