

## REVIEW ARTICLE

### Stem Cells in Dentistry

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#### ABSTRACT:

The human body is made up of three basic categories of cells: germ cells, somatic cells and stem cells. The materials required for tissue engineering include stem cells, morphogens (or growth factors) and a scaffold to guide cell growth. The present review article highlighted the role of stem cells in dentistry.

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#### INTRODUCTION

The human body is made up of three basic categories of cells: germ cells, somatic cells and stem cells. The materials required for tissue engineering include stem cells, morphogens (or growth factors) and a scaffold to guide cell growth. Scientific study into cell-based therapies has identified tremendous potential for the use of these stem cells to treat a number of diseases and disorders.<sup>1</sup>

It is now accepted that progenitor/stem cells reside within orofacial region. Stem cells residing in the orofacial region have been classed as the Mesenchymal stem cells (MSCs)/Adult stem cells (ASCs)/ Tissue stem cells (TSCs). Studies have identified several niches of multipotent mesenchymal progenitor cells, known as dental pulp stem cells, which have a high proliferative potential for self-renewal. These progenitor stem cells are now recognized as being vital to the dentine regeneration process following injury. More recently, researchers have discovered that stem cells harvested from deciduous teeth may be a source

of tissue regeneration and repair. Five different types of dental stem cells isolated from dental soft tissues are dental pulp, apical papilla, dental follicle and periodontal ligament.<sup>2</sup>

In general, tissue engineering strategies in the field of dentistry using dental stem cells are mainly applied for whole tooth regeneration or development of wide variety of orodontal structures including resorbed root, cervical, or apical dentin, periodontal structures and gingival, which can facilitate and restore the physiological structural integrity. In this regard, efforts to formulate bioengineered teeth were made by seeding dental epithelium and mesenchymal tissues or dissociating embryonic tooth germs to prefabricated tooth shaped scaffolds. Eventually, it led to the formation of basic tooth-like structures with the development of enamel, dentin, and pulp.<sup>2</sup>

#### STEM CELL TYPES

Stem cells can be broadly divided into

1. Embryonic stem cell
2. Adult stem cell
  - Hematopoietic stem cell
  - Mesenchymal stem cell
3. Induced pluripotent stem cell

### **SOURCES OF STEM CELLS<sup>3</sup>**

The oral and maxillofacial region can be treated with stem cells from the following sources

1. Bone marrow
2. Adipose tissue
3. Stem cells from oral and maxillofacial region

#### **Bone marrow**

Bone marrow stem cells (BMSCs) can be harvested from sternum or iliac crest. It is composed of both hematopoietic stem cells and mesenchymal stem cells (MSCs). The majority of oro-maxillofacial oral structures are formed from mesenchymal cells. The advantage of bone marrow is that it has a larger volume of stem cells and can be differentiated in to wide variety of cells. Isolation of BMSCs can be carried out only under general anesthesia with possible post operative pain.

#### **Adipose tissue**

They can be harvested from the lipectomy or liposuction aspirate. Adipose derived stem cells (ADSCs) contain a group of pluripotent mesenchymal stem cells that exhibit multilineage differentiation. Advantage of adipose tissue is that it is easily accessible and abundant in many individuals.<sup>4</sup>

#### **Dental Pulp Stem Cells (DPSC)**

Their source is dental pulp mesenchyme (neural crest mesenchyme). They are slow cycling cells having restricted potential and represent mature adult pulp stem cells. They have better immunologic/host acceptance. In vitro they formed odontoblasts, osteoblasts, endothelial cells, adipocytes, chondrocytes, neurons and smooth muscle cells while in vivo, various directions like odontogenic, myogenic, adipogenic, angiogenic and osteogenic are found and were able to form complete dentin pulp complex. Its in vitro developmental capability and in vivo therapeutic targeting is yet to be explored.

#### **Stem Cells From Human Exfoliated Deciduous Teeth (SHED):**

Their source is human exfoliated deciduous teeth (coronal pulp). They are multipotent cells with very high proliferative potential and higher cell doublings. In vitro they can differentiate odontogenically, osteogenically, adipogenically, chondrogenically, or neurally. In vivo they can form neurons, adipocytes, odontoblasts, and osteoinductive and endothelioid cells. But they failed to form complete dentin pulp complex in vivo.<sup>5</sup>

#### **Oral Epithelial Stem Cells (OESCs)**

They are derived from oral epithelial progenitor cells from basal layer of oral mucosa. They are unipotent stem cells and possess clonogenicity. They can form highly stratified and well organized graft. But they cannot differentiate into mesenchymal cell lineage.

#### **Gingiva derived MSCs (GMSCs)**

These cells are derived from lamina propria of gingiva. They possess clonogenicity, self-renewal and multipotent differentiation capacity similar to BMSCs and proliferate faster than BMSCs, display stable morphology after extended passages. These cells exhibit adipogenic, osteogenic and chondrogenic potential along with immunomodulatory effect on lymphocytes. Their applications further need to be explored.<sup>6</sup>

#### **Salivary gland stem cells (SGSCs)**

They are derived from the stromal tissue of salivary glands. They are useful for regeneration of salivary gland damaged from irradiation and can be guided to osteogenic, chondrogenic and adipogenic differentiation. It is difficult to isolate salivary gland stem cells from the collection of stromal cells.<sup>7</sup>

#### **Tooth Germ Progenitor Cells (TGPCs)**

They are the stem cells in the mesenchyme of the third molar tooth germ and possess very high proliferative activity. They can differentiate into lineages of three germ layers including osteoblasts, neural cells and hepatocytes. Their applications further need to be explored.<sup>8</sup>

#### **Stem cells from the oral and maxillofacial region**

Stem cells from oral and maxillofacial region predominantly contain mesenchymal stem cells. In oral and maxillofacial area different types of dental stem cells were isolated and characterized. They include

- Dental pulp stem cells (DPSCs).
- Stem cells from exfoliated deciduous teeth (SHED).
- Periodontal ligament stem cells (PDLSCs).
- Stem cells from apical papilla (SCAP).
- Dental follicle progenitor cells (DFPCs).<sup>9</sup>

#### **Dental stem cell advantages**

The advantages of stem cells from oral and maxillofacial region is that

1. Have high plasticity.
2. It can be cryopreserved for longer period (Ideal for stem cell banking).
3. It showed good interaction with scaffold and growth factors.
4. Stem cells transplantations can cause pathogen transmission and also need immunosuppression, so autologous stem cell source is the best option. Dental pulp stem cells will be better fitting tool due to easy surgical

access, the very low morbidity of the anatomical site after the collection of the pulp.<sup>10</sup>

### **Stem cells in periodontal regeneration**

Stem cells will be a promising tool for regenerating the periodontal structures such as periodontal ligament and other supporting elements. BMSCs have the ability to produce alveolar bone, periodontal ligament, and in vivo cementum after implantation into the periodontal defects.<sup>11</sup> Thus, it was proved BMSCs provides an alternative source for the treatment of periodontal diseases. Autologous mesenchymal stem cells from iliac crest in combination with platelet rich plasma from peripheral blood was used for periodontal regeneration. Significant closure of bone defect and improvement of attachment level was observed after one year follow up. It also showed good healing and regeneration of interdental papilla.<sup>12</sup>

### **Regeneration of craniofacial defects**

Stem cells can be useful in the regeneration of bone and to correct large craniofacial defects due to cyst enucleation, tumor resection, and trauma. The closure of a bone defect is commonly carried out with the transfer of tissue, which have disadvantages like, not able to restore the unique function of the lost part, donor site morbidity, accompanied by scarring, infection and loss of function.<sup>13</sup>

### **Tooth regeneration**

The regeneration of adult teeth will be possible in future with the newer advancement in stem cell therapy and tissue engineering. Regenerative procedures would be better fitting and alternative tool in place of dental implants. Experimental studies with animal models have shown that the tooth crown structure can be regenerated using tissue engineering techniques that combine stem cells and biodegradable scaffolds.<sup>14</sup>

### **Conclusion**

Future tissues like tissue engineered bone grafts, engineered joints and cranial sutures can be developed with stem cell therapy. A team of professionals including stem cell biologists, molecular biologists, geneticists, polymer and materials scientists, mechanical engineers and clinicians with knowledge of oral and maxillofacial disorders is needed to develop the field of craniofacial tissue engineering.

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