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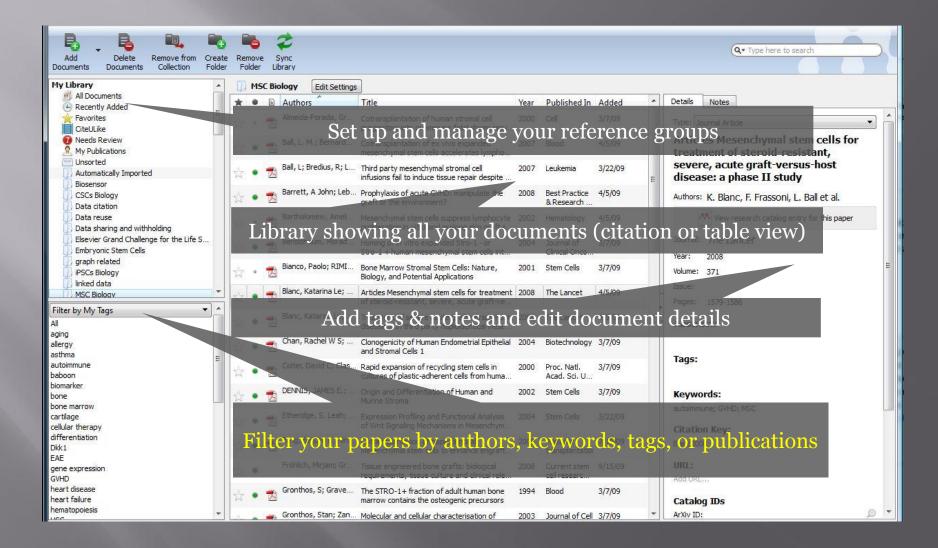
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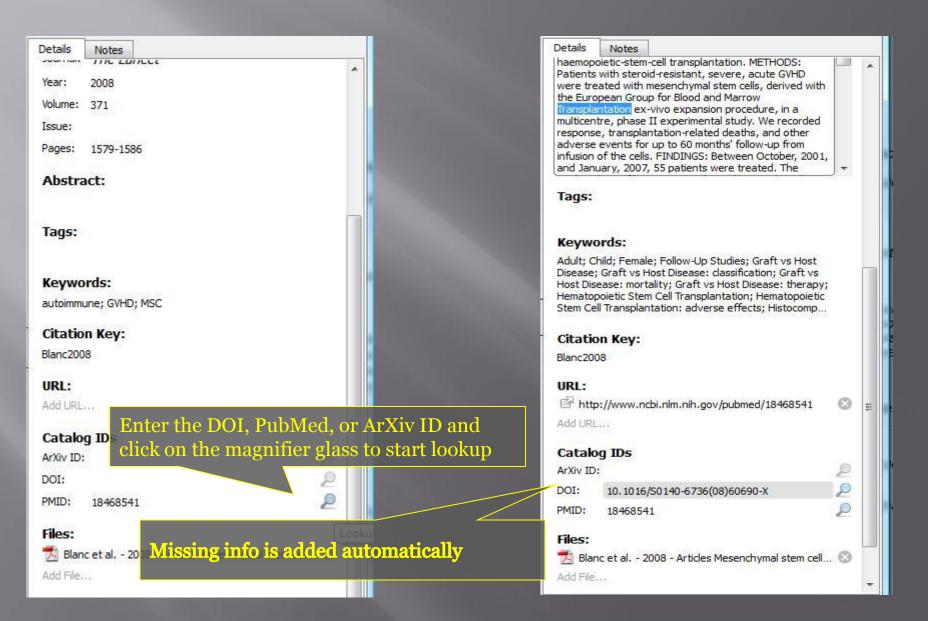
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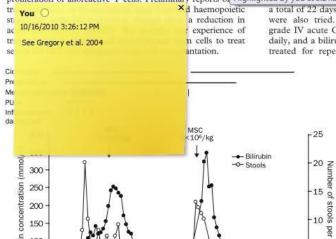
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Adult bone-marrow-derived mesenchymal stem cells are immunosuppressive and prolong the rejection of mismatched skin grafts in animals. We transplanted haploidentical mesenchymal stem cells in a patient with severe treatmentresistant grade IV acute graft-versus-host disease of the gut and liver. Clinical response was striking. The patient is now well after 1 year. We postulate that mesenchymal stem cells have a potent immunosuppressive effect in vivo.

Lancet 2004; 363: 1439-41 See Commentary page 1411

Severe acute graft-versus-host disease (GVHD) after allogeneic stem-cell transplantation is associated with high mortality. Bone marrow contains pluripotent mesenchymal stem cells that form bone, cartilage, adipose tissue, and muscle. These stem cells are not immunogenic and escape recognition by alloreactive T cells and natural killer cells. Mesenchymal stem, cells given intravenously have been well tolerated." I hermore, they are immunosuppressive and inhibit the



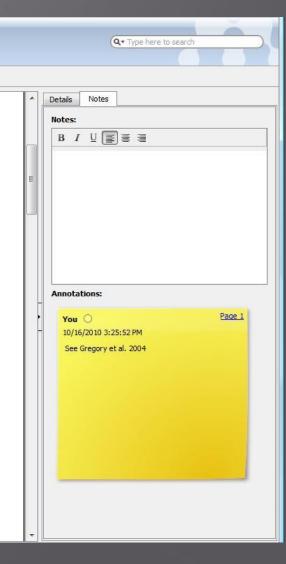
A 9-year-old boy with acute lymphoblastic leukaemia in third remission received a transplant of blood stem cells from an HLA-A, HLA-B, HLA-DRB1 identical, unrelated, female donor after conditioning with cyclophosphamide (120 mg/kg) and fractionated total body irradiation (3 Gy for 4 days). Immunosuppression included thymoglobulin (6 mg/kg) during the conditioning, followed by ciclosporin combined with four doses of methotrexate. On day 11 after allogeneic stem-cell transplantation, the patient developed a maculopapular rash of the thorax and back that progressed despite treatment with prednisolone (2 mg/kg daily). By day 22, the patient developed diarrhoea (>1000 mL per day) and abdominal pain requiring morphine. He stopped eating on day 24. Bilirubin and alanine aminotransferase concentrations rose (figure 1). Psoralen and ultraviolet-A light (PUVA) treatment (two to three times per week) for 3 weeks was followed by extracorporeal PUVA (one to four times per week) for 6 weeks. Infliximab (10 mg/kg) and daclizumab (1 mg/kg) for 4 weeks were ineffective proliferation of alloreactive T cells, Preliminary reports of Highlighted by you at 3:24:55 PM on Saturday, October 16, 2010 (day) for

a total of 22 days, mycophenolate mofetil, and methotrexate were also tried. By day 70, the patient had developed grade IV acute GVHD, including diarrhoea up to 20 times daily, and a bilirubin concentration of 250 mmol/L. He was treated for repeated bacterial, viral, and invasive fungal infections.

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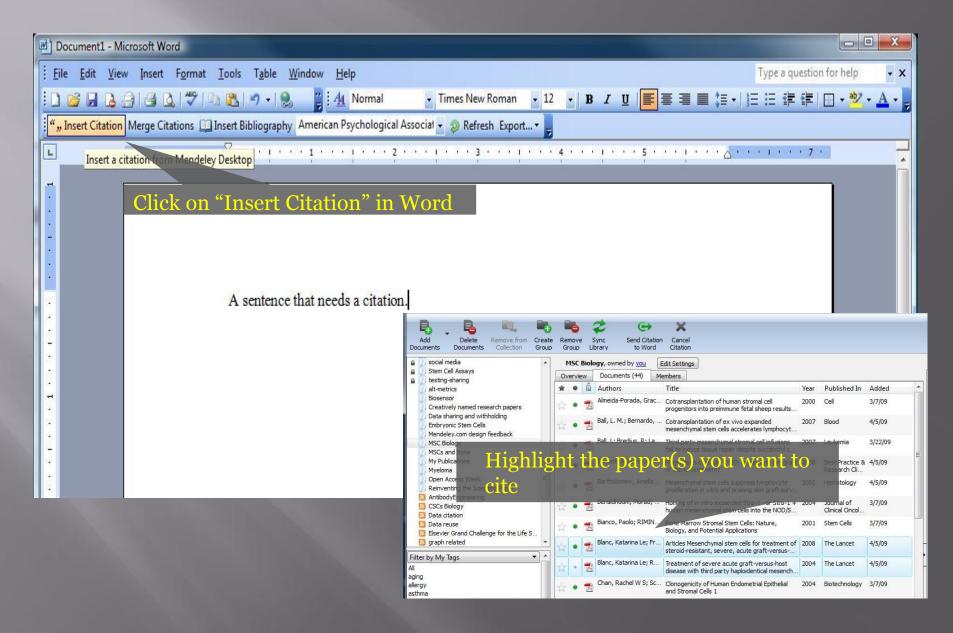
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We chose the mother as donor, because she was readily available and because MHC-compatibility is not necessary for mesenchymal stem-cell immunosuppression.3 After ethicscommittee approval and informed consent, mesenchymal stem cells were isolated as previously described.3 Briefly, we isolated mononuclear cells from a bone marrow aspirate by Percoll gradient centrifugation (Sigma, St Louis, MO, USA). The cells were plated at a density of 3×107 cells per 175 cm2 in polystyrene flasks in low glucose Dulbecco's modified Eagle's medium (Life Technologies, Gaithersburg, MD, USA), supplemented with 10% fetal

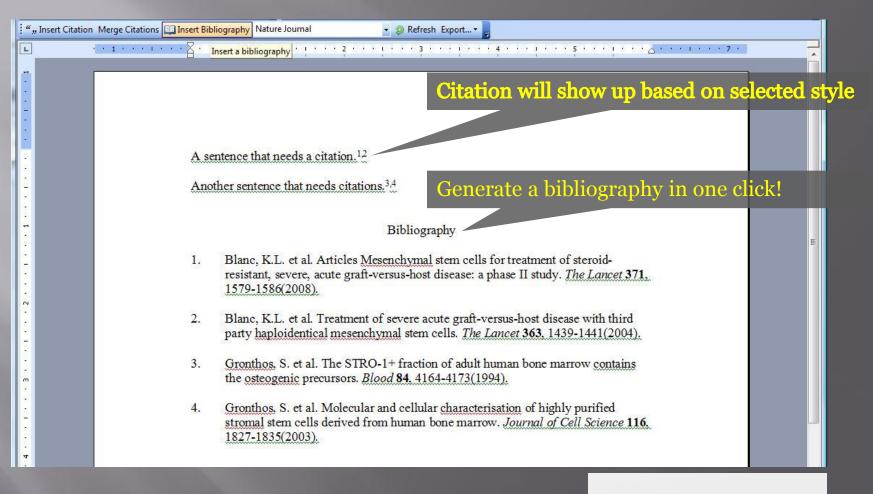


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View the group online – other users can request to join or simply follow the activity of the group

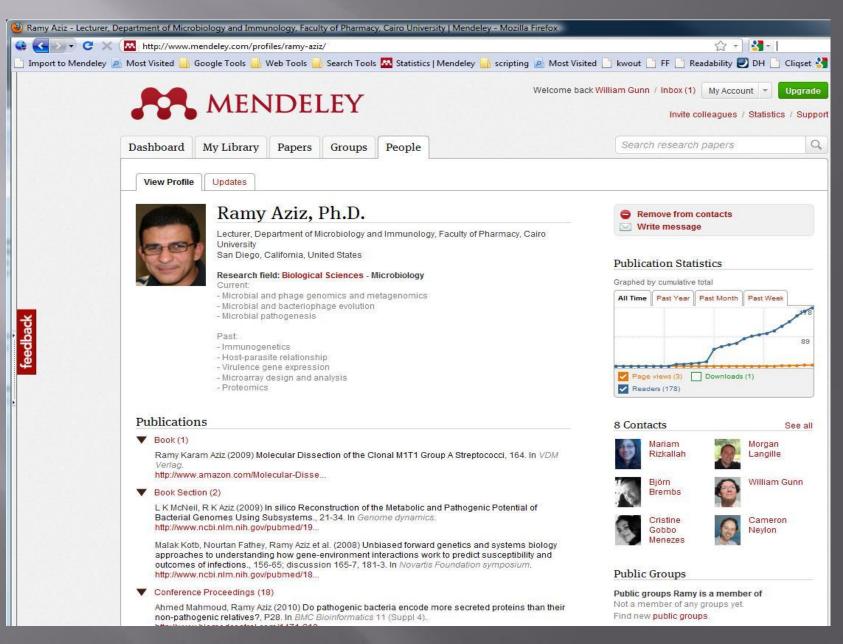
Create Groups in Mendeley Desktop

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Create Groups in Mendeley Desktop

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