

## Efficacy of Adipose-Derived and Bone Marrow-Derived Stem Cells in Spinal Fusion: A Comparative Study in a Rat Model

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### Introduction

Although bone marrow-derived stem cells (BMSCs) have been widely used in spinal fusion research, adiposederived stem cells (ADSCs) offer a number of advantages as a clinical cellsource, including: increased tissue volume availability, higher stem cell concentrations, and reduced donor site morbidity. In this study, the efficacy of ADSCs vs. BMSCs in achieving successful spinal fusion, when combined with a clinical-grade bone graft substitute, is compared in a rat model.

### Methods

ADSCs were isolated from the inguinal fat pads, while BMSCs were isolated from the long bones of syngeneic female 6-8 week old Lewis rats and cultured in vitro until passage 2 for subsequent transplantation. Posterolateral spinal fusion surgery at L4-5 was performed on 36 female Lewis rats (6-10 wk old) divided into 3 experimental groups: [1] Vitoss (Stryker) bone graft substitute only (VO group, n=12); [2] Vitoss + 2.5 x 10<sup>6</sup> ADSCs/side (n=12); and [3] Vitoss + 2.5 x 10<sup>6</sup> BMSCs/side (n=12). Fusion was assessed eight weeks post-surgery via micro-computed tomography (MicroCT) analysis, manual palpation, and histology. Manual palpation scoring was conducted by blinded researchers as follows: 0=non-fused; 1=partial fusion, some motion across operative joint;

## MicroCT imaging analyses showed that the average fusion volume in the ADSC group was significantly higher than in the BMSC and VO groups (44.3 mm<sup>3</sup> vs. 27.6 mm<sup>3</sup> and 30.0 mm<sup>3</sup>, respectively, p < 0.01). Similarly, average manual palpation score was the highest in the ADSC group compared with the BMSC and VO groups (1.5 versus 0.7 versus 0.8, p =0.03).

**Fusion Assessment** 

Palpation Score:

1 = Partial Fusior

CT Fusion Score:

Each Side:

0 = Nonfused 1 = Partial Fusion

2 = Solid Fusion

Score = Avg. Sum of Both Sides

Data presented as mean  $\pm$  STD, (n= 12), \* = (p < 0.05)

250

aMSL

2 = Solid Fusion

0 = Nonfused

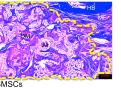
Score = Ava

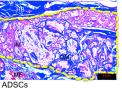
**Results** 

Histologically, ADSC and BMSC groups showed a higher concentration of osteogenic matrix and osteoblasts in the fusion mass compared to VO.

# Assessment of Fusion Mass Bone Quality







Masson's Trichrome Stain BM = Bone marrow HB = Host bone OM = Osteogenic matrix IM = Implant MF = Muscle fibers

### Conclusions

When combined with a clinical grade bone graft substitute in a rat model, ADSCs yielded increased fusion mass volume and rates of fusion than bone marrow-derived stem cells. Ongoing studies will explore whether freshly isolated ADSCs will yield similar results.

### Learning Objectives

By the conclusion of this session, participants should be able to:

1) Describe the potential role of BMSC and ADSC in spinal fusion.

 Discuss the translational importance of these treatments.

3) Identify advantages offered by these alternatives therapies.

#### References

[1] Kitchel et al. (2006) Spine J. 6(4) [2] Carter et al. (2009) Spine J. 9(6) [3] Yamada et al. (2012) Spine 37 [4] Muschler et al, 2004, Clin. Orthop.Relat. Res. 432:242 [5] Gan et al, 2008, Biomaterials. 29(29),3973 [6] Odri e al, 2012, Eur Spine J. 21(12):266 [7] Stolzing et al. (2008) Mech. Ageing Dev. 129(3) [8] Benisch et al. (2012) PLoS One 7(9) [9] Stenderup et al. (2003) Bone 33(6) [10] Salamanna et al. (2017) Stem Cells Int. 3537094 [11] Thesleff et al. (2011) Neurosurgery. 68(6) [12] Guven et al. (2011) Biomaterials.32(25) [13] Nunes et al. (2013) Sci Rep. 3:2141 [14] Shi et al. (2005) Plast Reconstr Surg. 116(6) [15] Chen et al. (2012) J Cell Mol Med.16(3)