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Effectiveness of autologous bone marrow-derived mononuclear cells delivery in wide spectrum of pathologies

The promising field of regenerative medicine is working to restore structure and function of damaged tissues and organs. The adult heart represents an attractive candidate for cell-based technologies. While there is a wealth of preclinical and clinical data showing the safety, feasibility and efficacy of stem cells in adults with acute myocardial infarction and heart failure, less is known about possible implementation of stem cell therapy in infants and children with heart failure due to dilated cardiomyopathy and pulmonary arterial hypertension. The challenges facing cardiac stem cell therapy are multiple. There are uncertainties around the destiny of stem cells after their injection into the blood stream. In particular, it regards migration and homing of implanted cells in the target tissues. As yet unclear is the possible role of sympathetic nervous system in the context of osteoreflexotherapy. There is still no definitive answer to the question on which is the preferred type of stem cells to be used for transplantation in different settings. Since 2008, when we first used autologous Bone Marrow-Derived Mononuclear Cells (BM-MNCs) in patient with acute myocardial infarction, we have investigated the use of stem cells not only for myocardial regeneration in adults and pediatric patients, but also in adult patients with diabetes mellitus and osteoarthritis. The objective is to determine the role of BM-MNCs in management of wide spectrum of pathologies, including critically ill pediatric patients, adult patients with acute myocardial infarction and heart failure and adult patients with osteoarthritis. Two patients (9 and 15 years old) with trisomy 21 and severe pulmonary arterial hypertension due to uncorrected large ventricular septal defects received imtra-pulmonary BM-MNCs implantation. Radionuclide scintigraphy showed improvement of lungs vascularization during 36 months follow-up. Seven patients (4 months-17 years) with dilated idiopathic cardiomyopathy received intramyocardial BM-MNCs injections. During follow-up (up to 7 years), we observed improvement of left ventricular ejection fraction (LVEF), decrease of left ventricular end diastolic dimension by echocardiography and cardio-thoracic index at chest X-ray exams, reduction of serum brain-natriuretic peptide serum levels and decrease of the stage of heart failure from stage IV to stage I, by NYHA classification. No peri-procedural harmful side effects were observed. We performed BM-MNCs intracoronary infusion in 101 adult patients with acute myocardial infarction with reduced LVEF and in 14 patients with chronic heart failure. Our results showed statistically significant improvement in LVEF at 12 months. We also infused BM-MNCs to the pancreas directly via branches of splenic artery or superior pancreaticoduodenal artery we have performed single intra-articular BM-MNCs injections in 70 patients with knee or hip joint osteoarthritis (stage II-III). No adverse effects after the BM-MNC injection were observed. Preliminary analysis showed decrease in pain and other symptoms and statistically significant improvement by clinical scoring system using different questionnaires. The results are promising and we suggest that BM-MNCs might be used for the stabilization of the adult and pediatric patients to improve symptoms and outcomes or serve as a bridge for heart or lung transplantation or delay joint replacement surgery. It also could be recommended in cases if other more traditional treatment options fail or are contraindicated.

Biography

Aris Lacis has worked as a General and Thoracic Surgeon in P. Stradina University Hospital in Riga (1964-1969), Thoracic and Cardiac Surgeon in the Latvian Centre for Cardiovascular Surgery (1969-1994) and the Head of Pediatric Cardiology and Cardiac Surgery Clinic in University Children's Hospital, Riga from 1994 to 2012. He is the Vice President of Latvian Society for Cardiovascular Surgery and the President of Latvian Association for Pediatric Cardiologists. He is the author of 395 scientific publications, 3 monographs and 13 patents. He is the Investigator in more than 10 clinical trials including cardiosurgical procedures performed under deep hypothermia, hybrid procedures, etc.

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