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임상

P-90

Canine Adipose Tissue-derived Mesenchymal Stem Cells Alleviate Severe Acute Pancreatitis by Regulating T cells in Rats

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Introduction: Severe acute pancreatitis (SAP) is associated with systemic complications and high mortality rate in dogs. Recently, mesenchymal stem cells (MSCs) have been investigated for their therapeutic potential in several inflammation models. The present study investigated the effects of canine adipose tissue-derived (cAT)MSCs in a rat model of SAP.

Materials and Methods: A rat model of SAP was induced by retrograde injection of 3% sodium taurocholate solution into the pancreatic duct. cATMSCs labeled with diiodoacetyl-3,3',3'-tetramethylindocarbocyanine perchlorate (1×10^7 cells/kg) were systemically administered and pancreatic tissue was collected 3 days later. Histopathological, quantitative real-time PCR, and immunocytochemical analyses were performed.

Results: Greater numbers of infused cATMSCs were detected in the pancreas of SAP as compared to sham-operated rats. cATMSC infusion reduced pancreatic edema, inflammatory cell infiltration, and acinar cell necrosis, and decreased pancreatic expression of the pro-inflammatory cytokines tumor necrosis factor- α , interleukin (IL)-1 β , -6, -12, -17, and -23 and interferon- γ , while stimulating expression of the anti-inflammatory cytokines IL-4 and IL-10 in SAP rats. Moreover, cATMSCs decreased the number of cluster of differentiation 3-positive T cells and increased that of Forkhead box P3-positive T cells in the injured pancreas

Conclusions: These results indicate that transplantation of cATMSCs might be developed as a potential therapy strategy of SAP in dogs.

P-91

Anti-inflammatory effects of Oct4/Sox2 overexpressing adiposetissue-derived mesenchymal stem cells

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Introduction: Recent works have been shown that *Oct4* and *Sox2* as main self-renewal factor enhance proliferation and pluripotency of adipose tissue-derived mesenchymal stem cells (ATMSCs). However, anti-inflammatory effects of ATMSCs overexpressing *Oct4* and *Sox2* (*Oct4/Sox2*-ATMSCs) have not been determined. The aim of the present study was to evaluate the anti-inflammatory effects of *Oct4/Sox2*-ATMSCs.

Materials and Methods: In vitro, green-fluorescent protein (GFP) (control) and *Oct4/Sox2*-ATMSCs were cultured for 48 h and the supernatant (conditioned media) was collected to treat lipopolysaccharide(LPS)-induced Raw 264.7 cells. Subsequently, the levels of inflammatory cytokines expression were determined using real-time PCR analysis. In LPS induced systemic inflammatory mice models, GFP- and *Oct4/Sox2*-ATMSCs (1×10^7 cells/kg) were injected intraperitoneally and monitored by survival rate and sick score (diarrhea, eye condition, activity and condition of their fur).

Results: *Oct4/Sox2*-ATMSCs group further decreased expression of pro-inflammatory cytokines (IL-1 β , IL-6, and TNF- α) from Raw 264.7 cells than GFP-ATMSCs group. Comparison to GFP- with *Oct4/Sox2*-ATMSCs injected mice, the total sick score was reduced to 1.53 fold, while the survival rate was increased by 11.1%.

Conclusions: Although further studies of mechanisms are needed, these results suggest *Oct4/Sox2*-ATMSCs may be developed as a novel therapy strategy of inflammatory diseases.

P-92

Temozolomide reduced the side population in human malignant mesothelioma cell lines

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Introduction: Malignant mesothelioma (MM) is a nearly incurable tumor with a median survival time of 6 to 9 months. To develop a measure to overcome therapeutic resistance of MM, the effects of temozolomide were investigated. The therapeutic