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# Mesenchymal Stem Cells Localize to Developing Saccular Aneurysms

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## Introduction

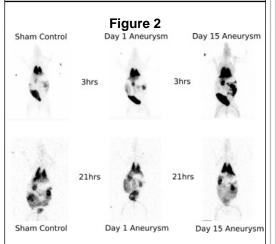
- Mesenchymal stem cells (MSCs) are multipotent immunomodulatory cells
- MSCs promote healing of formed aneurysms (1-3)
- Our lab has previously shown that IV MSC adminstration during saccular aneurysm formation inhibits tunica intima hyperplasia
- MSCs also altered serum cytokines
- Objective: to determine whether or not MSCs localize to the developing aneurysm

## **Methods**

- 8 rabbits randomly assigned to 2 control groups (1 rabbit each), a sham surgery group (2), an early aneurysm group (2) and a late aneurysm group (2)
- Aneurysm groups underwent elastase-induced surgical creation of right common carotid artery (CCA) saccular aneurysm; sham surgery rabbits had right CCA dissected but not manipulated
- Technetium-99m (Tc) used to tag MSCs; observed systemic distribution with planar gamma camera
- 1 control group received IV Tc only, compared to control group with Tc-MSCs
- All surgical groups received IV Tc-MSCs
- After imaging, tissue harvested for radiation quantification in gamma counter

# Figure 1 Tc Control (1) Tc IV Image @ Counter MSC Control (1) MSC-Tc IV Image @ Harvest for counter Sham Control (2) Day 1 Aneurysm (2) Aneurysm (2) Aneurysm (2) Aneurysm (2) Aneurysm (2) Aneurysm (2)

Experimental plan for in vivo MSC localization study. Eight rabbits were randomly assigned to 5 groups. Planar gamma camera imaging with Techentium-99m (Tc). Radiation in harvest tissues was quantified with gamma counter.

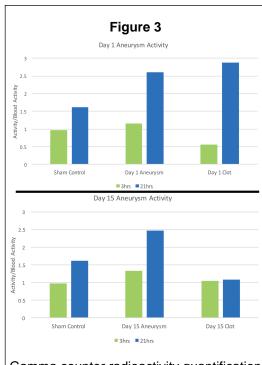


Planar scintigraphy of sham control, early aneurysm and late aneurysm rabbits at 3hrs and 21hrs after Tc-MSC injections showing similar MSC distributions.

Harvested tissues showed localization of MSCs to developing aneurysms.

## **Results**

- Systemic distribution of Tc-MSCs differed from Tc, indicating stable Tc tag
- Tc-MSCs localize largely to lungs
- Tc-MSC distribution similar in all surgical groups (Figure 2)
- Gamma counter showed small degree of MSC localization to early and late developing aneurysms and acute intraaneurysmal thrombus (Figure 3)



Gamma counter radioactivity quantification of harvested tissues relative to arterial blood radioactivity. (Top) Day 1 Aneurysm and (Bottom) Day 15 Aneurysm rabbits.

## **Conclusions**

- MSCs injected IV become entrapped in the lung
- Portion of MSCs traverse and localize to developing aneurysm and acute intra-aneurysmal thrombus
- MSC effects on developing aneurysm likely occur through combination of local release of factors and secretion of factors from within pulmonary capillary bed

# References

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- 3. Rouchaud A, Journe C, Louedec L, Ollivier V, Derkaoui M, Michel JB et al. Autologous mesenchymal stem cell endografting in experimental cerebrovascular aneurysms. *Neuroradiology*. 2013. 55(6):741-9.

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