## Does the ovulation pattern in consecutive menstrual cycles

 affect the sex ratio of subsequent conceived children?Misao Fukuda, Kiyomi Fukuda, Kenichi Tatsumi’, Takashi Shimizu\#, Miho Nobunaga\#, Anne Grete Byskov*, Claus Yding Andersen*

M\&K Health Institute
30-9 Kariya, Ako, Hyogo 678-0239, Japan
"Umegaoka Women's Hospital
1-33-3 Umegaoka, Setagaya-ku, Tokyo 154-0022, Japan
\#Shimizu Women's Clinic
2-2-4 Minamiguchi, Takarazuka, Hyogo 665-0011, Japan
*Laboratory of Reproductive Biology, Juliane Marie Center, Rigshospitalet, Section 5712, University Hospital of Copenhagen,

Blegdamsvej 9, DK-2100, Copenhagen, Denmark

## Ovulation patterns

O.: Ovulation + oocyte

O : Aovulation
Contralateral ovulation: LR, RL
Ipsilateral ovulation: RR, LL


1 cycle 2 cycles 3 cycles
Previous second cycle
Previous first cycle


Treatment cycle

LR
LLR
 RLR 0
0


R
RR LRR


## RRL

RL
LRL
L


LL


LLL


0
0

## 8 ovulation patterns and IVF outcomes in 260 SP and 315 CC cycles

| 1 cycle | R |  |  |  | L |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 cycles | LR |  | RR |  | RL |  | LL |  |
| 3 cycles | LLR | RLR | LRR | RRR | RRL | LRL | RLL | LLL |
| Oocyte retrieval rate(\%) | $\begin{gathered} 79 / 86 \\ (92) \end{gathered}$ | $\begin{gathered} 93 / 99 \\ (94) \end{gathered}$ | $\begin{gathered} 76 / 94 \\ (81) \end{gathered}$ | 62/81 <br> (77) | $\begin{gathered} \hline 71 / 74 \\ (96) \end{gathered}$ | $\begin{gathered} 90 / 103 \\ (87) \end{gathered}$ | $48 / 67$ (72) | 41/58 <br> (71) |
| Fertilization rate(\%) | 72/79 <br> (91) | $\begin{gathered} 79 / 93 \\ (85) \end{gathered}$ | 50/76 <br> (66) | $\begin{gathered} 43 / 62 \\ (69) \end{gathered}$ | $61 / 71$ <br> (86) | $\begin{gathered} 83 / 90 \\ (92) \end{gathered}$ | 35/48 <br> (73) | 23/41 <br> (56) |
| Polyspermia rate(\%) | $\begin{gathered} \hline 8 / 79 \\ (10) \end{gathered}$ | 0/93 <br> (0) | $4 / 76$ <br> (5) | 0/62 <br> (0) | $\overline{1 / 71}$ <br> (1) | $1 / 90$ <br> (1) | $0 / 48$ <br> (0) | 0/41 <br> (0) |
| Cleavage rate(\%) | $\begin{aligned} & 60 / 72 \\ & (83) \end{aligned}$ | $66 / 79$ <br> (84) | 36/50 <br> (70) | $\begin{gathered} 30 / 43 \\ (70) \end{gathered}$ | 54/61 <br> (89) | $\begin{aligned} & 72 / 83 \\ & (87) \\ & \hline \end{aligned}$ | $\begin{gathered} 25 / 35 \\ (71) \end{gathered}$ | $\begin{gathered} 18 / 23 \\ (78) \\ \hline \end{gathered}$ |
| Rate of preembryo formation | $60 / 72$ <br> (75) | 66/93 <br> (71) | 36/76 <br> (46) | $\begin{gathered} 30 / 62 \\ (48) \end{gathered}$ | 54/71 <br> (76) | $\begin{gathered} 72 / 90 \\ (80) \end{gathered}$ | $\begin{gathered} 25 / 48 \\ (52) \end{gathered}$ | 18/41 <br> (43) |
| Pregnancy rate(\%) | $\begin{aligned} & 17 / 78 \\ & (22) \end{aligned}$ | $\begin{gathered} 13 / 88 \\ (15) \end{gathered}$ | 5/78 <br> (6) | $3 / 64$ <br> (5) | $\begin{aligned} & 10 / 68 \\ & (15) \end{aligned}$ | $\begin{gathered} 12 / 94 \\ (13) \end{gathered}$ | $\begin{array}{r} \hline 8 / 59 \\ (14) \\ \hline \end{array}$ | 3/46) <br> (7) |
| Implantation rate(\%) | $\begin{gathered} 17 / 60 \\ (28) \end{gathered}$ | $\begin{gathered} 13 / 66 \\ (20) \end{gathered}$ | $\begin{aligned} & 5 / 36 \\ & (14) \end{aligned}$ | $\begin{array}{r} 3 / 30 \\ (10) \end{array}$ | $\begin{gathered} 10 / 54 \\ (19) \end{gathered}$ | 12/72 <br> (17) | $\begin{array}{r} 8 / 25 \\ (32) \end{array}$ | $\begin{array}{r} \hline 3 / 18 \\ (17) \end{array}$ |

8 ovulation patterns and pregnancy outcome in 872 SP ( 612 IUI + 260 IVF) and 746 CC ( 431 IUI + 315 IVF) cycles

| 1 cycle | $R$ |  |  |  | R |  |  | RR |  |
| :--- | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| 2 cycles | LR |  | RR |  |  |  |  |  |  |
| 3 cycles | LLR | RLR | LRR | RRR | RRL | LRL | RLL | LLL |  |
| SP+CC <br> Total | $54 / 202$ | $44 / 264$ | $34 / 217$ | $22 / 219$ | $33 / 207$ | $27 / 218$ | $16 / 148$ | $10 / 143$ |  |
| (\%) | $(26.7)$ | $(16.7)$ | $(15.7)$ | $(9.6)$ | $(15.9)$ | $(12.4)$ | $(10.8)$ | $(7.0)$ |  |
| Live <br> birth | $44 / 202$ | $36 / 264$ | $23 / 217$ | $20 / 219$ | $25 / 207$ | $20 / 218$ | $13 / 148$ | $8 / 143$ |  |
| (\%) | $(21.8)$ | $(13.6)$ | $(10.6)$ | $(9.1)$ | $(12.2)$ | $(9.2)$ | $(8.8)$ | $(5.6)$ |  |

SP+CC Total: LLR vs RLR, LRR, RRR, RRL, LRL. RLL, LLL. All P<0.012

## Comparison of statistical power of pregnancy rate among different ovulation pattern groups

Contralateral or ipsilateral ovulation pattern in the immideate previous cycle:

$$
\begin{gathered}
\text { Contralateral : LLR + RLR + RRL + LRL }=158 / 891 \text { (17.7 \%) } \\
\text { Ipsilateral : } \mathrm{LRR}+\mathrm{RRR}+\mathrm{RLL}+\mathrm{LLL}=82 / 727 \text { (11.3\%) } \\
\mathrm{P}=0.0004
\end{gathered}
$$

Contralateral or ipsilateral ovulation pattern in the previous second cycle:
Contralateral : LLR + LRR + RRL + RLL = 137/774 (17.7 \%)* Ipsilateral : RLR + RRR + LRL + LLL = 103/844 (12.2 \%)* $P=0.0024$

Right-sided ovulation (R) vs Left-sided ovulation (L)

$$
R: L L R+R L R+L R R+R R R=154 / 902 \text { (17.1\%)\# }
$$

L: RRL + LRL + RLL + LLL = 86/716 (12.0 \%)\#
$P=0.0055$
Thus the LLR pattern results in enhanced pregnancy rates

## Ovulation patterns and offspring sex ratio (males/males+females)

| Fertile women | Infertile women |  |  |
| :---: | :---: | :---: | :---: |
| $\begin{array}{rl} R & 210 / 410 \\ (0.512) \end{array}$ | 1 cycle | 2 cycles | 3 cycles |
|  | $\begin{aligned} \text { R } 226 / 390 \\ (0.579) \end{aligned}$ | $\begin{gathered} \text { LR } 93 / 148 \# \\ (0.628) \end{gathered}$ | LLR 48/63 |
|  |  |  | (0.762) |
|  |  |  | RLR 24/49 |
|  |  |  | (0.490) |
|  |  | $\begin{gathered} \text { RR 64/116 } \\ (0.552) \end{gathered}$ | LRR $16 / 38$$(0.421)$ |
|  |  |  |  |
|  |  |  | RRR 21/38 |
|  |  |  | (0.553) |
| $\begin{array}{r} \text { L 132/258 } \\ (0.512) \end{array}$ | $\begin{array}{r} \text { L } \begin{array}{r} 128 / 240 \\ (0.533) \end{array} \end{array}$ | $\begin{array}{r} \text { RL } 55 / 100 \\ (0.550) \end{array}$ | $\begin{array}{r} \text { RRL } 25 / 46 \\ (0.543) \end{array}$ |
|  |  |  |  |
|  |  |  |  |
|  |  | $\begin{gathered} \text { LL } \begin{array}{c} \text { 25/62\# } \\ (0.403) \end{array} \end{gathered}$ | $\begin{gathered} \\ \text { RLL } \\ 8 / 2.577) \\ (0.348) \end{gathered}$ |
|  |  |  |  |
|  |  |  |  |
|  |  |  | LLL 5/16 |
|  |  |  | (0.313) |
| C=LR+RL=148/248 (0.597)* |  |  |  |
| $\mathrm{I}=\mathrm{RR}+\mathrm{LL}=89 / 178$ (0.500)* |  |  |  |
| *: P=0.0488 \#: P=0.0036 |  |  |  |
| LLR vs RLR, LRR, RRR, RRL, RLL, LLL All P<0.05 |  |  |  |

Offspring sex ratio following anovulation of infertile women

Induced+spontaneous anovulation

|  | males | females | males/males+females |
| :--- | :---: | :---: | :---: |
| Before+after | 137 | 111 | $137 / 248(0.552)^{\star} \#$ |
| First cycle | 24 | 10 | $24 / 34(0.706)$ |
| Second cycle | 24 | 7 | $24 / 31(0.774) \#$ |
| First+second cycle | 48 | 17 | $48 / 65(0.738)^{*}$ |

*: P=0.0070, \#: P=0.0372

Offspring sex ratio following lactational anovulation of fertile women


Summary: Ovulation patterns, pregnancy potential and offspring sex ratio


Contralateral ovulation (LLR)
Right ovary
o o o- 0
Left ovary
o о о оо0000ОО○ 000.000

Ipsilateral ovulation (LLL)
Right ovary
Ovulation Pregnancy Offspring Strong CLpatterns rate (\%) sex ratio
$\bigcirc_{\text {LLR }}$ 27\%(54/202) $0.762(48 / 63)$ ooooดЯO○○ LLR 27\%(54/202) 0.762(48/63)

- o o

Left ovary

o o O.OOo

## Conclusion

Ovulation from the right ovary in the group of infertile women tended to result in boys more often than the left ovary, which, however, was not observed in the group of fertile women. Ovulation jumping from one ovary to the other during two consecutive cycles seems to enhance the frequency of boys and especially the ovulation pattern left-leftright (LLR) observed during three consecutive cycles results in boys in three out of four
pregnancies. Induced and spontaneous anovulation also seems to enhance the frequency of boys in the first and second subsequent cycles, increasing the ovulation pattern of LLR.

