

# SRI 64th Annual Scientific Meeting

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**Presenting/Contact Author:** Sonia Herraiz

**Department/Institution:** Fundación IVI

**Address:** C/ Catedrático Agustín Escardino nº 9, PARC CIENTIFIC  
UNIVERSITAT DE VALENCIA Edificio 3, CUE. 2ª Planta.

**City/State/Zip/Country:** Paterna, Valencia, Spain

**Phone:** +34626347082 **Fax:** **E-mail:** Sonia.Herraiz@ivi.es

**Keyword 1:** Bone Marrow Derived Stem Cells

**Keyword 2:** Ovarian rejuvenation

**Keyword 3:** Ovarian niche vascularization

**1. Abstract Categories:** 15.5. Regenerative Medicine

**2. Previously Presented:**

Has this abstract been previously presented as it is written? No

Has this abstract been partially presented? No

Presentation Date:

Where was this abstract presented:

**3. Data Requirement Questions**

**My submitted abstract(s) contains original data, written in standard scientific form, complete with numeric values and statistical analyses when appropriate:** Yes

**If my abstract contains microarray data, all analyses must be accompanied by confirmation of expression changes with either transcript or protein data:** Not Applicable

**All data derived using the same paradigm (set of patients or experiments) will not be separated into multiple abstracts:** Yes

**I understand that failure to comply with said requirements will result in abstract dismissal:** Yes

**4. I will comply with the SRI Abstract Withdrawal Policy:** Yes

**Title: Infusion Of Human Bone Marrow-Derived Stem Cells Improved Ovarian Function In Chemotherapy-Damaged Ovaries In Mice.**

Sonia Herraiz, †<sup>1</sup>, Anna Buigues, †<sup>1</sup>, Mónica Romeu<sup>1</sup>, César Díaz-García<sup>1</sup>, Aaron J Hsueh<sup>2</sup> and Antonio Pellicer<sup>1</sup>. <sup>1</sup>Fundación IVI, IIS la Fe, Valencia,

Spain and <sup>2</sup>School of Medicine, Stanford University, Stanford, CA, United States.

**Introduction:** Aging, poor ovarian response and other damaging-acquired conditions like oncologic treatments, lead to an impaired ovarian function. Nevertheless, even when the damaged ovaries have lost their ability to ovulate, they might contain a residual pool of quiescent follicles that could be activated to growth. Infusion of bone marrow-derived stem cells(BMDSC) could provide an ovarian niche for follicular rescue and rejuvenation.

**Objective:** To assess regenerative effects of human BMDSC on chemotherapy-damaged ovaries in mice.

**Methods:** Twelve 8-week old female NOD-SCID mice were treated with two different chemotherapy regimens to induce ovarian damage. Standard treatment (1xChT,n=6) consisted of a single injection of 12mg/kg busulfan (Bu) and 120mg/kg Cyclophosphamide (Cy) while the reduced dose (0.1xChT,n=6) was 1.2mg/kg Bu+12mg/kg Cy. A week later (Day 0) animals from both groups were randomized to receive an injection of PBS (Control,n=3each) or  $1 \times 10^6$  Human Rhodamine B-labeled BMDSC (BMDSC groups) via tail vein. Controlled ovarian stimulation was induced on day 14 with 10IU of PMSG+hCG. Then ovaries were recovered to evaluate follicle growth, proliferation, apoptosis and vascularization.

**Results:** In the standard chemotherapy groups, antrum cavity formation (BMDSC:10.6±1.8% vs. Control:5.9±2.6%,p=0.01) as well as % of pre-ovulatory follicles (1.6±0.9% vs. 0.2±0.1% respectively,p<0.01) were increased in mice receiving BMDSC when compared to controls. In the reduced dosage groups, BMDSC also increased antrum (BMDSC:8.9±5.2% vs. Control:5.2±1.9%,p=NS) and pre-ovulatory follicles (1.7±0.9 vs. 0.5±0.8 respectively,p=0.04).

When ovarian stroma was examined, improvement in micro-vessel density (1xChT-BMDSC:5.1±0.8% vs. 1xChT-Control:1.8±0.9%,p=0.01), increases in cell proliferation (2.3±0.5% vs. 1.0±0.3%,p=0.04) and decreases in apoptosis (0.5±0.2% vs. 5.1±3.6%,p=0.04) were detected after BMDSC infusion in the standard dose. In the 0.1xChT dose, BMDSC also increased cell proliferation (BMDSC:1.4±0.4 vs. Control:0.7±0.2%,p=0.04) but decreased apoptosis (0.5±0.2 vs. 5.9±3.6 respectively,p=0.04).

**Conclusions:** Human BMDSC infusion improved ovarian function by promoting follicular growth to the pre-ovulatory stage, increasing vascularization and cell proliferation as well as suppressing apoptosis in chemotherapy-damaged ovaries in mice.

PROMETEOII/2014/045

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1. **How long have you been a member of the society?** I am not a member
2. **Please tell us what kind of society membership you currently hold**  
In-Training member

3. **What is your primary reason for joining the SRI?**  
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**Mentoring:**  
**Networking:**  
**To present my own data:**
4. **Gender?** Female
5. **Age Range** 36-45
6. **Race/Ethnicity**  
**White/Caucasian:** Yes  
**Black/African American:**  
**Mexican American:**  
**Latin American:**  
**Hispanic or Latino:**  
**Asian American or Pacific Islander:**  
**Asian:**  
**American Indian or Alaskan Native:**  
**Other (please specify):**
7. **Under-Represented Minority:** No
8. **Degree**  
**M.D.:**  
**Ph.D.:** Yes  
**D.Phil.:**  
**M.Sc.:**  
**D.Sc.:**  
**BA/BS:**  
**Other (please specify):**
9. **Career Sector:** In-training
10. **Moderating a concurrent session** No
11. **Academic Status** Postdoctoral Fellow
12. **What percent of time do you spend giving Clinical Care?** 10
13. **Department/ Division** Basic Science Department
14. **Please list your sub-specialty** Basic Reproductive Sciences
15. **How would you best represent your primary research?** Translational research

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