

## **Tissue digestion and cell fraction separation**

Samples were obtained after approval by the Institutional Review Board of the Hospital Clinic and Clinica Planas, Barcelona. Between 40 and 80 g of fresh mammary tissue from five women aged 20-45 years was used as starting material (**Supplementary Table 1**). Briefly, the material was minced into small pieces and digested at 37°C overnight (ON) in a shaker in DMEMF12 medium with 200 U/mL of collagenase type IV (Sigma, MO, USA) and 100 U/mL of hyaluronidase (Sigma, MO, USA), as we previously described (1). Subsequently, organoids, single epithelial cells and stromal cells were separated by differential centrifugation at 40 g for 1 min, 100 g for 2 min and 200 g for 4 min, respectively.

## **Bibliography**

1. Zubeldia-Plazaola A, Ametller E, Mancino M, *et al.* Comparison of methods for the isolation of human breast epithelial and myoepithelial cells. *Front Cell Dev Biol* 2015;3: 32.