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Can a dopamine agonist prevent severe ovarian hyperstimulation syndrome (OHSS) in women undergoing intracytoplasmic sperm injection (ICSI) treatment cycles?

Submission date
21/05/2011

Recruitment status
No longer recruiting

Registration date
19/07/2011

Overall study status
Completed

Last Edited
30/06/2017

Condition category
Pregnancy and Childbirth

[_] Prospectively registered
[_] Protocol
[_] Statistical analysis plan
[X] Results
[_] Individual participant data

Plain English summary of protocol

Background and study aims
Ovarian hyperstimulation syndrome (OHSS) is a complication that can affect women taking injectable hormone medications to stimulate the development of eggs in the ovaries (controlled ovarian hyperstimulation). This can occur in women undergoing in vitro fertilization (IVF), which is a process by which egg cells are fertilized by sperm outside the body. Most cases of OHSS are mild, but a small proportion is severe. The symptoms include swollen and painful ovaries, rapid weight gain, abdominal pain, vomiting and shortness of breath. The aim of this study is to find out whether the drug cabergoline can prevent severe OHSS in women undergoing IVF who are at a high risk of developing OHSS.

Who can participate?
Women undergoing IVF who are at a high risk of developing OHSS

What does the study involve?
Participants undergo controlled ovarian hyperstimulation and are randomly allocated to be treated with cabergoline or not. The eggs are retrieved 34 - 36 hours later. The eggs are fertilized and the resulting embryos are transferred into the womb 72 hours later. The incidence, onset and severity of OHSS are compared between the two groups.

What are the possible benefits and risks of participating?
Patients who take part in this study visit the hospital more frequently and therefore are monitored more closely. Side effects from cabergoline are extremely rare. It has been reported that cabergoline is linked with the development of cardiac valvuopathy (abnormal thickening and stiffness of the heart valves).

Where is the study run from?
Dr Samir Abbas Medical Center (Saudi Arabia)

When is the study starting and how long is it expected to run for?
July 2007 to June 2009

Who is funding the study?
Investigator initiated and funded

Who is the main contact?
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Contact information

Type(s)
Scientific

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Additional identifiers

EudraCT/CTIS number

IRAS number

ClinicalTrials.gov number

Secondary identifying numbers
SAC 05-12011

Study information

Scientific Title
Cabergoline and OHSS - a randomized controlled study

Acronym
COHSS

Study objectives
Capillary permeability is the end step of the cascade of the pathophysiology of OHSS which is associated with third space fluid accumulation and fluid shift. Vascular endothelial growth factor (VEGF) is one of the vasoactive mediators which increases capillary permeability and expressed

at a higher level in the granulose cells. The administration of a dopamine agonist in immature rats at low doses simultaneously with human chorionic gonadotropin (HCG) prevented an increase in vascular permeability and did not affect angiogenesis; the effect was due to the availability of dopamine type 2 receptors. Dopamine agonists prevent the phosphorylation of VEGF receptor 2 and reduce the in vitro and in vivo release of vasoactive angiogenic agents. As a result, vascular permeability is also reduce. Consequently, dopamine agonist has been supposed to be a potential new strategy to prevent OHSS and reduce the severity.

Ethics approval required
Old ethics approval format

Ethics approval(s)
Samir Abbas Ethics Board, December 2006

Study design
Single-center non-blinded randomized controlled study

Primary study design
Interventional

Secondary study design
Randomised controlled trial

Study setting(s)
Hospital

Study type(s)
Prevention

Participant information sheet
Not available in web format, please use the contact details to request a patient information sheet

Health condition(s) or problem(s) studied
Ovarian hyperstimulation syndrome (OHSS)

Interventions
1. Long mid luteal GnRH agonist protocol, 0.1 mg triptorelin SC. (Decapeptyl; Ferring; Germany) has been used for pituitary down regulation in both groups
2. Once pituitary down regulation has been confirmed, controlled ovarian hyperstimulation (COH) was started using fixed dose of HMG, 150- 225 IU (Menogon 75 IU,IM injections, Ferring, Germany), for 5 days, then the dose was adjusted according to response
3. When 3 leading follicles reached 18 mm, final oocyte maturation was triggered with a single dose of 5000 IU of hCG
4. On day of hCG administration, couples were randomized using computer generated list with closed opaque envelops into two groups, cabergoline group (Group I; n=100), received 0.25 mg daily for 8 days and non-cabergolone group (Group II; n=100), did not receive cabergoline
5. Transvaginal guided oocyte retrieval was performed 34 - 36 hours later
6. Both groups have been administrated 500 ml of hydroxyethyl starch (HES) over 30 minutes as a routine strategy in our center on the day of ovum pickup
7. Ultrasound guided transfer (ET) of 2-3 embryos was performed 72 hours later

8. Luteal phase was supported with 400 mg progesterone vaginal pessaries, twice daily up to the day of pregnancy test (Cyclogest; Cox Pharmaceuticals, Whiddon Valley, UK)
9. Haemoconcentration, presence of ascitis, measuring the perpendicular diameter of free fluid in Douglas Pouch, and the ovarian volume have been reported in both groups on day of ET

Intervention Type
Drug

Phase
Not Applicable

Drug/device/biological/vaccine name(s)
Cabergoline

Primary outcome measure
Incidence, onset and severity of OHSS

Secondary outcome measures
1. Oocyte recovery rate
2. Number of mature oocytes
3. Fertilization rate
4. Clinical pregnancy rate (defined as presence of fetal heart pulsation 2 weeks after a positive β-HCG test)

Overall study start date
01/07/2007

Completion date
01/06/2009

Eligibility

Key inclusion criteria
1. 200 infertile couples undergoing ICSI and at risk of developing OHSS have been included between January 2007 and July 2009
2. The risk to develop OHSS has been defined as follows:
2.1. Dopamine E2 level on day of hCG > 3500 pg/ml
2.2. With ≥ 20 follicles > 12 mm

Participant type(s)
Patient

Age group
Adult

Sex
Female

Target number of participants
200

Key exclusion criteria
Patients with dopamine E2 ≥5000 pg/ml

Date of first enrolment
01/07/2007

Date of final enrolment
01/06/2009

Locations

Countries of recruitment
Saudi Arabia

Study participating centre
Dr Samir Abbas Medical Center
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Sponsor information

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Funder(s)

Funder type
Other

Funder Name
Investigator initiated and funded

Results and Publications

Publication and dissemination plan
Not provided at time of registration

Intention to publish date

Individual participant data (IPD) sharing plan

IPD sharing plan summary
Not provided at time of registration

Study outputs

Output type	Details	Date created	Date added	Peer reviewed?	Patient-facing?
Results article	results	01/09/2010		Yes	No