

scar contracture are softer and that skin hyperpigmentation disappears after fat grafting.<sup>3</sup> Such regenerative/revitalizing effects of fat grafting are considered to be derived from ASCs in the tissue. Therefore, condensation of ASCs in the graft may be also crucial for such regenerating/revitalizing applications.

#### BASIC SCIENCE

##### *Difference Between Aspirated Fat Tissue and Intact Adipose Tissue*

Adipose tissue contains various types of cells including adipocytes and ASCs, as well as connective tissue (see the article by Mashiko and Yoshimura elsewhere in this issue for details). When surgeons aspirate fat, only fragile parts of adipose tissue are harvested through a suction cannula, whereas the honeycomb-like fibrous structures remain intact in the donor site.<sup>4</sup> The fibrous structure is predominantly composed of connective tissues and large vasculatures, which are considered to contain many ASCs. We have found that aspirated fat tissue contains only one-half the number of ASCs compared with intact fat tissue.<sup>5</sup> Stage-specific embryonic antigen-3-positive cells, which may be highly multipotent stem cells (muse cells),<sup>7</sup> locate around large vasculatures. These cells are also deficient in aspirated adipose tissue (unpublished data, Doi K et al, 2012). The relative deficiency of ASCs in aspirated fat tissue may be owing to (1) a substantial portion of ASCs being left in the donor tissue and (2) some ASCs being released into the fluid portion of liposuction aspirates, possibly owing to the act of an endogenous enzyme.<sup>1,8</sup> Thus, aspirated fat tissue is regarded as relatively ASC poor compared with intact fat tissue. This low ASC/adipocyte ratio may be a reason for long-term atrophy after fat grafting.<sup>2</sup>

##### *Importance of Adipose-Derived Stem/Stromal Cells in the Grafted Tissue for Adipose Regeneration after Fat Grafting*

ASCs have the potential to modulate or suppress immunoreaction,<sup>9</sup> differentiate into adipocytes,<sup>9,10</sup> vascular endothelial cells, or others and release angiogenic growth factors, such as hepatocyte growth factor and vascular endothelial growth factor, especially under hypoxic conditions.<sup>11</sup> ASCs were reported to contribute to angiogenesis during the adipose remodeling process after ischemia or fat grafting.<sup>9-11</sup> Our recent study using green fluorescent protein mice revealed that regenerated adipocytes after fat grafting are mostly originated from ASCs in the graft tissue, but not from other host-derived stem/progenitor cells, although new ASCs can be provided partly by bone marrow or

other tissues.<sup>12</sup> It was suggested that only ASCs originally located adjacent to dying adipocytes can become adipocytes, although other ASCs can contribute in other ways, such as angiogenesis or release of growth factors.

#### RELEVANCE TO CLINICIANS

##### *Graft Tissue Condensation*

Liposuction aspirates contain some components unnecessary for adipose tissue engraftment/regeneration; water, oil (broken adipocytes), and blood cells (red blood cells and white blood cells). It is recommended to remove such components and reduce the graft volume without reducing the number of viable adipocytes and ASCs; this is called condensation of graft tissue. Tissue condensation is important, especially when there is a maximum limit in graft volume, such as with breast augmentation. There are 3 major methods for graft tissue condensation: decantation (gravity sedimentation), filtration with or without a vacuum, and centrifugation. Among the 3, centrifugation is most effective to remove the water content without losing ASCs, although some adipocytes can be broken by the mechanical force and the resulting condensed fat may become more viscous and need higher pressure to inject through a small cannula (Fig. 1).<sup>13</sup> Oil released from damaged adipocytes causes inflammation-like foreign materials, suggesting that removal of oil should be important for better healing after fat grafting.

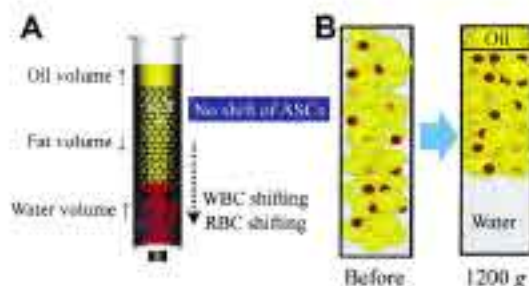


Fig. 1. Tissue and adipose-derived stem/stromal cell (ASC) condensation by centrifugation. (A) By centrifugation, fat volume becomes compact, water volume increases and oil will be clearly separated as a top layer. Many of red blood cells (RBCs) and white blood cells (WBCs) in the aspirated fat shift into the water portion after centrifugation, but most of ASCs remain in the fat portion. (B) By centrifugation at 1200×g for 3 minutes, fat volume decreases by 30%. Some adipocytes are broken and become oil as a top layer, but all ASCs remain intact and are concentrated in the condensed fat tissue.