NATURAL IVF CYCLE WITH AND WITHOUT HCG ADMINISTRATION

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Ovarian hyperstimulation

- Higher pregnancy rates
- but
- $\circ \ \textbf{expensive}$
- o complications!



(OHSS, multiple pregnancies, extrauterine pregnancies, ...)

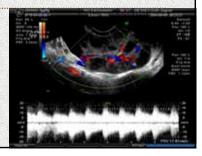
Complications of ovarian hyperstimulation

- o OHSS (24%)
- MULTIPLE PREGNANCIES (22% gemini, 4,3% trigemini, 0,4% quadrigemini)
- SPONTANEOUS ABORTIONS ? (20,9% in IVF pregnancies, 10% in spontaneous pregnancies)

Potential health risks of repeated hyperstimulation procedures

OVARIAN FUNCTION?

o OVARIAN
CARCINOMA ??



Complications of ovarian hyperstimulation

- High costs (35000 USD / newborn) Daya, 1995.
- Prolonged i.m. and/or s.c. administration of stimulation drugs
- Longer and more painful aspiration procedure

NATURAL CYCLE IVF

COST EFFECTIVE: average cost per one newborn is 5 times lower if the pregnancy is achieved by IVF in the natural cycle

1.Daya S, Gunby J, Hughes EG. Natural cycles for in-vitro fertilization: cost-effectiveness analysis and factors influencing outcome. Hum Reprod 1995;10:1719-1724

NATURAL CYCLE



ALTERNATIVE TO THE STIMULATED CYCLE

TIMING OF OOCYTE RETRIEVAL

- Most studies (19) ultrasound measurements starting cycle day 6-11 and ovulation triggering with HCG when the follicle is 15-20 mm
- The interval between HCG injection (2500 – 10000) and oocyte retrieval 31-36 h

Pelink et al, 2002. Hum Reprod Update

TIMING OF OOCYTE RETRIEVAL – studies using HCG

- o Foulot (1989. 71 patients)
- o Aboulghar (1995. 58 patients)
- o Tomažević (1996. 73 patients)
- o Janssens (2000.- 50 patients)
- o Nargund (2001. 52 patients)
- o Tomažević (2007. 397 cycles)
- o Vlaisavljević (2007.- 1024 cycles)
- o Schimberni (2008. 500 cycles)

TIMING OF OOCYTE RETRIEVAL

- In 2 studies monitoring is carried out by serial measurements of E2 and LH, twice daily until the detection of LH surge (Zayed, 1997. Hum Reprod, Lenton 2007.Reprod Biome Online)
- Egg collection is ideally timed to be 34-36 h from the onset of the spontaneous LH surge

TIMING OF OOCYTE RETRIEVAL

- Lenton 2007 comparison with (222 cycles)/without HCG (495 cycles)
- Without: 81% successfull OR, clinical pregnancies 14,8% per ET, 9,4% per oocyte, 7,7% per cycle
- With HCG: 76% successfull OR, clinical pregnancies 13,5% per ET, 8,9% per oocyte, 6,7% per started cycle

BUT

- o 7-day IVF laboratory service
- Egg collection schedueled from 09,00 till 17,00 h (working hours 07,00 – 21,00)

TIMING OF OOCYTE RETRIEVAL

- Efficacy of natural cycle IVF: a review of the literature (Pelinck et al, 2002.)
- o 20 studies, total of 1800 cycles
- With+without hCG
- o 129 pregnancies
- o 15,8% per ET
- o 7,2% per started cycle

TIMING OF OOCYTE RETRIEVAL

- o Without HCG:
- Lenton 495 cycles 38 pregnancies
 14,8% per Et 7,7% per started cycle
- Zayed 145 cycles 12 pregnancies
 13,5% per ET 6,5% per started cycle

TIMING OF OOCYTE RETRIEVAL

- Sveti Duh Hospital with 5000/6250
 IU HCG (US-follicle diameter, endometrium)
- o 896 started natural cycles
- 63 pregnancies (15,9% per ET
 9,4% per aspiration
 7% per started cycle)

Conclusion TIMING OF OOCYTE RETRIEVAL

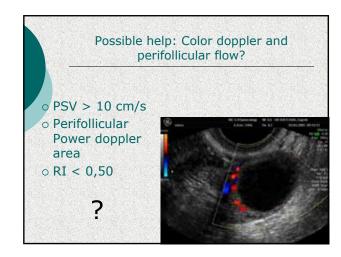
5000 - 10000 HCG!

Possible help in order to reduce the number of premature ovulations and negative aspiration procedures:

- Individualization of patients
- o E2 measurements
- o LH measurements
- o Indomethacin pretreatment (50 mg orally in the morning and 100 mg rectally at night prior to aspiration procedure-*Lenton 2007.* or everyday from follicle diameter of 14 mm *Kadoch 2008.*)

Possible help:

- Aspiration needles with double lumen with flushing of follicles (91% positive aspirations with flushing / 68,5% with single lumen, Daya 1995)
- o Use of GnRH antagonists



Conclusion TIMING OF OOCYTE RETRIEVAL 5000 - 10000 HCG!

