ROYAN INTERNATIONAL RESEARCH AWARD







In the Name of GOD





Dr Saeid Kazemi Ashtiani The Late Founder of ROYAN Institute



Cover Legend: It Is Not the Moon! It is the migration of Neural Crest Cells emerged from human embryonic stem cells, Royan H6 plated on poly L-ornitin / Laminin coated dish. (Photographer: Ali Fathi- Royan Institute)

Organizer:

Royan Institute

Street Address: East Hafez Alley, Banihashem Square, Tehran, Iran

Post Address:

P. O. Box: 16635-148, Tehran,

Phone: +98 (21) 22 33 99 36 **Fax:** +98 (21) 22 33 99 58 E-mail: info@royaninstitute.org

- Rahim Tavassolian
- Editor: Sima Farrokh
- Technical-Artwork Editor: Hassan Moghimi
- Graphic Designer: Mohammad Abarghouei
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On behalf of the Iranian Academic Center for Education, Culture and Research (ACECR) I take this opportunity to commemorate the late Dr Saeid Kazemi Ashtiani, the founder of Royan Institute whose endeavor along with his colleagues led to holding Royan Research Award in the field of infertility and stem cell research.

Organizing Royan 16th International Research Award and making it generally accepted by the national and international researchers is a confirmation to this event's success till now.

Royan Institute as one of the successful research centers of ACECR has brilliant services and records in national and international level in Reproductive Biomedicine, Stem Cell Research, and Animal Biotechnology. In addition, through holding Royan Research Award and the Congresses besides, it attempts to enhance the scientific level of the aspiring young of knowledge and science within the country and establish fruitful international relationships.

I, hereby, appreciate my colleagues at Royan Institute who every year hold this great scientific event. I hope this scientific movement, whose leaders are prominent Iranian researchers, be effective in promoting the science and solving the society's health problems.

Tayyebi HR, PhD

President of Academic Center for Education, Culture and Research (ACECR)



Introduction



Fortunately, Royan 16th international research award has been accompanied with the agreement made between our country and the delegates from international community in regard with peaceful nuclear program. And since the first Royan Award prize giving ceremony coincided with the unpleasant terrorist news of September 11th, 2000, it left a bad effect on attracting many international scientists through the years. However, we should certainly appreciate the scientists who despite such negative and inappropriate propaganda, have attended this event as well as Royan International Congress these years. So, with the improvement of political situation, we hope to see more scientists make their presence in the events of this kind in Iran, and the most important, to establish principle scientific collaborations.

Royan award has been established with the aim of creating appropriate scientific relationships with universities and research centers as well as encouraging young researchers to follow scientific and innovative activities. With great fortune, despite all aforementioned problems, Royan Award has established close relationship with the scientists and researchers worldwide. Also, the presence of award winners in Iran along with having a visit at Royan Institute has had proper feedbacks like the one by Dr Sophie Rousseaux published in the international journal of developmental biology which led the scientists in the world to knowing more about Iran's scientific movement.

This year, award secretariat has received 204 projects from about 47 countries. As previous years, the projects were evaluated by national and international referees and finally 5 international and 2 national winners were announced which will be awarded through the ceremony.

Another important issue in this year, is the presence of Prof Robert Langer's at Royan to receive the third Kazemi Prize. Kazemi Prize is an award to commemorate the efforts made by Dr Saeid Kazemi Ashtiani; the late founder of Royan Institute and revive his memories. It is established to make the young, who love science, familiar with the current scientific brilliants and geniuses in the world. We do deeply appreciate prof Langer for accepting our invitation, despite his busy schedule and remote distance to Iran.

Royan Award owes its success to strong and perseverant executive board, bound and prominent jury board, and the participants in this event. I wish to extend my gratitude to all of them and hope to replace the issues which endanger international peace and stability with the scientific competitions; like this.



Royan International Research Award was founded by the late director of Royan Institute, Dr Saeid Kazemi Ashtiani with the aim of encouraging researchers, appreciating their efforts and preparing a friendly scientific atmosphere for them to exchange their knowledge and experiences. Kazemi had wonderful ideas to bring researchers together and motivate them to increase their efforts and perform high level researches via this research award. Royan's staff lost their beloved director in January 2006 by heart attack, May he rest in peace.

This annual award is extending into a higher quality event every year, increasing the scientific level and number of the submitted papers. The research papers are evaluated through an intense jury procedure by Award's national and international Jury board to whom our special thanks goes. Each year the prominent researches with outstanding help in solving problems in reproduction and stem cell fields, are announced, appreciated and rewarded.

As comparing the researches in different fields is very difficult and finding the best researches with variations in methods, implements and results is almost impossible, from the eighth award the same prizes are distributed among winners in different fields of reproductive biomedicine and stem cell such as: female infertility, epidemiology, ethics, andrology, embryology, reproductive imaging, reproductive genetics, stem cell biology and technology, regenerative medicine, and biotechnology.

Nomination and Selection Procedure of Award

The submitted research articles are categorized according to nine scientific groups: female infertility, reproductive genetics, epidemiology, ethics, embryology, andrology, reproductive imaging, stem cell biology and technology and biotechnology. Each article is ranked according to its relevancy, impact factor, and an innovation score.

After the articles are sorted, each scientific group selects their nominees and sends them to national and international referees for evaluation.

Each referee evaluates at most 5 research articles, related to his/her field of interest, qualitatively in Likert scale according to these norms:

- Relevancy to the award subjects
- Creativity and innovation
- Methodology and research design
- Problem solving
- Applicability on human

Evaluation of the articles by the juries has been discussed in the board of juries and their decisions have been approved by scientific board of the institute. Finally, international and national winners are selected and invited to present their researches in Royan twin congress on Reproductive Biomedicine and Stem Cell Biology and Technology which is held almost in September every year and will receive their prizes in prize award ceremony.

Note: It is obligatory for the winners to attend the ceremony and present their research articles in the congress.



The First Royan International Research Award | September 2000 | Received Papers: 72





Reproductive Biomedicine & Stem Cell



First Place: Mohamed Mitwally, Canada
 Comparison of an Aromatase Inhibitor with Clomiphene Citrate for Induction of Ovulation

Second Place: Ali Ahmady, Canada
 Cell and Molecular Investigation of the Fertilizing Ability of Dead Sperm

• Third Place: Weihau Wang, USA
Spindle Observation in Living Human Eggs with Pollaries Microscope and Its Use in Assisted Human
Reproduction

• Fourth Place: Simon Marina Avendano, Spain HIV-Seropositive Can Be Fathers without Infecting the Women or Child

• Fifth Place: Jaffar Ali, Qatar Formulation of a Protein-Free Medium for Human Assisted Reproduction

Iranian Winners:

Mohammad Hossein Nasr-Esfahani
 Sperm Chromatin Status and Male Infertility

Mahnaz Ashrafi
 Effect of Metformin on Ovulation and Pregnancy Ratein Women with Clomiphen Resistant PCOS

Mohammad Ebrahim Parsanezhad
 Section of the Cervical Septum Doesn't Impair Reproductive Outcome



The Second Royan International Research Award | September 2001 | Received Papers: 78









m september 2001

Reproductive Biomedicine & Stem Cell

International Winners:

• First Place: Ri-Cheng Chian, Canada

A New Treatment for Women with Infertility Due to Polycystic Ovarian Syndrome: Immature Oocyte Retrieval Followed in-vitro Maturation

- Second Place: Ma'asouma Makhseed, Kuwait
 The Possible Immunological Basis of Repeated Pregnancy Loss
- Third Place: Esmail Behboodi, USA
 Production of Goats by Somatic Cell Nuclear Transfer
- Fourth Place: Sayeed Unisa, India
 Reproductive, Demographic and Behavioral Causes of Infertility in India
- Fifth Place: Ahmed Mohammed Saleh, Saudi Arabia Effect of Laparoscopic Ovarian Drilling on Serum Vascular Endothelial Growth Factor (VEGF), and on Insulin Response to Oral Glucose Tolerance Test in Women with PCOS

Iranian Winners:

Hossein Baharvand

Improvement of Blastocyst Development in-vitro and Overcoming the Blastocyst Collapse and Its Effective Factor(s) in Sequential Culture Media

Marzieh Nojomi
 Epidemiology of Infertility in the West of Tehran 2000-2001

Gholamreza Pourmand
 Effect of Renal Transplantation on Sperm Quality and Sex Hormones Level



Page

The Third Royan International Research Award | September 2002 | Received Papers: 212









September 2002

Reproductive Biomedicine & Stem Cell

International Winners:

- First Place: Marco Filicori, Italy
 Novel Approaches to Ovulation Induction: The Critical Role of Luteinizing Hormone Activity in Regulating
 Folliculogenesis
- Second Place: Klaus G. Steger, Canada Influence of Histone-Protmine-Exchange on Male Infertility
- Third Place: Franck Pellestor, France Chromosomal Investigations in Human Gametes: Study of the Interchromosomal Effect in Sperm of Chromosomal Rearrangement Carriers and Mechanisms of Non Disjunction in Oocytes
- Fourth Place: Ghazala S. Basir, Hong Kong
 The Effect of High Estradiol Levels on Endometrial Development in Assisted Reproduction Technology:
 Evaluation of Sonographic Doppler Haemodynamic and Morphometric Parameters
- Fifth Place: Mohamed Ali Bedaiwy, USA
 Transplantation of Intact Frozen-Thawed Mammalian Ovary with Vascular Anastomosis: A Novel Approach

Iranian Winners:

- Saeed Alborzi
 Laparoscopic Salpingoovolysis. Is There Any Place for Second Look Laparoscopy?
- Saeed Rahbar
 Laser Assisted Hatching in Young Women Significantly Increases Pregnancy and Implantation Rates
- Shir Ahmad Sarani
 Morphological Evidence for the Implantation Window in Human Luminal Endometrium Special Winner in Reproductive Health
- V. I. Sodestrom- Anttila, Finland
 Embryo Donation-Outcome & Attitude Among Embryo Donors & Recipient



The Fourth Royan International Research Award | September 2003 | Received Papers: 222







September 2003

Reproductive Biomedicine & Stem Cell

International Winners:

- First Place: Yong-Mahn Han, South Korea Abnormal Structural Integrity and Reprogramming in the Cloned Embryos
- Second Place: Lucille E. Voullaire, Australia Chromosome Abnormality In Human Embryos Diagnosed Using Comparative Genomic Hybridization: Its Relationship to Infertility
- Third Place: Mauro Maccarrone, Italy Low Fatty Acid Amide Hyrolase and Anandamide Levels Are Associated with Failure to Achieve an Ongoing Pregnancy after IVF and Embryo Transfer
- Fourth Place: Ali Honaramooz, USA Sperm from Neonatal Mammalian Testes Grafted in Mice
- Fifth Place: Jan M.R. Gerris, Belgium Elective Single Embryo Transfer Halves the Twinning Rate without Decrease in the Total Ongoing Pregnancy Rate of an AVF/ICSI Program

Iranian Winners:

- Mohammad Ebrahim Parsanezhad
 - Ovarian Stromal Blood Flow Changes After Laparoscopic Ovarian Cauterization in Women with Polycystic Ovary Syndrome
- Mojdeh Salehnia
 Vitrification of Ovarian Tissue
- Jaleh Zolghadri

Successful Pregnancy Outcome with IUI in Patients with Unexplained Recurrent Miscarriage, Whose Male Partners Have Low Score Hypo-Osmotic Swelling Test





The Fifth Royan International Research Award | September 2004 | Received Papers: 199









September 2004

Reproductive Biomedicine & Stem Cell

International Winners:

- Second Place: Alfonso Guiterrez-Adan, Spain
 Long Term Effect of in vitro Culture of Mouse Embryos with Serum on mRNA Expression of Imprinting Genes,
 Development and Behavior
- Second Place: Maciej K. Kurpisz, Poland
 Reactive Oxygen Species and "Male Factor" of Infertility
- Third Place: Michel von Wolf, Germany
 Glucose Transporter Proteins (GLUT) in Human Endometrial-Expression, Regulation and Function through
 out the Menstrual Cycle and in Early Pregnancy
- Fourth Place: Sophie Lambard, France
 Human Male Gamete Quality: Place of Aromatase and Estrogens
- Fifth Place: Naojiro Minami, Japan A Novel Maternal Effect Gene, Oogenesin: Involvement in Zygotic Gene Activation and Early Embryonic Development in the Mouse

Iranian Winners:

- Seyed Javad Mowla
 - Catsper Gene Expression in Postnatal Development of Mouse Testis and in Subfertile Men with Deficient Sperm Motility
- Mohammad A. Khalili
 Restoration of Spermatogenesis by Adenoviral Gene Transfer into Injured Spinal Cords of Rats
- Mojdeh Salehnia
 Ultrastructural, Histochemical and Morphometric Studies of Mouse Reproductive Tract after Ovarian Induction



The Sixth Royan International Research Award | September 2005 | Received Papers: 198









= 3eptember 2003

Reproductive Biomedicine & Stem Cell

International Winners:

- First Place: Kathyjo Ann Jackson, USA Therapeutic potential of stem cells
- Second Place: Carmen Belen Martinez-Madrid, Belgium
 Ficoll Density Gradient Method for Recovery of Isolated Human Ovarian Primordial Follicles
- Third Place: Federico Alejandra Calegari, Germany
 Tissue-Specific Manipulating of Gene Expression of Mouse Embryos Using in Utero Electroporation
- Fourth Place: Maryam Kabir-salmani, Japan Different Roles of $\alpha_5\beta_1$ and $\alpha_\nu\beta_3$ Integrins in the IGF-I-Induced Migration of the Human Extravillous Trophoblast Cells
- Fifth Place: Zhenmin Lei, USA
 Testicular Phenotype in Luteinizing Hormone Knockout Animals and the Effect of Testostrone Replacement
 Therapy

Iranian Winners:

- Seyed Javad Mowla
 - The Profile of Gene Expression Changes During the Neural Differentiation of Bone Marrow Stormal Cells (BMSCs)
- Jaleh Zolghadr

Pregnancy Outcome Following Laparoscopic Tubal Ligation of Hydrosalpinx Tube in Patients with Early Recurrent Abortion

Finally, this year we got more papers and the jury procedure was more difficult. The papers were very close together in scientific level, so a hairsplitting jury procedure was needed to find out the best of them.



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The Seventh Royan International Research Award | September 2006 | Received Papers: 221







■ September 2006

Reproductive Biomedicine & Stem Cell

International Winners:

• First Place: James Affram Adjaye, Germany

A) Whole-Genome Approaches for Large-Scale Gene Identification and Expression Analysis in Mammalian Preimplantation Embryos & B) Primary Differentiation in the Human Blastocyst: Comparative Molecular Portraits of Inner Cell Mass and Trophectoderm Cells

• Second Place: Tian-hua Huang, China

Detection and Expression of Hepatitis B Virus X Gene in One and Two-Cell Embryos from Golden Hamster Oocytes in-vitro Fertilized with Human Spermatozoa Carrying HBV DNA

• Third Place: Adrian Richard Eley, UK

Opoptosis of Ejaculated Human Sperm Is Induced by Co-Incubation with Chlamydia Trachomatis Lipopolysaccaride

• Fourth Place: Lone Schmidt, Denmark

Does Infertility Cause Marital Benefit? An Epidemiological Study of 2250 Women and Men in Fertility Treatment

• Fifth Place: Louis Chukwuemeka Ajonuma, Hong Kong

Molecular and Cellular Mechanisms Underlying Abnormal Fluid Formation in the Female Reproductive Tract: The Critical Role of Cystic Fibrosis Transmembrane Conductance Regulators

Iranian Winners:

Mohammadreza Baghban Eslaminejad

Polarized Culture Systems and Their Effects on Embryo Development

Mansoureh Movahedin

New Approaches to Assess the Success and Enhance the Efficiency of Male Germ Cell Transplantation in the Mouse

Ashraf Alleyassin

Comparison of Unilateral and Bilateral Transfer of Injected Oocytes into Fallopian Tubes: A Prospective Randomized Clinical Trial



The Eighth Royan International Research Award | September 2007 | Received Papers: 248





December 1997

Reproductive Biomedicine & Stem Cell

International Winners:

Best research project in stem cell field

 Chiba Shigeru, Japan
 Role of Notch Signaling in Normal and Neoplastic Hematopoietic Stem Cells and Clinical Application of Notch Signal Modifiers

Best research project in reproductive genetics field

Françoise Dantzer, France
 Poly (ADP-Ribose) Polymerase-2 Contributes to the Fidelity of Male Meiosis I and Spermiogenesis

Best research project in female infertility field

• Seyed Mohammad Moazzeni, Iran
Dendritic Cells and Pregnancy: A Bidirectional Relationship to Protect the Semiallogenic Fetus

Best research project in embryology field

 Bjorn Johannes Oback, New Zealand
 Nuclear Donor Choice, Sperm Mediated Activation and Embryo Aggregation: A Multi-Pronged Approach to Sequentially Improve Cattle Cloning Efficacy

Best research project in andrology field

• Reddanna Pallu, India
Role of Cyclooxygenases in Male Reproduction

Iranian Winners:

Ramin Radpour

Novel Mutations and (TG)M(T)N Polymorphism in Iranian Males with Congenital Bilateral Absence of the Vas Deferens

Mohammad Ebrahim Parsanezhad
 Hysteroscopic Metroplasty of the Complete Uterine Septum, Duplicate Cervix, and Vaginal Septum

Mehri Azadbakht

Apoptosis in Mouse Embryos Co-Cultured with Polarized or Non-Polarized Uterine Epithelial Cells Using Sequential Culture Media





The Ninth Royan International Research Award | September 2008 | Received Papers: 202









December 19 and 19 and

International Winners:

Best research project in stem cell field

Su-Chun Zhang, USA
 Human Embryonic Stem Cells As a Tool of Discovery

Best research project in reproductive genetics field

• Smita Mahale, India
Structural, Functional and Molecular Aspects of Follicle Stimulating Hormone Receptor: Applications in Designing Receptor Targets and Management of Female Infertility

Best research projects in female infertility field (share)

- Federico Prefumo, Italy
 Uterine Doppler Investigations and Trophoblast Biology in Early Pregnancy
- Saeed Alborzi, Iran
 Laparoscopic Metroplasty in Bicornuate and Didelphic Uterus

Best research project in embryology field

• Leen.Vanhoutte, Belgium Nuclear and Cytoplasmic Maturation of in vitro Matured Human Oocytes After Temporary Nuclear Arrest by Phosphodiesterase 3-Inhibitor

Best research project in andrology field

• T.O.Ogata, Japan Haplotype Analysis of the Estrogen Receptor Alpha Gene in Male Genital and Reproductive Abnormalities

Iranian Winners:

Ali Fathi

The Molecular Mechanisms Controlling Embryonic Stem Cells (Escs) Proliferation and Differentiation

Fardin Fathi

Characterizing Endothelial Cells Derived from the Murine Embryonic Stem Cell Line CCE



The Tenth Royan International Research Award | September 2009 | Received Papers: 253







International Winners:

Best research project in stem cell field

• Yi Liu, China Dental Stem Cells-Based Tissue Regeneration in a Large Animal Model

Best research project in reproductive genetics field

 Wai-sum OO, China Adrenomedullin in Male and Female Reproduction

Best research projects in female infertility field (share)

• Sherman Silber, USA

A Series of Monozygotic Twins Discordant for Ovarian Failure: Ovary Transplantation (Cortical versus Microvascular) and Cryopreservation

What Harbours the Cradle of Life? The Progesterone-Dependent Immunomodulation

Best research project in embryology field

Geetanjali Sachdeva, India

• Melinda Halasz, Hungary

Molecular Assessment of the Uterine Milieu during Implantation Window in Humans and Non-human Primates

Best research project in andrology field

Paolo Chieffi, Italy

PATZ1 Gene Has a Critical Role in the Spermatogenesis and Testicular Tumours

Iranian Winners:

Hossein Mozdarani

Reduction of Induced Transgenerational Genomic Instability in Gametes Using Vitamins E and C, Observed As Chromosomal Aneuploidy and Micronuclei in Preimplantation Embryos

Seyed Javad Mowla

OCT4 Spliced Variants Are Differentially Expressed in Human Pluripotent and Nonpluripotent Cells

Mohammad Reza Safarinejad

Evidence Based Medicine on the Pharmacologic Management of Premature Ejaculation



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The Eleventh Royan International Research Award | September 2010 | Received Papers: 358







Reproductive Biomedicine & Stem Cell

International Winners:

Best research project in regenerative medicine field

• Stefano Pluchino, Italy
Human Neural Stem Cells Ameliorate Autoimmune Encephalomyelitis in Non-human Primates

Best research project in stem cell biology & technology field

Hooman Sadri-Ardekani, Iran-The Netherlands
 Propagation of Human Spermatogonial Stem Cells in vitro

Best research project in female infertility field

Louis Chukwuemeka Ajonuma, Nigeria
 New Insights into the Mechanisms Underlying Chlamydia Trachomatis Infection Induced Female Infertility

Best research project in reproductive genetics field

 Anu Bashamboo, France Mutations in NR5A1 Associated with Ovarian Insufficiency

Best research project in embryology field

Mohammad Hossein Nasr-Esfahani, Iran
 New Era in Sperm Selection for ICSI Procedure

Iranian Winners:

Serajoddin Vahidi

Prevalence of Primary Infertility in the Islamic Republic of Iran in 2004-2005

Tahereh Ma'dani

Improvement of Pregnancy Rate in ART Cycles

Mehrdad Noruzinia

MTHFR Promoter Hypermethylation in Testicular Biopsies of Patients with Non-obstructive Azoospermia: The Role of Epigenetics in Male Infertility

Abbas Piryaei

Differentiation Capability of Mouse Bone Marrow-Derived Mesenchymal Stem Cells into Hepatocyte-Like Cells on Artificial Basement Membrane Containing Ultraweb Nanofibers and Their Transplantation into Carbon Tetrachloride Injured Liver Model



The Twelfth Royan International Research Award | September 2011 | Received Papers: 280







■ September 2011

Reproductive Biomedicine & Stem Cell

International Winners:

Best research project in regenerative medicine field

• Lorenzo Piemonti, Italy

Bone Marrow as Ideal Microenvironment for Human Islet Transplantation to Treat Type 1 Diabetes
(ClinicalTrials.gov Identifier: NCT01345227)

Best research project in stem cell biology & technology field

Hiromitsu Nakauchi, Japan
 Heterogeneity and Hierarchy Within the Most Primitive Hematopoietic Stem Cell Compartment

Best research project in female infertility field

• Elizabeth Stewart, USA
Safely Extending Focused Ultrasound Surgery for Uterine Leiomyomas to Women Who Desire Future
Pregnancies

Best research project in reproductive genetics field

Paul Thomas, Australia
 Identification of SOX3 As an XX Male Sex Reversal Gene in Mice and Humans

Best research project in embryology field

Steve Tardif, UK
 Infertility with Impaired Zona Pellucida Adhesion of Spermatozoa from Mice Lacking TauCstF-64

Best research project in epidemiology & ethics fields

Heping Zhang, USA

Decision Trees for Identifying Predictors of Treatment Effectiveness in Clinical Trials and Its Application to Ovulation in a Study of Women with Polycystic Ovary Syndrome

Iranian Winners:

Morteza S. Hosseini
 Development of an Optimized Zona-Free Method of Somatic Cell Nuclear Transfer in the Goat

Jaleh Zolghadri

Relationship Between Abnormal Glucose Tolerance Test and History of Previous Recurrent Miscarriages, and Beneficial Effect of Metformin in These Patients: A Prospective Clinical Study

Batool Rashidi

Simvastatin Effects on Androgens, Inflammatory Mediators, and Endogenous Pituitary Gonadotropins Among Patients with PCOS Undergoing IVF: Results from a Prospective Randomized Placebo-Controlled Clinical Trial



The Thirteenth Royan International Research Award | September 2012 | Received Papers: 169







International Winners:

Best research project in stem cell biology & technology field

• Chengcheng (Alec) Zhang, USA Ex Vivo Expanded Hematopoietic Stem Cells Overcome the MHC Barrier in Allogeneic Transplantation

Best research project in andrology field

• Kristian Almstrup, Denmark Screening of Subfertile Men for Testicular Carcinoma in Situ by an Automated Image Analysis-based Cytological Test of the Ejaculate

Best research projects in female fertility field (share)

• Wenjie Zhu, China

Transvaginal Ultrasound-guided Ovarian Interstitial Laser Treatment in Anovulatory Women with Polycystic Ovary Syndrome: A Randomized Clinical Trial on the Effect of Laser Dose Used on the Outcome

Kaei Nasu, Japan

Role of Mevalonate-Ras Homology (Rho)/Rho-associated Coiled-Coil-Forming Protein Kinase-mediated Signaling Pathway in the Pathogenesis of Endometriosis-associated Fibrosis

Best research project in reproductive genetics field

Signe Atlmäe, Sweden

Interactorme of Human Embryo Implantation: Identification of Gene Expression Pathways, Regulation, and Integrated Regulatory Networks

Best research project in embryology field

Laura Cecilia Giojalas, Argentina

Sperm Chemotaxis towards Progesterone, a Guiding Mechanism That May Be Used to Select the Best Spermatozoa for Assisted Reproduction

Iranian Winner:

Alireza Pouya

Human Induced Pluripotent Stem Cells Differentiation into Oligodendrocyte Progenitors and Transplantation in a Rat Model of Optic Chiasm Demyelination



The Fourteenth Royan International Research Award | September 2013 | Received Papers: 206







September 2013

Reproductive Biomedicine & Stem Cell

International Winners:

Best research project in stem cell biology & technology field

Antonio Uccelli, Italy
 Mesenchymal Stem Cells Shape Microglia Effector Functions Through the Release of CX3CL1

Best research project in reproductive genetics & andrology fields

Pierre F Ray, France
 Search for Genetic Causes of Male Infertility

Best research project in female infertility field

Paola Panina Bordignon, Italy
 The Selective Vitamin D Receptor Agonist Elocalcitol Reduces Development of Endometriosis and Formation of Peritoneal Adhesion in a Mouse Model

Best research project in embryology field

Mariano Buffone, USA
 Role of Actin Cytoskeleton During Mouse Sperm Acrosomal Exocytosis

Iranian Winners:

Ashraf Moini

Risk Factors Associated with Endometriosis Among Iranian Infertile Women

Malek Hossein Asadi

OCT4B1, A Novel Spliced Variant of OCT4, Is Highly Expressed in Gastric Cancer and Acts as an Antiapoptotic Factor

Hossein Mozdarani

Genome Instability and DNA Damage in Male Somatic and Germ Cells Expressed as Chromosomal Microdeletion and Aneuploidy Is a Major Cause of Male Infertility

Armin Towhidi

Omega-3 Fatty Acids Accompanied with A-Tocopherol Improved Fresh and Post-thaw Sperm Quality in Ruminants





The Fifteenth Royan International Research Award | September 2014 | Received Papers: 222



■ September 2014

Reproductive Biomedicine & Stem Cell

International Winners:

Best research project in regenerative medicine field

Anne S. Baron-Van Evercooren, France
 Role of Endogenous Neural Precursor Cells in Multiple Sclerosis

Best research project in stem cell biology & technology field

Milena Bellin, Netherlands
 Human Pluripotent Stem Cells for Modelling and Correcting Long-QT Syndrome

Best research project in andrology & reproductive genetics fields

Sophie Rousseaux, France
 Male Genome Programming, Infertility and Cancer

Best research project in female infertility field

Christiani Andrade Amorim, Belgium
 New Steps Towards the Artificial Ovary

Best research project in embryology & biotechnology fields

Guoping Fan, USA
 Transcriptome Dynamics of Human and Mouse Preimplantation Embryos Revealed by Single Cell RNA-sequencing

Best research project in ethics field

Kristien Hens, Netherlands
 Towards the Transparent Embryo? Dynamics and Ethics of Comprehensive Pre-implantation Genetic Screening

Iranian Winners:

Seyedeh Nafiseh Hassani

The Augmented BMP Pluripotency Pathway via TGF- β Suppression Maintains the Ground State of Embryonic Stem Cells Self-Renewal

Rouhollah Fathi

Optimal Strategy Toward Fertility Preservation: In vivo and in vitro Post-thaw Options in Gamete, Embryo and Ovarian Tissue Cryostorage



No	Name	Country	Title
1	Abe, Toshiaki	Japan	Controlled Multi-drug Release by a Novel Polymeric Device for Local Lesions
2	Aberdam, Daniel	France	Pluripotent Stem Cells as Therapeutic Tools for Limbal Stem Cell Deficiencies and Drug Testing
3	Al-Ali, Haifa Kathrin	Germany	Allogeneic Stem Cell Transplantation for Myelofibrosis in the JAK1/JAK2 Inhibitor Era
4	Alborzi, Saeed	Iran	A Comparison of Pelvic Magnetic Resonance Imaging, Trans-vaginal and Trans-rectal Sonography with Laparoscopic Findings in the Diagnosis of Deep Infiltrating Endometriosis
5	Alvarez-Buylla, Arturo	USA	Supraependymal Axons Influence Neural Stem Cells in the Adult Brain
6	Amarasekara, Dulshara	Sri Lanka	Cost-Effective Diagnosis of Male Oxidative Stress Using the Nitroblue Tetrazolium Test: Useful Application for the Developing World
7	Andric, Silvana	Yugoslavia	Systemic in vivo Blockade of α 1-Adrenergic Receptors Mitigate Stress-Triggered Disturbance of Camp and Cgmp Signaling in Testosterone-Producing Leydig Cells
8	Aplin, John	UK	Use of in vitro Models to Discover Cellular and Molecular Mechanisms of Embryo Implantation
9	Arnaout, M. Amin	USA	Stress Hematopoiesis Is Regulated by the Kruppel-Like Transcription Factor ZBP-89
10	Ashton, Randolph	USA	Clinically Translatable Protocols for Deriving a Spectrum of Region-Specific Neural Stem Cells
11	Asimakopoulos, Byron	Greece	Conventional Semen Parameters and DNA Fragmentation
12	Aumari, Amira	Syria	The Experimental Contribution for Neural Differentiation Study of Mouse Embryonic Stem Cells: Sox1 (46C) in vitro
13	Bae, Jeehyeon	Korea	Positive Cross Talk Between FOXL2 and Antimüllerian Hormone Regulates Ovarian Reserve
14	Banihani, Saleem	Jordan	Cryoprotective Effect of I-Carnitine on Motility, Vitality and DNA Oxidation of Human Spermatozoa
15	Bartlewski, Pawel	Canada	Correlations Among Ultrasonographic and Microscopic Attributes of the Testis
16	Behrens, Axel	UK	Generation of Functional eta -cells from Adult Pancreatic Ductal Cells
17	Betts, Dean	Canada	Post-transcriptional Regulation of Telomerase Isoforms in Human Embryonic Stem Cells Cultured Under Varying Oxygen Microenvironments
18	Betts, Dean	Canada	Microenvironmental Regulation of Telomerase Isoforms in Human Embryonic Stem Cells
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Endometrial Secretome and Its Role in Uterine Functions

Objective:

Earlier it was believed that uterine fluid (uf) or secretions are not relevant after the embryo implantation in humans. However, recent reports suggest that uterine secretions continue to play important role till the first trimester of human pregnancy. Evidences also suggest that uf mirrors endometrial functions or dysfunctions. Considering the relevance of uf in endometrial functions, studies were undertaken 1) to develop human uf proteomes;2) to identify uf proteins that display differential abundance during the receptive phase and also 3) to assess the functional relevance of differentially abundant proteins using an animal model.

Material and Methods:

2D-PAGE and gel-free Isobaric Tag for Relative and Absolute Quantitation (iTRAQ) were employed. Samples were collected in the pre-receptive (i.e. day two post-ovulation, n=7) or receptive phase (i.e. day six post-ovulation, n=7) of the menstrual cycles, from regularly cycling healthy fertile women. Receptive phase samples were also collected from women with unexplained infertility. Further, to test the functional significance, rats (Rattus norvegicus) were used as an experimental model. Samples were collected in the proestrous (n=8) and metestrous (n=8) phases of estrous cycles and also from pregnant rats (n=18) during day 3-5 post-coitum (p.c.).

Results:

Our studies demonstrated higher abundance of uf alpha-1 antitrypsin precursor and apolipoprotein A-1 in the receptive phase, than in the nonreceptive phase, in regularly cycling women (Parmar et al, 2008). iTRAQ revealed identities of 127 proteins in the human uf. Of these, 27 proteins displayed differential abundance in the receptive (R) phase, compared to the pre-receptive (PR) phase. High Mobility Group Binding Protein 1(HMGB1), one of the differentially abundant proteins displayed less abundance in the R phase than in the PR phase; in secretions as well as in endometrial tissues. Interestingly rats also revealed a lesser abundance of HMGB1 in the receptive phase, compared to that in the nonreceptive phase uf (Bhutada et al, 2013). Thus, human and rat data indicated an association of endometrial receptivity with a decline in the levels of uf HMGB1. A significant decline was also observed in the expression of endometrial HMGB1 on the day of implantation in pregnant rats. Further, recombinant HMGB1 (0.25-1.5ug/ horn) was administered on day three p.c. in mated rats. The horns administered with HMGB1 showed pregnancy failure, whereas those with saline remained unaffected. This indicated the detrimental effect of an excess of extracellular HMGB1 on pregnancy in rats. Also, morphological changes in the endometrium, an increase in the expression of luminal epithelial NF κ β ; and also various inflammatory molecules such as Receptor for Advanced Glycation End Products, Tumor necrosis factor-alpha and interleukin-6; were observed in HMGB1 treated rats, when compared with untreated rats (Bhutada et al, 2014). Significantly higher expression of endometrial HMGB1 was also observed during the receptive phase in the women with unexplained infertility, compared with healthy proven fertile women.

Conclusion:

Our study, for the first time, employed iTRAQ, a gel-free approach to characterize human uf proteome. The study also demonstrated that an excess of extracellular HMGB1 in the receptive phase induces inflammatory changes in the endometrium which interfere with pregnancy.

Keywords:

Uterine fluid, Proteomics, HMGB1, Inflammation, Pregnancy



Geetanjali Sachdeva, PhD

■ India sachdevag@nirrh.res.in

Dr Geetanjali Sachdeva did doctorate in Human Genetics from Jawaharlal Nehru University, Delhi, India. She worked as a postdoctoral fellow at National Institute of Immunology, New Delhi and also at National Cancer Institute, National Institutes of Health, Frederick, USA. She has been a faculty at National Institute for Research in Reproductive Health, Indian Council of Medical Research, Mumbai since 1997. Her major research interests are the mechanisms contributing to endometrial receptivity, early pregnancy, and endometriosis. Recently her group has ventured into the area of conventional and nonconventional steroid signaling prostate cancer pathogenesis. Dr Sachdeva has 45 peer-reviewed publications and five book chapters to her credit. She has received national and international awards for her research. She is a member of the editorial board of the PLOS One journal. She is also a reviewer for journals like Human Reproduction, Biology of Reproduction, Life Sciences, Reproductive Biomedicine Online and Reproduction.



nternational Winner

Embryology

Page



Priyanka Parte, PhD

India partep@nirrh.res.in

Dr Priyanka Parte achieved her PhD in Biochemistry in the year 2000 after completing two Masters Degrees in Applied Biology and Reproductive Biology, respectively. She began her research career in the area of Neuroendocrinology where worked on neuroendocrine regulation of Hypothalamus-pituitary-gonadal axis and was involved in studying the role of estrogen in spermatogenesis and male fertility. On a Rockefeller Foundation fellowship from Oct 1994-Sept 1996 she worked with Dr Richard Sharpe and Phillipa Saunders at the MRC Reproductive Biology Unit, University of Edinburgh, Scotland, UK, where she studied the effect of endocrine disrupters on spermatogenesis. She did her post-doctoral research with Prof David Kupfer, Dept of Biochemistry and Molecular Pharmacology at the UMASS Medical School at Worcester, MA from May, 2003 to May 2004 on an NIH Fellowship. Back to India, she concentrated her research efforts in the area of Gamete Biology. Presently her research is focused towards understanding the molecular mechanisms underlying sperm motility and developing tools towards selection of good quality sperm for assisted reproductive technologies. She has 33 publications till date in peer reviewed International journals.

Tubulin Reversible Acetylation – Driving the Moves and the Moves Behind the Drive

Objective:

Asthenozoospermia accounts for almost 50% of the cases of male infertility. Our study investigating phosphoproteins differentially expressed in asthenozoosperm has identified the phosphoproteins relevant to sperm motility and the signature molecules likely to be altered in asthenozoospermia. The 66 phosphoproteins differentially expressed included four alpha tubulin isoforms which were differential expressed in individuals with poor sperm motility; an increase in TUBA3E and TUBA4A and decrease in TUBA3C and TUBA8. Isoforms TUBA3C, TUBA4A and TUBA4A are reported to be abundantly present in the testis. Of these, TUBA3C and TUBA4A are known to be acetylable isoforms. Their differential expression in asthenozoosperm prompted us to investigate the role of reversible acetylation of alpha tubulin in sperm motility.

Material and Methods:

Alpha tubulin, acetylated α tubulin, and isoforms TUBA3C, TUBA4A, and TUBA8 were investigated in Percoll separated human sperm and HDAC6 in rat sperm by Western blot analyses, Flow cytometry, Real-time RT-PCR and IIF localization and data statistically analyzed. The observations were analyzed in silico for obtaining further insights. Interaction between HDAC6 and α Tubulin was elucidated by IIF co-localization and Co-immunoprecipitation studies. Sperm HDAC6 activity, motility and status of Ac α -tubulin was investigated in the presence of HDAC inhibitors Trichostatin A, Tubastatin A and Sodium Butyrate.

Results:

The differential expression of these isoforms was validated in the normal- and asthenozoosperm at protein and transcript level. Investigation of acetyl α tubulin expression revealed a reduction in tubulin acetylation in asthenozoosperm. The decrease in TUBA3C and increase in TUBA4A transcripts, both being acetylable isoforms of alpha tubulin, could be elucidated on the basis of transcription factors binding to the promoters of the respective isoforms while the reduction in acetyl alpha tubulin in asthenozoosperm could be explained on the basis of decreased TUBA3C and the association of HDAC6 with TUBA4A. However the presence of HDAC6 on sperm had not been hitherto reported. In this study we have demonstrated for the first time the presence of HDAC6 transcript and protein in testicular- and caudal-sperm of rat and further by co-localization and coimmunoprecipitation studies we showed that HDAC6 interacts with alpha-tubulin and they colocalize in the mid piece and principal piece of sperm flagella. Using HDAC inhibitors we further demonstrated that HDAC6 in sperm is catalytically active and inhibitors of HDAC6 increase acetylation and restrict sperm motility.

Conclusion:

Our data suggests an association between reversible α tubulin acetylation and sperm motility. We show that alpha tubulin acetylation is reduced in sperm of asthenozoospermic individuals. Paradoxically, our experiments in the rat sperm show that inhibition of HDAC6 increase alpha tubulin acetylation but restrict sperm motility. The persistent expression of HDAC6 on the sperm flagella in the presence of HDAC6 inhibitor hints at a possible role for HDAC6 possibly as a microtubule associated protein(MAP). On the basis of our observations with the human and rat sperm and the available literature on microtubule stability in atat1 KO mice, we conclude that HDAC6 may act as a MAP and maintain dynamic instability in sperm flagella, and that dynamic instability may be a pre-requisite for normal sperm motility.

Keywords:

Acetylated Alpha-tubulin, Deacetylase Activity, HDAC6, HDAC Inhibitor, Sperm Motility





International Winner

Biotechnology

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Identifying and Overcoming an Epigenetic Barrier for SCNT Reprogramming

Objective:

Despite successful cloning of many mammalian species, the cloning efficiency is extremely low compared to that of IVF raising the possibility of the existence of epigenetic barrier preventing successful cloning.

Material and Methods:

Using comparative transcriptome analysis comparing transcriptomes of IVF and SCNT embryos and that of donor cells, we identified epigenetic barrier and find a simple way to overcome the barrier.

Results:

We identified that the Suv39h deposited H3K9me3 in somatic cells is an epigenetic barrier preventing zygotic genome activation and thus preventing successful development of the SCNT embryos. We find that by injecting mRNA encoding a H3K9me3 demethylase can overcome this epigenetic barrier to achieve high efficient cloning.

Conclusion:

Suv39h deposited H3K9me3 in somatic cell is an epigenetic barrier for SCNT, which can be overcome by injecting mRNA of Kdm4d into the one-cell embryo to achieve high efficient cloning.

Keywords:

SCNT, Epigenetic Barrier for Cloning, Suv39h, Kdm4d, ZGA



Yi Zhang, PhD

■ USA yzhang@genetics.med.harvard.edu

Dr Zhang is currently an Investigator of the Howard Hughes Medical Institute and a Fred Rosen Chair Professor of the Department of Genetics and Department of Pediatrics of the Harvard Medical School. He is also a senior Investigator of PCMM of the Boston Children's Hospital. The major interest of Dr Zhang is to understand the epigenetic regulation in early development, stem cell reprogramming, differentiation, reward-related learning and memory, differentiation, and how dysregulation of chromatin modifying enzymes contribute to various human diseases. His group contributed to the identification and characterization of several classes of epigenetic enzymes that include histone methyltransferases, containing histone demethylases, histone H2A ubiquitin E3 ligase, and the Tet family of 5-methylcytosine dioxygenases. Dr Zhang was Top 10 author of high-impact papers by ScienceWatch in Genetics and Molecular Biology, and is also one of the most influential scientists in the world in the past 10 years.

International Winner

Reproductive Genetics



Masoud Zamani Esteki, PhD

■ Belgium masoud.zamaniesteki@med. kuleuven.be

Masoud Zamani Esteki computational geneticist working at the Centre for Human Genetics, working University Hospital, Leuven (Belgium). He has recently accomplished his PhD in Molecular and Developmental Genetics at the same university. In preparation of his doctoral research, he first obtained an MSc degree in Bioinformatics followed by an Advanced Master degree in Artificial Intelligence, Engineering and Computer Science (ECS), both from KU Leuven. Prior to that, he acquired a BSc degree in Genetics at the Shahid Chamran University of Ahvaz (Iran). During the course of his PhD research, he developed, applied and translated an ensemble of innovative genome analysis approaches, including the haplarithmisis concept, which uncovers the allelic architecture of entire genomes in DNA derived from many cells down to a single cell.

Concurrent Whole - Genome Haplotyping and Copy-Number Profiling of Single Cells

Objective:

Methods for haplotyping and DNA copy-number typing of single cells are paramount for studying genomic heterogeneity and enabling genetic diagnosis. Before analyzing the DNA of a single cell by microarray or next-generation sequencing, a whole-genome amplification (WGA) process is required, but it substantially distorts the frequency and composition of the cell's alleles. As a consequence, haplotyping methods suffer from error-prone discrete SNP genotypes (AA, AB, BB) and DNA copy-number profiling remains difficult because true DNA copy-number aberrations have to be discriminated from WGA artifacts.

Material and Methods:

Here, we developed a single-cell genome analysis method that reconstructs genome-wide haplotype architectures as well as the copy-number and segregational origin of those haplotypes by employing phased parental genotypes and deciphering WGA-distorted SNP B-allele fractions via a process we coin haplarithmisis.

Results:

Our approach proved accurate on 55 embryos from 12 couples carrying either autosomal dominant, recessive or X-linked Mendelian disorders, or simple or complex translocations. The method allowed diagnosing an embryo for multiple monogenic disorders at once, and, in contrast to current PGD for translocation cases, it enabled distinguishing embryos that inherited normal chromosomes from embryos that inherited a balanced configuration of the rearranged derivative chromosomes.

Conclusion:

We demonstrate that the method can be applied as a generic method for preimplantation genetic diagnosis (PGD) on single cells biopsied from human embryos, enabling diagnosis of disease alleles genome wide as well as numerical and structural chromosomal anomalies. Moreover, meiotic segregation errors can be distinguished from mitotic ones. The method, therefore, facilitates genetic selection of embryos, and broadens the range of classic PGD.

Keywords:

Single-Cell Genomics, Haplotyping, Copy Number Typing, Reproductive Genomics, Embryo Genomics



Anternational Winner

Stem Cell Biology and Technology



DNA Oxidation Towards Totipotency in Mammalian Development

Objective:

Mammalian development begins with a single cell resulted from the fertilization of a sperm and an oocyte. The early embryonic genome undergoes profound epigenetic reprogramming to prepare for development. We try to understand the biological significance and mechanisms of epigenetic reprogramming.

Material and Methods:

We used biochemical assays to detect oxidation product of 5-methylcytosine in DNA and identify the responsible oxidases. The significance of 5mC oxidation was confirmed by mouse gene targeting experiments.

Results:

We find that 5-methylcytosine (5mC), the most abundant type of base modification in DNA, is oxidized to 5-hydroxymethylcytosine (5hmC) as well as 5-carboxymethylcytosine (5caC) in mouse zygotes. In vitro, the Tet family of dioxygenases oxidize 5mC to 5caC under physiologically relevant conditions (e.g. in the presence of 1mM ATP). In zygotes, the Ten-eleven-translocation protein Tet3 is responsible for the genome-wide oxidation of 5mC to 5hmC and 5caC. Deficiency of zygotic Tet3 impedes demethylation at the paternal Oct4 and Nanog genes and delays the reactivation of Oct4 in early embryos. The heterozygous mutant embryos lacking maternal Tet3 suffer increased developmental failures. Importantly, oocytes lacking Tet3 also show impaired reprogramming of injected somatic cell nuclei. In addition, MEFs deficient in all Tet genes were unable to be reprogrammed by Yamanaka factors

Conclusion:

We conclude that Tet-mediated oxidation is important for DNA demethylation and gene activation in the early embryo following natural fertilization, as well as for the reprogramming in somatic cell nuclear transfer and factor-based iPSC generation.

Keywords:

Epigenetic Reprogramming, DNA Demethylation, Enzymatic DNA Oxidation, Cell Pluripotency



Guoliang Xu, PhD

China glxu@sibs.ac.cn

Dr Xu is a principal investigator at the Institute of Biochemistry and Cell Biology (IBCB), Chinese Academy of Sciences, Shanghai. He trained for PhD at the Max Planck Institute for Molecular Genetics, Berlin. He conducted his postdoctoral training at Columbia University, New York. Dr Xu established the DNA metabolism research group in 2001 under the auspices of Max Planck research group at Shanghai IBCB, where his team investigates the importance of DNA methylation (5-methylcytosine) in stem cells and development. In 2011 his team described a role for Tet DNA dioxygenases in oxidation of 5-methylcytosine 5-carboxylcytosine involvement of a DNA glycosylase -TDG in active demethylation, a critical step for the epigenetic reprogramming of early embryos and throughout development.

National Willi

Reproductive Genetics



Maryam Shahhoseini, PhD

■ Iran m.shahhoseini@royaninstitute.org

Maryam Shahhoseini received her BSc degree in Microbiology in 1998 from Department of Biology, University of Tehran. In a straight line, she started her MSc degree in Biochemistry at the Institute of Biochemistry & Biophysics (IBB), University of Tehran, and was graduated in 2001 as the 1st rank student. She continued her education at PhD level in Biochemistry at the same department (IBB) and was graduated 2007 as the 1st rank student. Immediately after dissertation, she joined Royan Institute as an academic Now, she is the Associate Professor of Genetic Department, and the leader of "Epigenetic Sub-group" in Royan Institute.

Since 2007, she has managed several genetic/epigenetic projects in the fields of Reproduction and Stem Cell Biology. Parallel, she has been the invited academic lecturer of Department of Biotechnology, Faculty of Sciences, University of Tehran, teaching Molecular Biotechnology, and Genetics, Molecular Biotechnology, and Gene Expression Regulation courses.

Expression Profile of Macrophage Migration Inhibitory Factor (MIF) Signaling Pathway as a Potentional Biomarker in Pathophysiology of Endometriosis

Objective:

MIF via its receptor, CD74, initiates a signaling cascade that leads to proliferation and survival of cells. Also, MIF binding to CD74 activates p38 signaling pathways that lead to positive effect on the expression of COX-2. The aim of this study was to evaluate the gene expression profile of MIF, CD74 and COX-2 in normal, ectopic and eutopic endometrium during menstrual cycle. The expression level of MIF protein in peripheral blood samples of patients was another variable factor checked in this study.

Material and Methods:

Quantitative real-time polymerase chain reaction (Q-PCR) was performed using cDNA and primers for MIF, CD74 and COX-2. Also, protein level of MIF in blood serum was measured by ELISA assay.

Results:

The mean relative expression of MIF, CD74 and COX-2 genes were significantly higher in ectopic endometrium in compare to eutopic and control endometrium. However, there were significantly variations in mRNA expression of these genes in normal, ectopic and eutopic endometrium during menstrual cycle. Also women with endometriosis had significantly higher circulating levels of MIF protein as compared to normal controls.

Conclusion

Higher expression of MIF, CD74 and COX-2 genes in ectopic endometrium can be considered as a molecular biomarker for endometriosis development and pathophysiology. Variation in the expression of these genes in normal, ectopic and eutopic endometrium during menstrual cycle could play an essential role in reproduction, inflammation and endometrium reconstruction.

Keywords:

MIF, CD74, COX-2, Endometriosis



Mational Winner

Stem Cell Biology and Technology



Bioinspired Substrates Direct the Fate of Stem Cells

Objective:

Bioinspired materials can mimic the stem cell environment and modulate stem cell differentiation and proliferation. In this context, biomaterials can mimic the biological microenvironments (i.e., niches) of stem cells and specifically affect the in vitro differentiation that is necessary for clinical application. In vivo, the appropriate differentiation, proliferation, and maintenance of potency are regulated by either stem cells or their specific niches. In this study, biomimetic micro/nanoenvironments were fabricated by cell-imprinted substrates based on mature human keratinocyte morphological templates and also smart nanoenvironments were obtained by cell-imprinted substrates based on mature and dedifferentiated chondrocytes as templates. Material and Methods:

This substrate was characterized by SEM, AFM, Flouresent and Confocal microscopy. Toxicity of PDMS to stem cells was evaluated using an MTT-assay. The gene expression analysis of differentiated cells, were detected by Real Time PCR, array analysis and computer simulation study.

Results:

The data obtained from atomic force microscopy and field emission scanning electron microscopy revealed that the keratinocyte-cell-imprinted poly (dimethylsiloxane) casting procedure could imitate the surface morphology of the plasma membrane, ranging from the nanoscale to the macroscale, which may provide the required topographical cell fingerprints to induce differentiation. Gene expression levels of the genes analyzed (involucrin, collagen type I, and keratin 10) together with protein expression data showed that human adipose-derived stem cells (ADSCs) seeded on these cell-imprinted substrates were driven to adopt the specific shape and characteristics of keratinocytes. The observed morphology of the ADSCs grown on the keratinocyte casts was noticeably different from that of stem cells cultivated on the stem-cell-imprinted substrates. Since the shape and geometry of the nucleus could potentially alter the gene expression, we used molecular dynamics to probe the effect of the confining geometry on the chain arrangement of simulated chromatin fibers in the nuclei. Additionally, rabbit adipose derived mesenchymal stem cells (ADSCs) seeded on these cell-imprinted substrates were driven to adopt the specific shape (as determined in terms of cell morphology) and molecular characteristics (as determined in terms of gene expression) of the cell types which had been used as template for the cell-imprinting.

Conclusion:

The results obtained suggested that induction of mature cell shapes onto stem cells can influence nucleus deformation of the stem cells followed by regulation of target genes. This might pave the way for a reliable, efficient, and cheap approach of controlling stem cell differentiation toward human cells for wound healing applications.

Keywords:

Stem Cells, Biomimicking, Cell Fate, Differentiation, Smart Substrates



Morteza Mahmoudi, PhD

■ Iran mahmoudi-m@tums.ac.ir

Morteza Mahmoudi received his PhD from Institute for Nanoscience and Nanotechnology at Sharif University Technology with specialization applications the biomedical of superparamagnetic iron nanoparticles. He is Director of NanoBio Interaction Laboratory at University of Medical Sciences. He is the leading expert on the bionanointerfaces and stem cell nano-based engineering. He was visiting Professor at University of Illinois at Urbana Champaign and Stanford School of Medicine. He has received numerous awards for his work on nanobiointeractions and stem cell nanotechnologies including 11th Kharazmi Young Festival Award (2009), Razi Festival Young Researcher National Award (2012), Sixth-Ninth Festival National Awards of Iran Nanotechnology Initiative (2010-2014), Distinguished Researcher of Iran's Ministries (2013), and Stanford University Seed Grants Women's Heart Health Program (2015). He has also introduced new technology in stem cell differentiation using pseudo 3D smart substrates



Last Name, First Name, Degree	Country
Abroun, Saeid, PhD, Post Doc	Iran
Aflatoonian, Reza, MD, PhD	Iran
Aflatoonian, Abbas, MD	Iran
Afsharian, Parvaneh, PhD	Iran
Aghdami, Nasser, MD, PhD	Iran
Ahmadi, Firoozeh, MD	Iran
Ajonuma, Louis Chukwuemeka, MD, PhD	Hong Kong
Akhlaghpoor, Shahram, MD	lran
Aleyasin, Ashraf, MD	Iran
Al-Hasani, Safaa, DVM, PhD	Germany
Alimoghaddam, Kamran, MD	lran
Alini, Mauro, PhD	Switzerland
Alipour, Hiva, DVM, PhD Student	Denmark
Alizadeh Moghadam Masouleh, AliReza, PhD	lran
Almadani, Seyed Navid, MD	lran
Almstrup, Kristian, PhD	Denmark
	(Slovak Republic)
Altmae, Signe, PhD	Spain
Amirchaghmaghi, Elham, MD, PhD	lran
Ao, Asangla, PhD	Canada
Ashrafi, Mahnaz, MD	lran
Awwad, Johnny, MD	Lebanon
Bahadori, Mohammad Hadi, PhD	lran
Baharvand, Hossein, PhD	lran
Bakhtiari, Mitra, PhD	lran
Baron-Van Evercooren, Anne, PhD	France
Bazrgar, Masood, PhD	lran
Beigi Boroujeni, Mandana, PhD	lran
Benagiano, Giuseppe, MD, PhD, FACOG, FRCOG	Switzerland
Braun, Thomas, MD, PhD	Germany
Bulte, Jeff, PhD	United States
Chieffi, Paolo, MD, PhD	ltaly
Dadkhah, Farid, MD	lran
Dazzi, Francesco, MD, PhD	United Kinadom
De Geyter, Christian, MD	Switzerland
de Rooij, Dirk G., PhD	Netherlands

Last Name, First Name, Degree	Country
Ebner, Thomas, PhD	Austria
Ebrahimi, Marzieh, PhD	Iran
Ebrahimi, Bita, PhD	Iran
Eftekhari-Yazdi, Poopak, PhD	Iran
Engin, Gulgun, MD	
Esmaeili Borzabadi, Vahid, MSc	
Evans, John, PhD	
Evers, Johannes, MD, PhD, FRCOG	Nothorlands
Fakhr Taha, Masoumeh, PhD	Iran
Familari, Mary, PhD	Australia
Farrahi Faramarz MD	Iran
Fathi, Rouhollah, PhD	Iran
Fathi, Ali, PhD	Iran
Fleming, Steven, PhD	Australia
Francesco Saverio, Tedesco, MD, PhD	
Franco Junior, Jose G., MD, PhD	Rrazil
Fulka, Jr., Josef, PhD, DSc	
Gee, Adrian, PhD	11 1 10 10 1
Gentile, Luca, PhD	_
Geraedts, Joep, PhD	Netherlands
Ghaderian, Sayyed Mohammad Hossein, Post Doc fellow	MD, PhD, lran
Ghaedi, Kamran , PhD, Doc fellow	Iran
Giojalas, Laura, PhD	Argentina
Gourabi, Hamid, PhD	lran
Hamidieh, Amir Ali, MD	Iran
Hammarberg, Karin, RN, PhD	Australia
Hassani, Seyedeh Nafiseh, PhD	Iran
Heimberg, Harry, PhD	Relaium
Henkel, Ralf, BEd, PhD	South Africa
Hescheler, Jurgen, MD	Germany
Honaramooz, Ali, DVM, PhD	Canada
Hoseini far, Hani, MSc	Iran
Hoseini far, Hani, MSc Hosseini, Morteza, PhD	lran Iran
Hoseini far, Hani, MSc	Iran





Last Name, First Name, Degree	Country
Kabir-Salmani, Maryam, PhD, Post Doc	Iran
Kallen, Bengt, MD, PhD	Sweden
Kamali, Koorosh, MD, MPH, PhD	Iran
Kamali, Mohammad, PhD	Iran
Karimian, Leila, MSc	Iran
Karimzadeh Meybodi, Mohammad Ali, MD	Iran
Katabuchi, Hidetaka, MD, PhD	Japan
Khalili, Mohammad Ali, PhD	Iran
Khang, Gilson, PhD	Korea, Republic of
Kupesic Plavsic, Sanja, MD	United States
Liu, Ji-Long, PhD	United Kinadom
Maccarrone, Mauro, PhD	Italy
Ma'dani, Tahereh, MD	lran
Martino, Gianvito, MD	Italy
Mathur, Premendu, PhD	India
McNatty, Ken, PhD, DSc	New Zealand
Minami, Naojiro, PhD	lanan
Moein. Mohammad Reza. MD	Iran
Mohammadi Roushandeh, Amaneh, PhD	Iran
Mohseni Meybodi, Anahita, PhD	Iran
Moini, Ashraf, MD	Iran
Movaghar, Bahar, PhD	Iran
Mowla, Seyed Javad, PhD	Iran
Mozdarani, Hossein, PhD	Iran
Mukhopadhyay, Asok, PhD	India
Nakao, Toshihiko, DVM, MS, PhD	Japan
Nasr-Esfahani, Mohammad Hossein, PhD	Iran
Nasu, Kaei, MD, PhD	Japan
Ng, Ernest, MD	Hona Kona
Nielsen, Hans Ingolf, PhD, MEd	Denmark
Niknejadi, Maryam, MD	Iran
Nikzad, Hossein, PhD	lran
Nouri, Mohammad, PhD	
Nussler, Andreas, PhD	Germany
Ory, Steven, MD	United States
Ott, Michael, MD Pandit. Abhay, MS. PhD. MPH	Germany
Panina- Bordignon, Paola, PhD	Italy
Petraglia, Felice, MD	Italy
Piemonti, Lorenzo, MD	ltaly
Pourmand Gholamreza MD	Iran
Pourmand, Gholamreza, MD	lran
Puri, Chander P., PhD, FAMS, FNASc	India
namezanzaden, ratemen, MD	II ai i
Rashidi, Batool, MD	Iran
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Last Name, First Name, Degree	Country
Ray, Pierre F., PhD	France
Redi, CarloAlberto, PhD	Italy
Rezazadeh, Mojtaba, PhD	Iran
Ringe, Jochen, PhD	
Rizk, Mohamed Mostafa, MD	Favnt
Rostami, Sirous, MD	Iran
Rousseaux, Sophie, MD, PhD	_
Sabbaghian, Marjan, PhD	Iran
Sadeghi, Mohamad Reza, PhD	Iran
Sadighi Gilani, Mohammad Ali, MD	Iran
Safarinejad, Mohammad Reza, MD	Iran
Salman Yazdi, Reza, DCLS	Iran
Saric, Tomo, MD, PhD	C
Satarian Leila PhD	Iran
Schlegel, Peter, MD	
Shahhosaini Maryam PhD	Iran
Chabrard: Abdalbassain DLD	luo io
Shahzadeh Fazeli, Seyed Abolhassan, MD, PhD	Iran
Chius Marzigh MD	Iran
Cinn Douglas DA	lanan
Caflitia Nikalaaa AAD DLD	C v a a a a
Stewart Flizabeth MD	United States
Strauer, Bodo, MD, FRCP, FACC, FESC	Germany
Sumi Shoichiro MD PhD	lanan
Tahamtani Vacor DhD	Iran
Taheri Panah, Robabeh, MD	Iran
Tardif Steve PhD	United States
Totonchi Mehdi PhD	Iran
Ural. Ali MD	Turkev
Van der Veen, Fulco PhD	Netherlands
Van Lohuizen, Maarten, PhD	
Verlhac, Marie-Helene, PhD	France
Viville, Stéphane, PhD	France
Viville, Stéphane, PhD Vosough Taghi Dizaj, Ahmad, MD	Iran
Wang, Dong-An, PhD	9 9
Yadav, Prem, PhD	India
	Iran
Zamanian, Mahdi, PhD Zamanian, Mohammadreza, MD, PhD	Iran
Zerbini, Gianpaolo, MD, PhD	ltalv
Zhang. Chengcheng. PhD	United States
Zhang , Chengcheng, PhD Zhou , Jiawei, PhD	China
Zolghadri, Jaleh, MD	Iran
Loightan, Jaich, MD	11 (111



Board scientific coi

Last Name, First Name, Degree	Country
A. Greenfeld, Dorothy, *	USA
Abbasi, Mahmood, PhD	Iran
Abbasy, Homayoun, MD	Iran
Aflatoonian, Abbas, MD	Iran
Aflatoonian, Reza, MD, PhD	Iran
Afsharian, Parvaneh, PhD	Iran
Agarwal, Ashok, PhD, HCLD	USA
Aghdami, Nasser, MD, PhD	Iran
Ahmadi, Firoozeh, MD	Iran
Akhlaghpoor, Shahram, MD	Iran
Akhondi, Mehdi, PhD	Iran
Akhoond, Mohamad Reza, PhD	Iran
Alborzi, Saeed, MD	Iran
Aleyasin, Ashraf, MD	Iran
Al-Hasani, Safaa, DVM, PhD	Germany
Al-Inany, Hesham, MD, PhD	Egypt
Almadani, Seyed Navid, MD	Iran
Almstrup, Kristian, PhD	Denmark
Amir Jannati, Naser, MD	Iran
Ansari-Pour, Naser, PhD	Iran
Ao, Asangla, PhD	Canada
Aramesh, Kiarash, MD	Iran
Arefi, Soheila, MD	Iran
Asghari, Fariba, MD	Iran
Ashrafi, Mahnaz, MD	Iran
Azin, Seyed Ali, MD, PhD	Iran
Azin, Seyyed Mohammad, PhD Student	Iran
Bagheri Lankarani, Narges, PhD	Iran
Baharvand, Hossein, PhD	Iran
Bazrgar, Masood, PhD	Iran
Behnam Manesh, Shima, MA	Iran
Benagiano, Giuseppe, MD, PhD, FACOG,FRCOG	Switzerland
Boiani, Michele, PhD	Germany
Brandacher, Gerald, MD	USA
Brivanlou, Ali H., PhD	USA
Caplan, Arthur, PhD	USA
Chian, Ri-Cheng, PhD	Canada
Dadkhah, Farid, MD	Iran
Dazzi, Francesco, MD, PhD	Uk
Ebner, Thomas, PhD	Austria

Last Name, First Name, Degree	Country
Ebrahimi, Marzieh, PhD	Irar
Ebrahimi, Bita, PhD	Irar
Eftekhari-Yazdi, Poopak, PhD	Irar
Eimani, Hussein, PhD	Irar
Englert, Yvon, MD*	Belgium
Eshrati, Babak, PhD	Irar
Esmaeili Borzabadi, Vahid, MSc	Irar
Farrahi, Faramarz, MD	Irar
Farzadi, La'ya, MD	Irar
Ferrara, James L.M., MD, DSc	USA
Frith, Lucy, BA (Hons) MPhil, PhD	UŁ
Geraedts, Joep, PhD	Netherlands
Ghaedi, Kamran, PhD, Doc fellow	Irar
Ghafari, Firoozeh, MD	Irar
Ghorbani, Behzad, MD	Irar
Giojalas, Laura, PhD	Argentina
Golestanha, Seyyed Ali, MD	lrar
Gourabi, Hamid, PhD	lrar
Heimberg, Harry, PhD	Belgium
Hens, Kristien, PhD	Netherlands
Hillier, Stephen, MD	Uł
Honaramooz, Ali, DVM, PhD	Canada
Hoseini far, Hani, MSc	lrar
Hosseini Salekdeh, Seyed Ghasem, PhD	lrar
Hosseini, Ahmad, PhD	lrar
Hosseini, Jalil, MD	lrar
Hosseini, Roya, MD	lrar
Hosseini, Morteza, PhD	lrar
Jalali, Mohsen, PhD	lrar
Johnson, Peter, PhD	Uk
Kalantar, Seyed Mehdi, PhD	lrar
Kalantary, Mojgan, MD	lrar
Kamali, Koorosh, MD, MPH, PhD	Irar
Kamali, Mohammad, PhD	lrar
Karimian, Leila, MSc	lrar
Karimzadeh Meybodi, Mohammad Ali, PhD	lrar
Kazemeyni, Seyed Mohammad, MD	Irar
Khalili, Gholamreza, MD, PhD	lrar
Khalili, Mohammad Ali, PhD	lrar
Kiani Cahar DhD	Irar





ast Name, First Name, Degree	Country

Last Name, First Name, Degree	Country
Kupesic Plavsic, Sanja, MD	USA
Kurpisz, Maciej, MD, PhD	Poland
Larijani Tahoroh MD	Iran
Larijani Paghor MD	Iran
Linchultz Larry MD	ΙΙςΔ
Ma'dani Taharah MD	Iran
Malakafzali Haccoin DhD	Iran
Marghati Savad Taha PhD	Iran
Milanifar Alireza MD PhD Student	Iran
Maghaddam Matin Maryam PhD	Iran
Maghaddacali Paza DhD	Iran
Mohammad Kazam PhD	Iran
Mahsani Maybadi Anahita DhD	Iran
Moini, Ashraf, MD	Iran
Momtaz Mohamed MD	Favnt
Moncage TK PhD	South Africa
Moreh Ian S MD	ΙΙςΔ
Movaghar, Bahar, PhD	Iran
Movahedin, Mansoureh, PhD	Iran
Movassagh, Hooman, LLB, LLM, PhD Nakatsuji, Norio, PhD	UJA Janan
Namazi Hamidreza MD PhD Student	Iran
Nasr-Esfahani, Mohammad Hossein, PhD	
Nazari Tavakkali Capid DED	Iran
Nematollahi-mahani, Seyed Noureddin, PhD	Iran
NIT I I NA NA	1
Nouri Mohammad PhD	
Nouri, Mohammad, PhD Oback, Björn, PhD	New Zealand
Oghabian, Mohamad Ali, PhD	Iran
Omani Samani Raza MD PhD Student	Iran
Parsanezhad, Mohammad Ebrahim, MD	
Parsapour, Alireza, MD, PhD Student	Iran
Dobay Alico DED	Australia
Pannings Guido DDD	Polaium
Petraglia, Felice, MD	
Polan, Mary Lake, MD, PhD, MPH	ltaly
Poureisa, Masoud, MD	Iran
Raffaella Fabbri PbD	Italy
Rajabi, Sareh, PhD Ramezanzadeh, Fatemeh, MD Rashidi, Batool, MD Ray, Pierre E, PhD	Iran
Ramezanzadeh, Fatemeh, MD	Iran
Rashidi, Batool, MD	Iran
Rashidi, Batool, MD Ray, Pierre F, PhD	France
Padi Carlo Alborto DbD	l+alv
Ranning Sigard PhD	Natharlands
	Iran
Rezazadeh, Moitaba PhD	Iran
Richters, Juliet, PhD	ltalv
Richters, Juliet, PhD Sabbaghian, Marjan, PhD Sabeti, Shokofeh, MD	Iran
Sabeti, Shokofeh, MD	Iran
Sadaghi Mohamad Reza PhD	Iran
Jacqii, Monamad Neza, Filb	

Last Name, First Name, Degree	Country
Sadighi Gilani, Mohammad Ali, MD	lran
Saeidi, Hojjatollah, PhD	lrar
Safdarian, Leila, MD	Iran
Salamati, Masoumeh, MD	Iran
Salehnia, Mojdeh, PhD	Iran
Salehpour, Saghar, MD	Iran
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Satarian, Leila, PhD	
SepidarKish, Mahdi, PhD	Iran
Shahhoseini, Maryam, PhD	Iran
Shahvardi Ahdolhossain PhD	lran
Shahzadeh Fazeli, Seyed Abolhassan, MD, PhD	Iran
Shamsi Gooshki Ehsan MD BbD	Iran
Shamei nour Maneur DhD Student	Iran
Shariatinacah Sadach DhD Student	
Shariatinasab, Sadegh, PhD Student Shenfield, Françoise,*	
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Spaudiora, Corrado, PhD	Ildiy
Starring Christon, MD	GCIIIIdiiy
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Taheri Panah, Robabeh, MD	Irar
Tahmasebpour, Ahmad Reza, MD	Irar
Tarzamni, Mohammad Kazem, MD	Iran
Tehraninejad, Ensieh, MD	Iran
Thomson, Jeremy, PhD	Australia
Thornhill, Alan, PhD	Uk
Tian, Xiuchun Cindy, PhD	1.10.4
Totonchi Mehdi PhD	Iran
Vahidi, Serajoddin, MD	Irar
Vahidi, Serajoddin, MD van der Veen, Fulco, PhD Vermeesch, Joris, PhD	Netherlands
Vermeesch, Joris, PhD	Belgium
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Voet, Thierry, PhD	Belaium
Vosough Taghi Dizai. Ahmad. MD	Irar
Voet, Thierry, PhD Vosough Taghi Dizaj, Ahmad, MD Weichert, Alexander, MD	Germany
Wert, Guido de PhD	Netherland
Wert, Guido de, PhD Yazdani, Kamran, MD	Iran
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Zaneui Alididki, Fdizdileli, MU	ı
Zahedi Anaraki, Farzaneh, MD Zamanian, Mohammadreza, MD, PhD Zolghadri, Jaleh, MD	ırar
Zolghadri, Jaleh, MD	Iran



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Last Name, First Name, Degree

Jafarpour, Farnoush, PhD

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Alizadeh, Seyyed Kamal, BSc

Arabipoor, Arezoo, MSc

Behnam-Manesh, Shima, MA

Daliri, Leila, MSc

Ebrahimi, Bita, PhD

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Ezabadi, Zahra, MSc

Gourabi, Hamid, PhD

Hajihasan, Fatemeh, MSc

Hosseinifar, Hani, MSc

Last Name, First Name, Degree

Lotfipanah, Mahdi, BA
Mirshekar, Zeynab, BSc
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Shahverdi, Abdolhossein, PhD
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Zarrabi, Morteza, MD



Kazemi Prize



About Kazemi Prize

Dr Saeid Kazemi Ashtiani was born in March 1961 in Tehran. Upon completion of his high school at the age of 18, he was admitted to Iran Medical University to pursue his studies in the field of Physiotherapy. He graduated in 1991 and subsequently in 1993 he started his postgraduate education in the field of Anatomy (Embryology branch) in Tarbiat Modaress University. He received his Doctorate Degree with Distinction In 1998.

Dr Kazemi established Royan Research Institute in 1991. This institute renders advanced medical services to infertile couples. The center is also one of the most important and active research centers in the Middle East.

Dr Kazemi and his colleagues at Royan Infertility Research Center could achieve a tremendous success in 2003 by establishing human embryonic stem cell line. This great scientific achievement has earned a high position for Iran among the other top 10 countries having access to this advanced technology at that time. He was not only a scientist who led a lot of principle research projects in the field of stem cell and cloning but a great manager as well. He was the head of ACECR, Iran Medical University branch, head of Royan Research Institute, guest instructor and lecturer of many Iranian medical universities, manager and chief of quarterly scientific and research journal of Yakhteh, head of ethical research committee in Royan Institute, and an active member of Iranian society for reproductive biomedicine as well as Iran Anatomical Science

Society. Dr Saeid Kazemi also presided Royan International Award, which was held six times from 2000-2005. His short fruitful life was ended in 2006 when he died of a sudden heart attack.

To respect his efforts and revive his memories amongst national and international scientists as well as nonscientists, Iran supreme leader, Ayatollah Khamenei recommended establishing a yearly prize in biology entitled "Kazemi Prize" which will be awarded to a scientist who made an extraordinary progress in the biological sciences. Kazemi Research Award is for appreciation of extreme effort of the scientist who dedicates his/her life to make progress in human life and relief people's pain.

A nomination committee consisting of prominent national and international scientists is the working body that evaluates the nominees and presents its recommendations to the scientific board of the institute. The scientific board is responsible for the final selection of the prize laureates. In 2010 the first Kazemi Prize was awarded to Prof Rudolf Jaenisch one of the most innovative and creative scientists in the field of developmental biology, gene regulation, stem cell biology and stem cell-mediated therapies. In 2011 the second Kazemi Prize was awarded to Prof Hans Robert Schöler a world-renowned researcher who has made significant contributions to the field of stem cell biology over the past 35 years. And this year the prize will be awarded to Prof Robert S. Langer.

Professor Robert S. Langer is the David H. Koch Institute Professor. Dr Langer has written over 1,300 articles. He also has over 1,080 patents worldwide. Dr Langer's patents have been licensed or sublicensed to over 300 pharmaceutical, chemical, biotechnology and medical device companies. He is the most cited engineer in history (h-index 211).

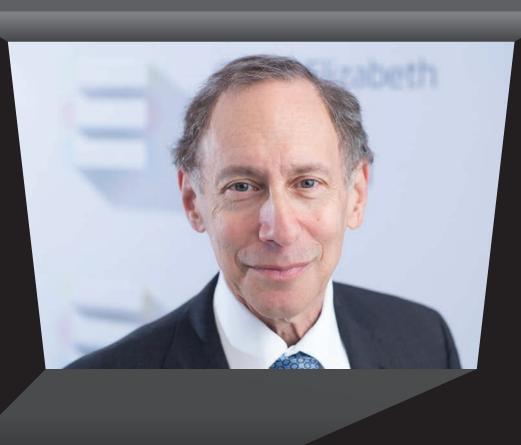
He served as a member of the United States Food and Drug Administration's SCIENCE Board, the FDA's highest advisory board, from 1995 -- 2002 and as its Chairman from 1999-2002.

Dr Langer has received over 220 major awards. He is one of 4 living individuals to have received both the United States National Medal of Science (2006) and the United States National Medal of Technology and Innovation (2011). He also received the 2002 Charles Stark Draper Prize, considered the equivalent of the Nobel Prize for engineers, the 2008 Millennium Prize, the world's largest technology prize, the 2012 Priestley Medal, the highest award of the American Chemical Society, the 2013 Wolf Prize in Chemistry, the 2014 Breakthrough Prize in Life Sciences and the 2014 Kyoto Prize. He is also the only engineer to receive the Gairdner Foundation International Award; 82 recipients of this award have subsequently received a Nobel Prize. In 2015, Dr Langer received the Queen Elizabeth Prize for Engineering. Among numerous other awards Langer has received are the Dickson Prize for Science (2002), Heinz Award for Technology, Economy and Employment (2003), the John Fritz Award (2003) (given previously to inventors such as Thomas Edison and Orville Wright), the General Motors Kettering Prize for Cancer Research (2004), the Dan David Prize in Materials Science (2005), the Albany Medical Center Prize in Medicine and Biomedical Research (2005), the largest prize in the US for medical research, induction into the National Inventors Hall of Fame (2006), the Max Planck Research Award (2008), the Prince of Asturias Award for Technical and Scientific Research (2008) and the Terumo International Prize (2012). In 1998, he received the Lemelson-MIT prize, the world's largest prize for invention for being "one of history's most prolific inventors in medicine." In 1989 Dr Langer was elected to the Institute of Medicine of the National Academy of Sciences, and in 1992 he was elected to both the National Academy of Engineering and to the National Academy of Sciences, and in 2012 he was elected to the National Academy of Inventors.

Forbes Magazine (1999) and Bio World (1990) have named Dr Langer as one of the 25 most important individuals in biotechnology in the world. Discover Magazine (2002) named him as one of the 20 most important people in this area. Forbes Magazine (2002) selected Dr Langer as one of the 15 innovators worldwide who will reinvent our future. Parade Magazine (2004) selected Dr Langer as one of 6 "Heroes whose research may save your life." Dr Langer as received honorary doctorates from many universities, for example: Harvard University, the Mt. Sinai School of Medicine, Yale University, University of Western Ontario (Canada), Boston University and Hanyang University (South Korea). He received his Bachelor's Degree from Cornell University in 1970 and his Sc.D. from the Massachusetts Institute of Technology in 1974, both in Chemical Engineering.

Kazemi Prize **2015**

Professor Robert S. Langer





Royan Institute www.royaninstitute.org



Royan Institute

Royan Institute is a world-renowned center committed to multidisciplinary, campus-wide, integration and collaboration of scientific, academic, and medical personnel for understanding male/ female infertility, embryo development, stem cell biology, and biotechnology. Royan Institute provides comprehensive services for the treatment of infertility, regenerative medicine/ cell therapy and production of recombinant proteins. Royan Institute was established in 1991 by the late Dr Saeid Kazemi Ashtiani (May he rest in peace) in Tehran, Iran. The center supports innovation, excellence and the highest ethical standards focusing on increasing the success rate of infertility treatment alongside embryo health. Furthermore, this center supports the placement of stem cell research findings into operation in cell therapy and disease treatment with the purpose of increasing the level of health.

Mission

The mission of Royan Institute, which is aligned with the country's comprehensive scientific roadmap and the Iranian Academic Center for Education, Culture and Research (ACECR) development plan, can be categorized in the following aspects:

- Research and development of science and technology in the fields of reproductive biomedicine, stem cells and biotechnology
- Education and promotion of scientific findings at national and international levels
- Commercialization of research findings to offer services and biological products for the purpose of resolving the country's specialized needs
- Treatment of infertile patients and difficult-to-treat diseases by the efficient use of research findings

Vision

Royan Institute is a center of excellence in research and technology at an international level, a pioneer in development of science, technology and innovation of biological sciences, and an internationally renowned authority on stem cells science, reproduction, biotechnology, and regenerative medicine alongside its effective role in improving the society's health.



Royan Consists of Three Research Institutes and a Core Facility

- 1. Royan Institute for Reproductive Biomedicine (RI-RB)
- 2. Royan Institute for Stem Cell Biology and Technology (RI-SCBT)
- 3. Royan Institute for Biotechnology (RI-B)
- 4. Laboratory Animal Core Facility

Royan Institute for Reproductive Biomedicine (RI-RB)

Royan Institute for Reproductive Biomedicine, founded in 1991, consists of six departments and one infertility clinic actively working on different aspects of infertility and the development of new methods for infertility treatment.

Its vision is to improve the population's health through infertility treatments and giving infertile families the hope of having children.

In this regard, RI-RB's mission is to research on different aspects of infertility and its treatment in order to increase the success rate alongside improving embryo health.

RI-RB Departments:

- Endocrinology and Female Infertility
- Andrology
- Embryology
- Reproductive Genetics
- Epidemiology and Reproductive Health
- Reproductive Imaging
- Infertility Clinic

Royan Institute for Stem Cell Biology and Technology (RI-SCBT)

Royan Institute for Stem Cell Biology and Technology (RISCBT) was established in 2002 to promote research on general stem cell biology in Iran. Since early 2010, it has continued its activities in:

Stem Cells and Developmental Biology 16 Research Programs

10 Core Facilities

Regenerative Medicine 6 Research Programs

5 Core Facilities

RI-SCBT's vision is to efficiently put stem cell research findings into operation in disease treatment with the aim of improving health. RI-SCBT's mission is to generate insights into the biology of stem cells through basic research and to provide the foundation needed for novel therapies from regenerative medicine.

Royan Institute for Biotechnology (RI-B)

Royan Institute for Biotechnology was initially established in 2004 as the first research branch of Royan Institute. It is located in Isfahan Province, which is famous for its architecture, rivers and handcrafts, and is known as the city of "blue tiles"

Royan Institute for Biotechnology was established with the purpose of advancing research in reproduction, development, cell and molecular biology, in addition to the fields of bioengineering and reproductive technology. In this regard, this Institute has focused on somatic cell nuclear technology (SCNT), interspecies-SCNT, transgenesis, the establishment of novel sperm selection methods for assisted reproductive technology, cell differentiation, production of recombinant proteins and the cell biology of peroxisomes. The endeavors of Royan Institute for Biotechnology have made us the pioneer of animal cloning in Iran and the Middle East. Therefore, this Institute is well known for its cloned animals, Royana and Hanna, the first cloned sheep and goat in Iran, and Bovana; the first calf born with IVF. Areas of interest at this Institute are: gene reprogramming during SCNT, transgenesis, sperm cell biology, the role of sub-cellular organelles in differentiation and recombinant protein technology. In addition, the Institute is providing a comprehensive and coordinated "bench to production" approach in recombinant protein technology, animal farming and the establishment of methods to increase the efficiency of assisted reproductive techniques.

RI-B Groups:

- Cellular Biotechnology- Genetic Laboratory
- Cellular Biotechnology- Stem Cell Laboratory
- Molecular Biotechnology- Recombinant Protein Laboratory
- Reproductive Biotechnology Andrology Laboratory
- Reproductive Biotechnology- Embryology Laboratory

The institute's vision is to attain new heights in biotechnology research, shaping biotechnology into a premier precision tool of the future for creation of wealth, ensuring social justice and efficiently bridging science with daily life.

Overview of the Institute

- The first IVF child born in Tehran (1993)
- The first ICSI child born in Tehran (1995)
- Iran's second success in open testicular biopsy to treat severe male infertility (1996)
- The first frozen embryo child born in Iran (1996)
- The first ICSI birth by frozen sperm of a gonadectomized man in Iran (1999)
- The first human embryonic stem cell line established in Iran and the region (2003)
- The first PGD child born in Iran (2004)
- First time use of adult stem cells in the treatment of MI during CABG in Iran (2004)
- Production of insulin producing cells from human embryonic stem cells (2004)
- Culture of human limbal stem cells on chorionic membrane (2004)
- Establishment of the first Private Cord Blood Bank in Iran (2005)
- The first IVM-IVF sheep born in Iran (2006)
- The first cloned sheep born in Iran (2006)
- Establishment of mouse and human induced pluripotent stem cells (iPS) (2008)
- The first cloned goat born in Iran (2009)
- A new method for treatment of Vitiligo by cell transplantation (2009)
- The first transgenic goats born in Iran (2010)
- The first calves born from vitrified in vitro developed embryos in Iran (2011)
- Establishment of cell therapy pre-hospital (2011)
- Establishment of Stem Cell Bank (2011)
- The first healthy child birth after Molecular PGD for beta-thalassemia in Iran (2012)
- Birth of eight cloned goats through the simplified method of SCNT in Iran (2013)
- Birth of the first cloned wild ram as an endangered species in Iran (2015)



Royan Institute for Reproductive Biomedicine (RI-RB)

CENDOCRING Endocrinology and Female Infertility Department of RI-RB



This department was established in 1995, and began to research on new strategies and advanced methods for the diagnosis and treatment of female infertility and recurrent abortion with the intent of increasing implantation rates.

Goals

- Evaluation and treatment of infertile couples
- New guidelines for improving IVF outcomes
- Achieving new strategies for diagnosing infertility causes
- Ovulation induction and COH
- Improving methods for oocyte and embryo culture
- Endometrial preparation
- The promotion of prenatal care





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Andrology Department of RI-RB

History and Introduction

This department was established in 1995 and started to research on male infertility factors. The first step in infertility management is to evaluate the couple. Male factor infertility accounts for approximately 50% of all infertility cases. Thus in order to study male factor infertility it is necessary to use appropriate diagnostic and therapeutic techniques. The intent of this research department is to develop new diagnostic methods and treatment for male factor infertility.

Goals

- Determining the etiology of spermatogenesis, sperm function and ejaculation disorders
- Determining the etiology of azoospermic, genetic, and maturation disorders
- Determining the etiology of dry and retrograde ejaculation

Main Activities

- Improving diagnostic and therapeutic methods
- Determining the etiology of spermatogenesis, as well as functional and ejaculation disorders



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Contract State of Partners of RI-RB

History and Introduction

The Department of Embryology, founded in 1995, is a part of Royan Institute's Reproductive Biomedicine. During the preceding decade, a fundamental description of human and animal experimental studies has emerged in the field of embryology.

The main focuses of this Department are:

- Increasing the quality of gametes and embryos
- Studying molecular aspects of gamete maturation and embryo development
- Performing embryo co-culture with various types of somatic cells
- · Studying molecular aspects of gamete and embryo freezing
- In vitro maturation of animal and human gametes
- Evaluating molecular and cellular events of embryo implantation
- Three-dimensional culture of cells to design an endometrial biomodel
- Three-dimensional culture of follicles in order to acquire good quality oocytes
- Performing nuclear transfers
- Performing animal cloning and transgenesis
- Finding the best method for preserving gametes, ovarian, and testicular tissues

Goals

- Increasing the number of high quality human embryos
- Producing transgenic animals with selected genes
- Establishing in vitro human follicle culture following ovarian tissue cryopreservation



The mission of the RI-RB Embryology Department is the performance of multiple research regarding different aspects of fertility preservation and different treatments of infertility in order to improve embryo health and increase the pregnancy success rate.

Its aim is to make the wish of having children for infertile couples come true, and to give a promising future to them.



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Reproductive Genetic Department of RI-RB

History and Introduction

Department of Genetics was established in 2001. Some routine activities of this department include: genetic counseling, lymphocyte karyotyping, preimplantation genetic diagnosis (PGD), as well as molecular diagnostic tests which involve the diagnosis of Y chromosomal micro deletions and certain mutations in candidate genes that may be related to the causes of abortions or failed ART.

The major research interests in this department are genetic causes of male and female infertility, recurrent spontaneous abortion (RSA), genetic factors leading to azoospermia, mutations leading to congenital agenesis of the vas deferens, preimplantation genetic diagnosis, pharmacogenetics plus epigenetic and gene expression profiles of early embryogenesis.

The production of recombinant proteins by genetic manipulation in different host cells in addition to the joint production of transgenic animals in a mutual project with Embryology Group is another main activity of this department. Activities carried out in collaboration with Royan Institute for Stem Cell Research are karyotyping of stem cell lines following various manipulations, epigenetic and genetic studies of stem cells and iPS cells, in addition to other common research interests.

Goals

- To improve implantation rates along with health of embryos by preimplantation genetic screening and diagnosis
- To assist physicians with prescribing medicine for controlled ovarian stimulation via pharmacogenetics
- Genetic follow up of newborns conceived by ART
- Evaluation of candidate genes related to recurrent abortion in the Iranian population
- Epigenetic studies of oocytes, sperm and embryos



The mission of the Genetic Department is basic research on genetic and epigenetic factors that may influence fertility, embryo development, and implantation, bringing these research results to the clinical setting with the purpose of improving the health of patients and newborns, as well as the production of pharmaceutical proteins through transgenic animals.

The vision of this department is to perfect diagnosis and treatment of infertility based on reproductive genetic knowledge, which will lead to healthy newborns in a short period of time.



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CEPidemiology and Reproductive Health Department of RI-RB

History and Introduction

The Epidemiology and Reproductive Health Department was established in 1999 with the aim of doing extensive researches on epidemiological aspects of infertility and reproduction in addition to reproductive and sexual health. This department is responsible to check all research proposals in Royan three research institutes and gives both methodological and statistical consultation.

This department undertakes multicentre research between Iran and other countries in the following areas:

- Frequency, incidence and influencing factors for all subfertility and infertility types
- Environmental and occupational factors affecting fertility and reproduction
- Psychosocial issues affecting infertile couples, their treatment and coping mechanisms
- Experiences, quality of life, marital and sexual satisfaction of infertile couples, even after
- IVF failure
- Ethical issues, legislation and guidelines in assisted reproduction
- Statistical models and methods for research in reproduction, genetics and the cellular and molecular fields
- Animal ethics





The mission of this department is the promotion of reproductive health in Iran.

Reproductive health is an important aspect of general health and involves people of all ages within the society, from an embryo to the elderly. Focusing on sexual and reproductive health guarantees the future health of society by ensuring healthy children and healthy adults. Finally, its job involves research into all reproduction related areas including social, medical, psychological and ethical issues, and therefore its vision is to ensure the health of the society.

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○ Reproductive Imaging Department of RI-RB

History and Introduction

Reproductive Imaging Department was established in 2008 to focus on infertility assessment as well as evaluation of pregnancies in both clinic and research fields.

During the last two decades, dynamic advances have been made in the evaluation and treatment of infertility. Imaging technique has been a significant breakthrough in the diagnosis and management of infertility. A broad range of imaging techniques, from the old and proven - such as hysterosalpingography- to the latest and cutting edge - such as three-dimensional hysonosterography, has been employed.

After applying these advanced techniques in this department, we are able to upgrade the management of infertility and obstetric care, and thereby, positively provide better services for infertile couples.

The main goal of imaging department is to provide comprehensive evaluation of infertility using the latest knowledge and innovative research in order to provide the highest quality of infertility management and to monitor emergency obstetric care.

The mission of this research department is to expand clinical and fundamental research in reproductive imaging in order to provide modern strategies and improve clinical services for infertile couples.



The main aim of these projects classified as:

- Diagnostic accuracy investigation of imaging modalities (hysterosalpingography, hysterosonography and three-dimensional ultrasound)
- Role of imaging in (male & female) infertility management
- Fetal screening
- Measurement standards/ultrasound measurement charts appropriate for Iranian fetuses

The vision of this department is performing national and international multicentral researches and having collaboration with universities and other infertility centers to provide educational courses in diagnostic ultrasound including transvaginal, color

Doppler, power Doppler, 3D/4D imaging and radiology for radiologist, gynecologist and fellowships.

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Royan Institute for Stem Cell Biology and Technology



Introduction

Royan Institute for Stem Cell Biology and Technology (RI-SCBT), formerly known as the

Department of Stem Cells was first established in 2002 to promote research on general stem cell biology in Iran. Thereafter, Department of Stem Cells expanded to sixteen main research groups that conduct studies on stem cells and developmental biology and molecular systems' biology. Moreover Department of Regenerative Medicine consists of five main research groups which conduct research focused on translational and clinical studies using cell therapy. Throughout, the vision of RI-SCBT has been to make stem cell research findings applicable in disease treatment to improve public health.

Therefore, today, RI-SCBT is providing a comprehensive and coordinated "bench to bedside" approach to regenerative medicine, as well as a greater understanding of fundamental biology of stem cells, developmental biology, development of translational research of stem cell therapeutics and administration of new cell-therapy approaches that can restore tissue function to patients.





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Royan Institute for Biotechnology (



History and Introduction

In 1983, the late Dr Kazemi Ashtiani, the founder of Royan Institute, along with Dr Nasr-Esfahani established Royan Institute for Biotechnology, as the third branch of Royan Research Institutes. At present, this branch homes around 100 researcher and students working in 5 departments to expand the science over their areas. The intensive seminar schedule in each department has encouraged interactivity and opportunity for scientific discussion between students and the scientists to facilitate the progress of science in their filed. Therefore, in 2010 through this interactive science, this department has achieved a number of important results, including establishment of zona free somatic cell nuclear transfer (SCNT) in goat, evaluation of epigenetic modifier on outcome of SCNT and vitrified embryos, introducing novel approach for selection of intact sperm for ICSI based on sperm functional characteristics, understanding the role of embryonic structure in neurogenesis, assessment of the role of PEP (a peroxisomal protein) and PPAR γ in neurogenesis and finally role of biotechnology in production of biological products.



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Laboratory Animal Core Facility



Introduction

The Laboratory Animal Science Core Facility of Royan Institute plays a national role in education of scholars performing ground researches on experimental animals, by organizing proficient gadget in all categories within the animal research fields. Each center has three major activities:

- Maintenance and breeding the animals
- Creating animal models with surgical manipulations or chemical interactions
- Research and develop animal modeling

Scientists of this service unit facility who are responsible for the design of animal experiments have to be graduated in Veterinary Medicine or one of biomedical science fields and must have taken a course on laboratory animal science which concentrates on humane and gentle handling of animals. They also should be aware of knowledge of alternative routes and ethical aspects of animal experimentation.

Modern laboratory animal science builds on the three Rs of Russell & Burch:

- Replacement: Replace animal experiments with alternatives whenever possible.
- Reduction: Reduce the number of experiments and number of animals in each experiment to an absolute minimum.
- Refinement: Refine experiments so that the animals undergo a minimum of discomfort.

The primary aim of the Laboratory Animal Facility is to ensure that the three Rs are followed in practice.

Goals

- Providing quality care for all animals used at Royan Institute
- Assisting researchers in their mission of quality research with respect to humane use of laboratory animals
- · Providing researchers with a relevant education to enable them achieve scientific eminences in selected areas
- Producing, supporting and maintaining laboratory animals required for research
- Managing the animal care and having commitment to them
- Managing a preventive medicine program for disease control
- Advising research departments on all aspects of experimental use of animals, including experimental design, surgical, pre and post-operative care, oocyte and embryo harvesting, and experimental animal modeling establishment



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Prof. Andreas Keller Chair for Clinical Bioinformatics Saarland University, University Hospital, Saarbrücken, Germany



Prof. Graham Parker Assistant Professor (Research), Dept. of Pediatrics, WSU School of Medicine, USA



Prof. Shahragim Tajbakhsh Head of the Department of Develop-mental & Stem Cell Biology, Vice-Direc-tor of the CNRS Unit URA2578 and Director of a «Laboratory of Excellence» Consortium, REVIVE, Pasteur Institue,



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Prof. Teruko Taketo Professor of Department of Surgery, McGill University, Urology Research Laboratory, Royal Victoria Hospital, West Montreal, Canada



Prof. Paul Mozdziak Professor of Cornell University College of Agriculture and Life Sciences, Ithaca, USA



Associate professor of Department of Genetics, Medicine and Development. University of Geneva, Medical School, Switzerland



Dr. Kenneth McElreaveyDirector of Human Developmental Genetics, Director, Of Reproduction of Fertility, and Populations
Department of Developmental Biology Pasetur institute, France



Prof. Richard Andersen
Elsie Inglis Professor of Clinical Reproductive Science
Head of Section, Obstetrics and Gynecology, University of Edinburgh, UK



Kazem NouriDirector of IVF-Unit, Co-director of Fertility preservation program and fertility surgery unit, Austria



Prof. Stuart S. Howards University of Virginia Charlottesville VA, USA Wake Forest Medical Center Winston Salem NC, USA



Prof. Hooman Sadri Ardekani Director of Male Fertility Research Program, Wake Forrest Institute for Regenerative Medicine (WFIRM) and Department of Orology Wake Forest School of Medicine, Winston-Salem,USA



Don Leigh Principal Scientist, Department of Molecular Genetics, Genea, Sydney, Australia



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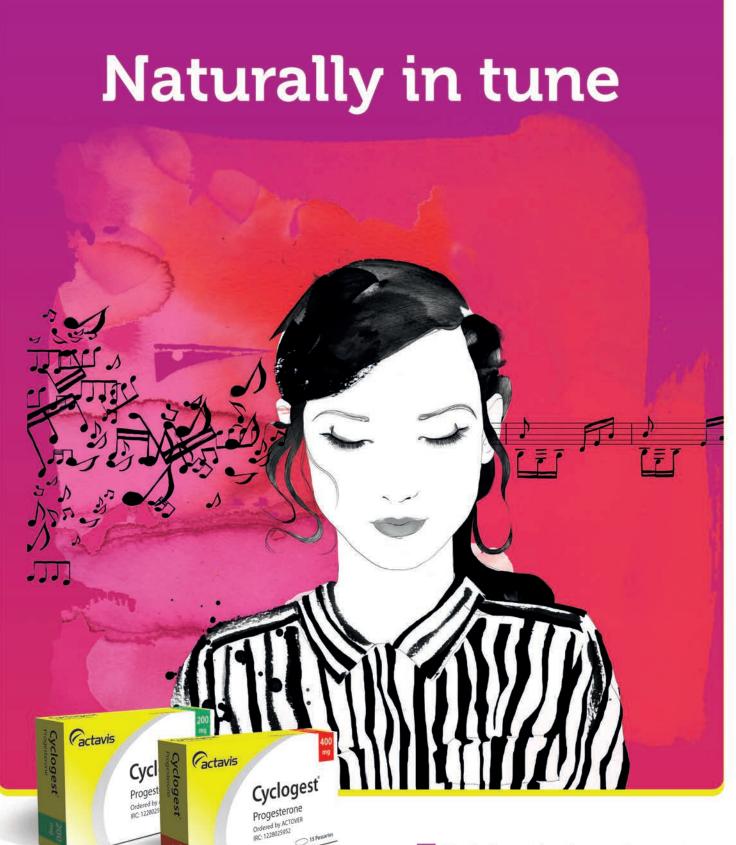




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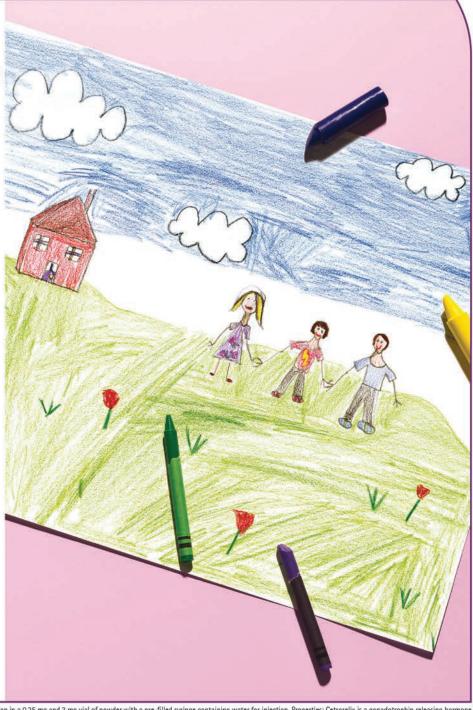
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Ref.: 1- Bodri et al, Fertility and Sterility Vol. 95, No. 1. January 2011. 2- Al-Inany et al, Vol 14. No 5.2007 640-649 Reproductive BioMedicine Online. 3- Al-Inany et al, Human Reproduction Update, Vol.17, No. 4 p. 435, 2011. 4- Al-Inany et al, The Cochrane Library 2011.





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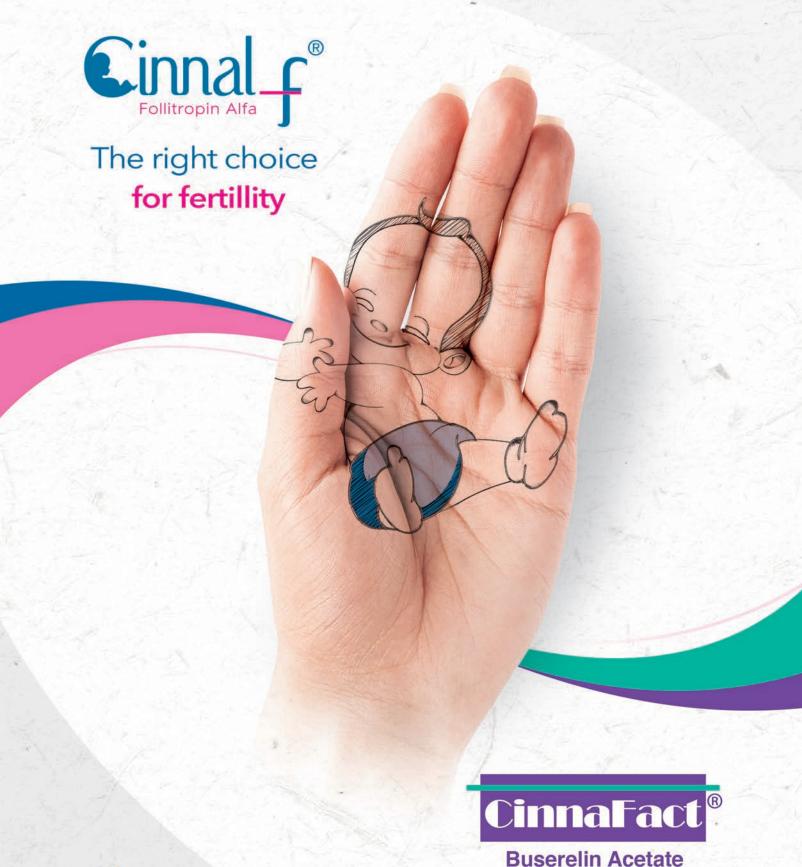


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