

Ultrasound in reproductive medicine

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**ULTRASOUND IN NATURAL CYCLE IVF: IS IT
GOOD ENOUGH?**

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To obtain a good oocyte is a fundamental step to
maximize the efficiency of natural cycle IVF

Major problem in natural cycle IVF is the high percentage
of failure to collect oocytes from each cycle

Due to mistakes in the timing of retrieval, to a deficient
follicular growth or to a precocious LH surge

US

- dominant follicle has a linear growth rate between 1,4 and 2,2 mm per day until the day of LH peak
- when LH surge it measures 18,1 – 22,6 mm

- direct correlation between follicular growth and E2 level
- endometrial shape and thickness reflects E2 levels

Timing of oocyte retrieval

TVUS + CD, PD, 3D

E2 measurements

LH measurements

Additional help:

Indomethacin (50 mg oral + 100 mg rectal night before OPU Lenton EA, 2007 or everyday from F>14 mm Kardoch 2008.);

Double lumen aspiration needles with follicle flushing (91%+OPU vs. 68,5% Daya 1995)

GnRH antagonists

Studies with US, E2, LH

- Tomažević T (2011) – 1167 cycles - 943 oocytes (80%) - 613 ET (65%) - pregnancy per cycle 9% - pregnancy per ET 17%
- Take home baby rate 7%
- E2>0,39 nmol/L (>104 pg/ml)
- f>16 mm
- Endometrial thickness > 5 mm
- When minimal criteria fulfilled LH urinary test ; if negative HCG
- Aspiration 31-32 h after HCG

Studies with US, E2, LH

- Janssens RMJ (2000) f>18 mm; E2; LH<15 IU/L
- 75 cycles; 14 (18,6% cancelled); 61 aspirations (81,3%); 50 oocytes (66,6%); 35 ET (46,6%); 9,3% pregnancy rate per cycle; 20% pregnancy rate per ET

US + E2 + LH

- Kim SH (1996) 80 cycles – 67 OR - 64 +OR – PR
10% /c; 12,5% /ET
- Paulson RJ (1992) 101 cycles - 78 OR - PR /c 10,9%;
PR /ET 14,5%
- Vlaisavljević V (2001) 362 cycles – 318 OR – 269+OR
– PR /c 10,2% - PR /ET 20,9%
- Tomažević T (2009) 397 cycles – 303 +OR – PR /c
12% - PR /ET 23%

Vlaisavljević V (2001) – three protocols for
monitoring 587 nIVF cycles

- A/ only US (f>18 mm)
- B/ f>18 mm or E2>0,91 nmol/L
- C/ f> 15 mm + E2>0,49 nmol/L
- Cancellation A=**40,5%**; B=27,8%; C=**9,8%**
- Oocyte recovery rate A= 78,9% B=88,9% C=90%
- PR / OPU C=12,8% B= 5,5% A=10%
- PR / ET C= 24,6% B=10% A= 16,6%

US + LH

- Busso C (2007) – 70 cycles - 52 aspirations - 77%
with oocyte (40) – 18 cancelations
- f > 18 mm and LH test negative
- 11,4% PR /c and 29,6% /ET

Studies with just US

- Schimberni M (2011) f >16 mm – 1000 cycles - 788 oocytes - 618 ET
- 12,8% /cycle and 20,7% /ET

Just US / Sv Duh (f>15 mm, end>6/7 mm)
2002-2012

- 1324 NC
- 1044 aspirations
- 830 oocytes (62% per cycle; 79,5% per aspiration)
- 620 ET
- 90 pregnancies
- PR 6,8% /c; 14,5% /ET
- Take home baby rate 5,6%

Just US / Sv Duh (f>15 mm, end>6/7 mm)
2009-07/2012

- according the legislation older patients
- no weekend OPU
- 23 - 50 years old, median 37
- 412 cycles, 353 oocytes
- 250 ET, 30 pregnancies
- PR /c 7,3 %
- PR /ET 12%

CD; PD; 3D; 4D

- time consuming
- perifollicular BF velocities gradually increase in the periovulatory period while the PI remains constant and RI in perifollicular vessels shows low to moderate values. This values suggest increase in blood flow (Bourne TH, 1991)
- the blood volume does not differ between follicles containing an oocyte and those with no oocyte in the aspirate

No significant increase in PSV between fertilization/nonfertilization cycles but decrease in PI and RI (Gavrić Lovrec V, 2001)

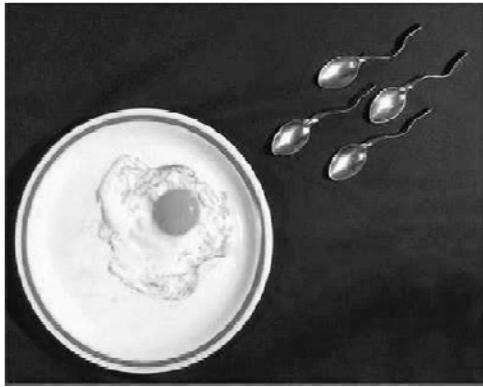
3D CD/PD

- controversial
- hypothesis – follicles containing oocytes able to produce a pregnancy – more uniform perifollicular vascular network (Vlaisavljević V, 2003)
- reduction of perifollicular arterial blood flow resistance after hCG administration is a good indicator of the recovery of mature oocytes in ART treatment (Nakagawa K, 2006)
- increase in perifollicular capillary network volume after hCG administration (Vlaisavljević V, 2010)

Conclusions

ULTRASOUND IN NATURAL CYCLE IVF: IS IT GOOD ENOUGH?

- **US + E2 + LH!** (PR/ET 12,5 – 24,6% - 42,8% bc ET !?)
- US + LH (PR/ET 29,6% !?)
- US (PR/ET 12-20,7%)



References

- Natural cycle IVF with and without terminal HCG: learning from failed cycles. Lenton EA. *Reprod Biomed Online*. 2007 Aug;15(2):149-55.
- Changes in perifollicular vascularity after administration of human chorionic gonadotropin measured by quantitative three-dimensional power Doppler ultrasound. Vlasisavljević V, Borko E, Radaković B, Zazula D, Dosen M. *Wien Klin Wochenschr*. 2010 May;122 Suppl 2:85-90
- Embryo transfer and luteal support in natural cycles. Vlasisavljević V. *Reprod Biomed Online*. 2007 Jun;14(6):686-92. Review.
- Measurement of perifollicular blood flow of the dominant preovulatory follicle using three-dimensional power Doppler. Vlasisavljević V, Reljić M, Gavrić Lovrec V, Zazula D, Sergent N. *Ultrasound Obstet Gynecol*. 2003 Nov;22(5):520-6.
- Dependence of the in vitro fertilization capacity of the oocyte on perifollicular flow in the preovulatory period of unstimulated cycles. Gavrić Lovrec VG, Vlasisavljević V, Reljić M. *Wien Klin Wochenschr* 2001;113(Suppl 3):21-26.
- Results of intracytoplasmic sperm injection of single oocyte in 362 unstimulated cycles. Vlasisavljević V, Kovacic B, Reljić M, Lovrec VG, Sajko MC. *J Assist Reprod Genet*. 2002 Mar;19(3):127-31.
- Three protocols for monitoring follicle development in 587 unstimulated cycles of in vitro fertilization and intracytoplasmic sperm injection. A comparison. Vlasisavljević V, Kovacic B, Reljić M, Lovrec VG. *J Reprod Med*. 2001 Oct;46(10):892-8.
- Value of the serum estradiol level on the day of human chorionic gonadotropin injection and on the day after in predicting the outcome in natural in vitro fertilization/intracytoplasmic sperm injection cycles. Reljić M, Vlasisavljević V, Gavrić V, Kovacic B, Cizek-Sajko M. *Fertil Steril*. 2001 Mar;75(3):539-43.
- Age, oestradiol and blastocysts can predict success in natural cycle IVF-embryo transfer. Tomazević T, Korosec S, Virant Klun I, Drobnic S, Verdenik I. *Reprod Biomed Online*. 2007 Aug;15(2):220-6.

References

- Intrafollicular blood flow during human ovulation. Bourne TH, Jurkovic D, Waterstone J, Campbell S, Collins WP. *Ultrasound Obstet Gynecol*. 1991 Jan 1;1(1):53-9.
- Natural-cycle in vitro fertilization in poor responder patients: a survey of 500 consecutive cycles. Schimberni M, Morgia F, Colabianchi J, Giallonardo A, Piscitelli C, Giannini P, Montigiani M, Sbracia M. *Fertil Steril*. 2009 Oct;92(4):1297-301.
- In vitro fertilization in unstimulated cycles: the University of Southern California experience. Paulson RJ, Sauer MV, Francis MM, Macaso TM, Lobo RA. *Fertil Steril*. 1992 Feb;57(2):290-3.
- Efficacy of natural cycle IVF: a review of the literature. Pelinck MJ, Hoek A, Simons AH, Heineman MJ. *Hum Reprod Update*. 2002 Mar-Apr;8(2):129-39. Review.
- Simultaneous program of natural-cycle in vitro fertilization and cryopreserved-thawed embryo transfer. Kim SH, Kim CH, Suh CS, Moon SY, Lee JY, Chang YS. *J Assist Reprod Genet* 2006;13:716-721
- In-vitro fertilization in a spontaneous cycle: easy, cheap and realistic. Janssens RM, Lambalk CB, Vermeiden JP, Schats R, Schoemaker J. *Hum Reprod*. 2000 Feb;15(2):314-8.
- Spontaneous ovulation rate before oocyte retrieval in modified natural cycle IVF with and without indomethacin. Kadoch IJ, Al-Khaduri M, Phillips SJ, Lapensée L, Couturier B, Hemmings R, Bissonnette F. *Reprod Biomed Online*. 2008 Feb;16(2):245-9.
- Factors associated with success in natural IVF cycle. Tomazević T, Virant Klun I, B Podlep, S Drobnic, I Bacer, Kermavner, B Valenincic, J Kmed, J Mivsek. 2011 *Giornale Italiano Vol 33 Issue 5*
