## **Dynamic Fertility Series**

Providing the Best of East and Western Medicine





Paul C Magarelli, M.D., Ph.D.
Board Certified
Reproductive Endocrinologist &
Infertility Specialist

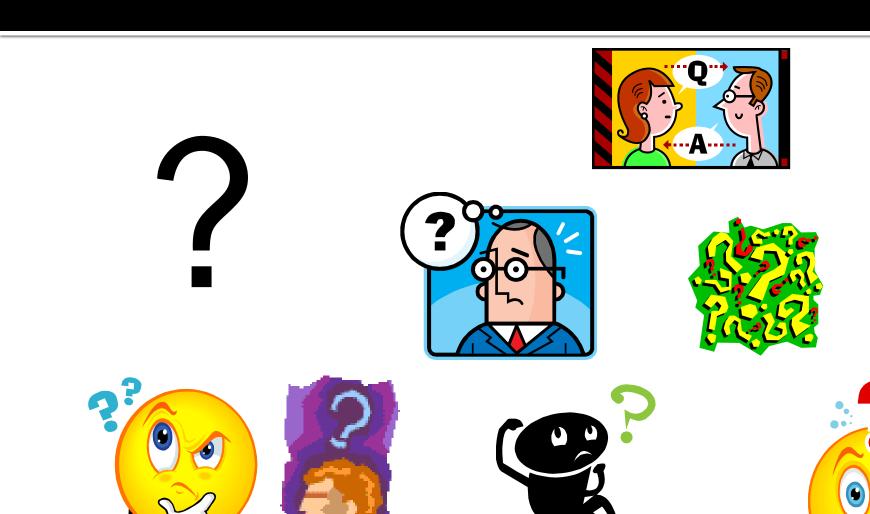


Reproductive Medicine & Fertility Centers 265 Parkside Drive, Suite 200 Colorado Springs, CO 80910



719-475-2229 info@RMFCFertility.com

### What questions can I answer for you?



### What can we do to help you create a baby?

- Tell you what's wrong
  - Lab Tests
  - Physical Exam and Ultrasound
- Give you choices and what each choice will entail
  - Treatment Consult
- Tell you the FULL costs
  - Financial Consult
- Help you with financials
  - Payment plans (only practice still doing this!)
  - Discounts for Military
  - Coupons worth 100's of dollars
  - Education seminars worth \$ 1,000 dollars



# Today's Fertility rituals!

Sex, then more sex, then timed sex, then more sex, then resentful sex, then planned sex, then concerned sex, then worried sex, then call friends, then more sex, then call MOM, then really worried sex, then begrudged sex, then call the doctor!



# **Embrace the Journey!**



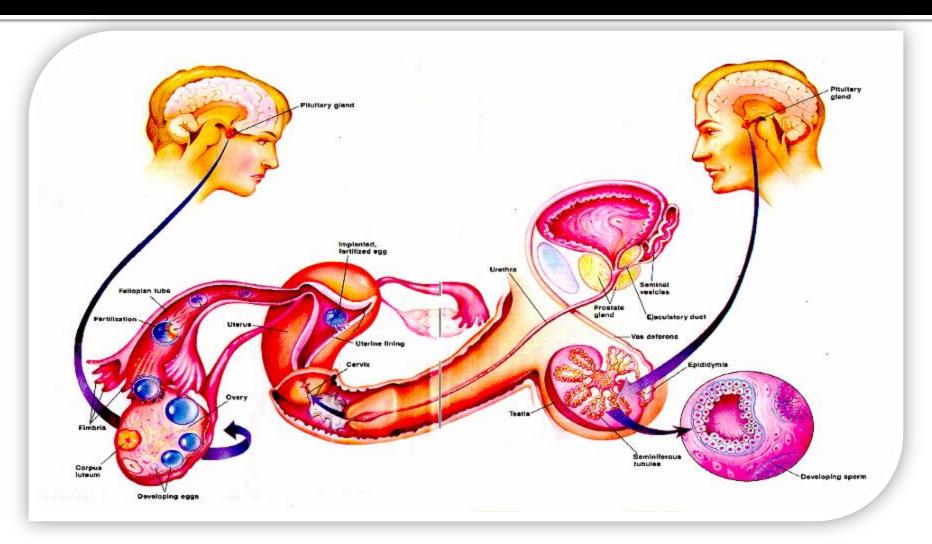


Our job is to point the way!

Our job is to help you avoid dead ends!



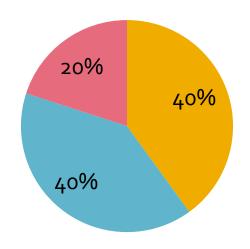
# Biology of Reproduction



## Causes of Infertility

#### Causes of Infertility

■ Male ■ Female ■ Couple



#### Male

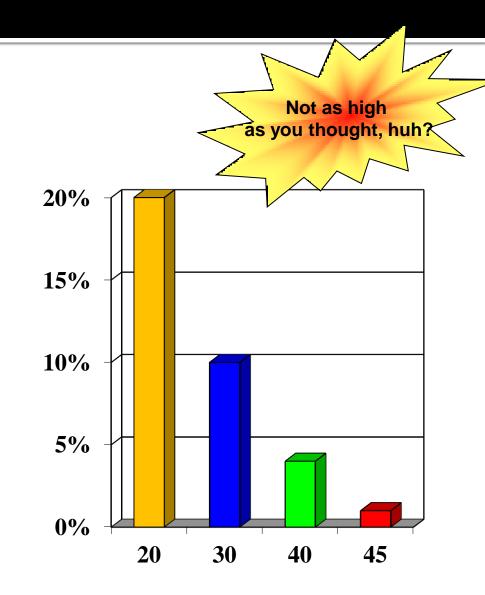
- Sperm Shape
- Sperm Motility
- Sperm Count
- Sperm Function
- Infection of Prostate
- Varicose veins in Testicles
- Genetics

#### Female

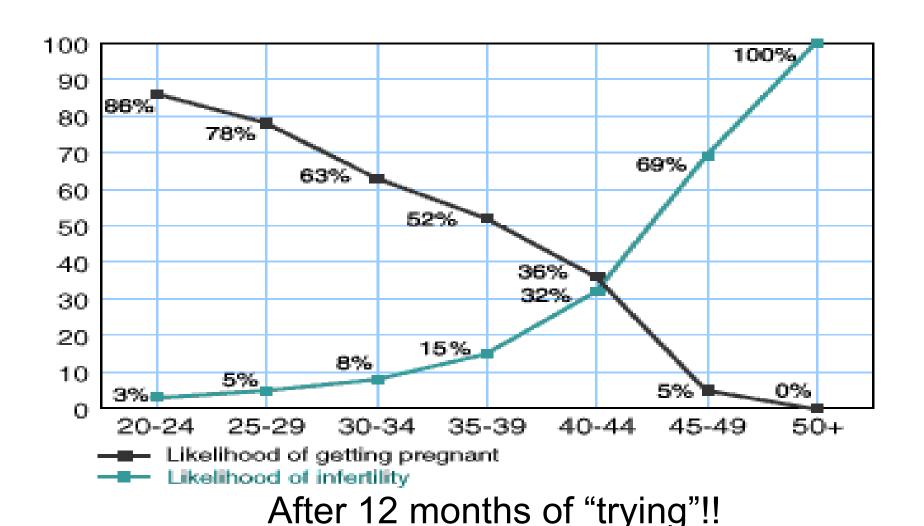
- Eggs
- Uterus
- Fallopian Tubes
- Hormones
- Infections
- Genetics

# What are your chances of getting pregnant by intercourse alone at age:

- Age 15
  - 20% per month
- Age 30
  - 10% per month
- Age 40
  - 3% per month
- Age 45
  - << 1% per month</li>



# Chance of Getting Pregnant versus Chance of Being Infertile

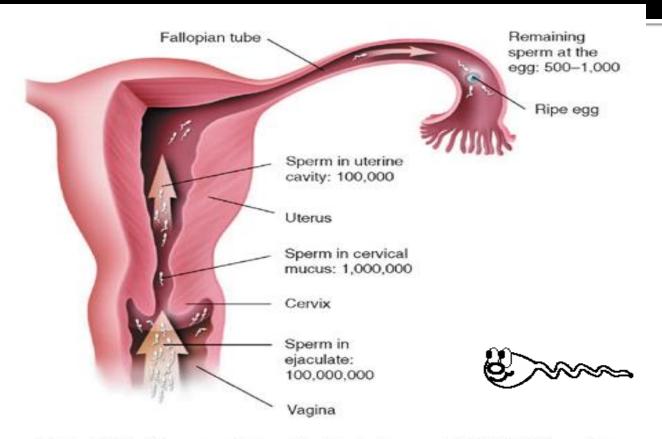




### **Question:**

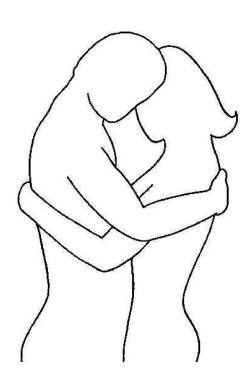
How many sperm must be ejaculated to have "enough" to fertilize an egg?

### **Answer: Lots!**



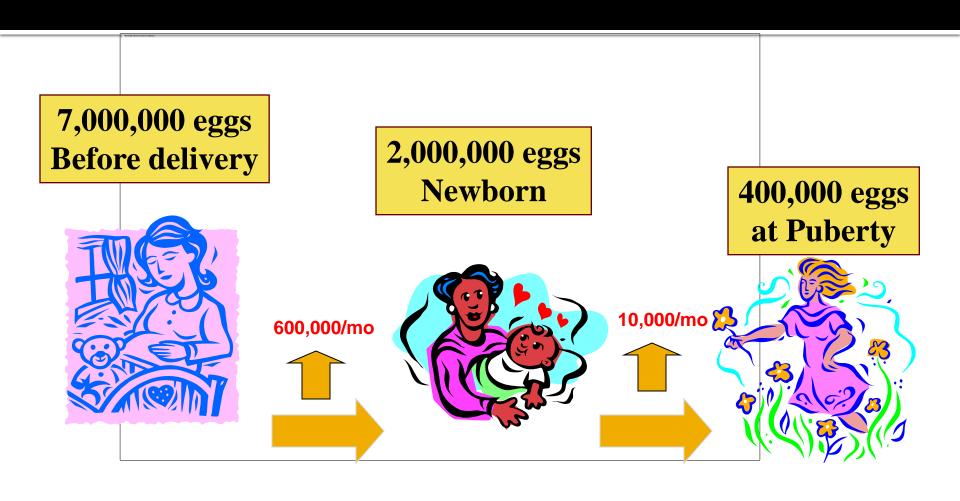
Of about 100 million sperm that are ejaculated, only around 500 to 1,000 remain by the time they reach the egg at the outer end of the fallopian tube.

### Why is it so hard to create a family?





# Answer: Female Eggs, a limited resource!





400,000 eggs

at Puberty

13 yo

We lose 1000 eggs per month

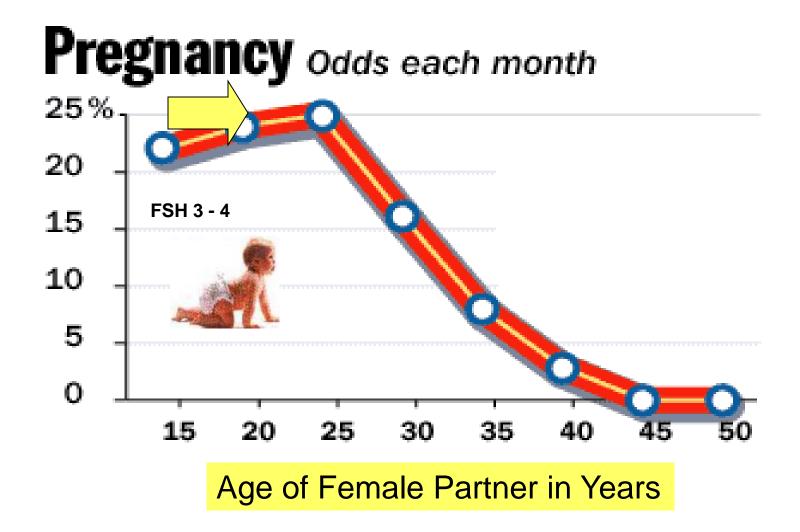
of each menstrual cycle, whether you ovulate or not or even if you are pregnant or on birth control pills!!

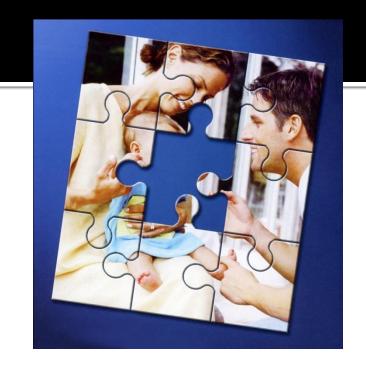
We only have 400 ovulations in our lives!



Zero (0) eggs at Menopause 50 yo

### Impact of Loss of Eggs



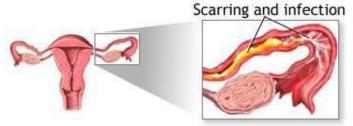


# The Four Questions

### **Initial Infertility Workup**

### **Four Simple Questions**

- Q1:
  - Are there sperm?
- Q2:
  - Are there eggs?
- Q3:
  - Can the sperm and egg meet?
- Q4:
  - Is there a safe place to grow a baby?



Low sperm count, anatomic defects or disease and fallopian tube scarring are the most common factors that may cause infertility



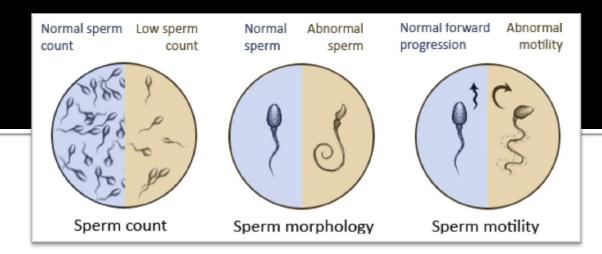
# Q1: Are there sperm?

### Semen Analysis

- How was it designed?
  - All fertile men in one room
  - All <u>infertile</u> men in another
  - Check sperm
  - Divide them into Fertile and Infertile
    - If the man did not fall into the Fertile group he is designated as
       Infertile



# Semen Analysis

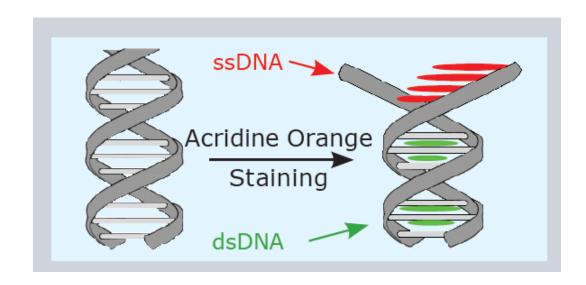


#### Count

- Need minimum of 20,000,000 sperm for each cc or ml of ejaculate
  - If 19,999,999 the male is INFERTILE
- Motility
  - Need minimum of 50% "swimmers"
    - If 49% the male is INFERTILE!
- Morphology
  - Need minimum of 15% "normal" shaped sperm
    - If 14% the male is INFERTILE!

# What about Sperm Function?

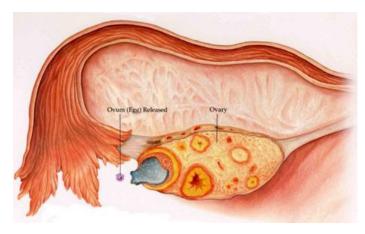
- Sperm DNA Fragmentation Assay (SDFA)
  - Can my sperm work using sexual intercourse?
  - Can my sperm work using IUI or Artificial Insemination?



### Q2: Are there Eggs?

How to test if eggs are available

- Day 3 FSH, Estradiol and LH serum levels
  - If FSH < 7 mIU/ml</li>
    - normal ovarian reserve of eggs
  - If 7 to 10
    - reduced ovarian reserve
  - If > 10
    - marked reduced ovarian reserve
  - If > 15
    - very poor ovarian reserve, increased risk of miscarriages, blighted ovum
  - If > 18
    - unlikely to have eggs that will results in live birth



#### Ovarian Assessment Report<sup>™</sup>

SPECIMEN #: 001244 PATIENT: Ms. Jane Doe DATE OF BIRTH: 05/07/76

Collected: 10/08/08 RECEIVED: 10/09/08

Est. Time: 9 am Est. TIME: 10 am

PATIENT VALUES:

AMH 0.3 ng/ml Inhibin B 30 pg/ml

MENSTRUAL CYCLE DAY 3 RANGES:

DEPENDENT UPON CHOSEN CLINICAL OUTCOME

FSH 8 IU/L <10 NORMAL 10-12 BORDERLINE

>12 HIGH

PHONE:

Fax:

LH 2.2 IU/L <2 Low 2-10 NORMAL >10 HIGH

PHYSICIAN: Mr. Mohamed Menabawey

One St Thomas Street, London Bridge London SE1 9RY United Kingdom

ADDRESS: The London Bridge Fertility, Gynaecology and Genetics Centre

+44 (020) 7403 3363

+44 (020) 7403 8552

30 pg/ml <12 LOW

Estradiol

12-126 NORMAL >126 HIGH

#### Egg Retrieval Score™ REDUCED

RANGE 0 to 5 reduced 6 to 10 fair 11 to 15 good 16 to 20 excellent

RESULTS

#### COMPARED TO OTHER WOMEN 31 to 35 YEARS OLD EGG RETRIEVAL SCORE

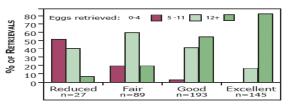
Below Average

COMMENTS: The Egg Retrieval Score indicates that this patient has a reduced chance of retrieving 5 or more eggs through ovarian stimulation. Compared to others in this patient's age group, the Egg Retrieval Score is below average. Ovarian reserve testing alone should not be used as the sole determinant for treatment recommendations.

#### OUTCOMES OF 454 EGG RETRIEVALS

CHANCE OF

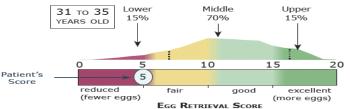
SUCCESSFUL RETRIEVAL



EGG RETRIEVAL SCORE

The Egg Retrieval Score<sup>™</sup>, which ranges from 1 to 20, is an index derived from age along with blood levels of several menstrual cycle day 3 hormones including AMH, inhibin B, FSH, and estradiol. In the graph above, the number of eggs obtained per retrieval from 244 women undergoing 454 egg retrievals is displayed in categories defined by the associated Egg Retrieval Score<sup>™</sup> (ERS): reduced, fair, good, and excellent. Within each category, column height represents the percent of retrievals resulting in 0 to 4 eggs, 5 to 11 eggs or 12 or more eggs. eggs, or 12 or more eggs.

#### Egg Retrieval Scores IN PATIENT'S AGE GROUP



Displayed in the graph above is a plot of the frequency (y-axis) of each Egg Retrieval Score (x-axis) in women of similar age to the patient receiving evaluation at a fertility clinic. The frequency distribution is divided into three sections by dotted lines to indicate below average (the lower 15%), average (middle 70%), and above average (upper 15%).

ReproSource Fertility Diagnostics<sup>™</sup> has performed extensive clinical studies validating the OAR<sup>™</sup>. The FDA recently issued a draft guidance which states that are diagnostic index derived from multiple laboratory values should be submitted to the FDA to determine if the index and manner of reporting requires review and clearance. In compliance with this new the CAR<sup>™</sup>, which also utilizes for "Research Use Only" components for AMH and inhibin B detection, is being submitted to the FDA to determine if clearance is required. Diagnostic index reports that are determined by the FDA to require clearance should, until clearance is obtained, carry the label "For investigational use only. The performance characteristics of this product have not been established."

Score

### Ovarian Assessment Report



#### Interpretation of the ERSTM

The Ovarian Assessment Report™ provides the most advanced testing system available today using a single blood sample for assessing the available egg supply for retrieval through ovarian stimulation ("ovulatory egg supply"). The ERS™ takes complex information from multiple factors related to ovulatory egg supply and combines them into a single score.

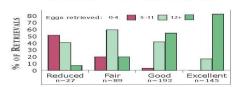
The ERS™ has been calibrated using research involving hundreds of women undergoing multiple egg retrievals <sup>1,2</sup>. The score allows patients and clinicians to better understand the chances of a successful egg retrieval which most fertility experts defined as yielding at least five eggs. However, this still does not mean that the eggs will be of sufficient quality to produce a healthy baby. Currently, age alone is the strongest single predictor of egg quality. How your age is related to chance of having good quality eggs is presented on the previous page of this report using a colored bar.

ERS™			CATEGORY
0	to	5	Reduced
6	to	10	Fair
11	to	15	Good
16	to	20	Excellent

The ERS™ ranges from 1 to 20 with chances of good egg supply increasing with higher scores. Above are ERS ranges divided into categories representing the chance of good egg supply.

#### GOLD STANDARD CLINICAL RESEARCH

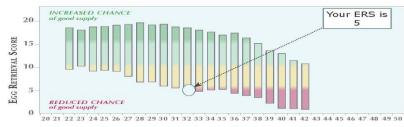
#### OUTCOMES OF 454 EGG RETRIEVALS



EGG RETRIEVAL SCORE BY CATEGORY

In the graph above, the number of eggs obtained per retrieval from 283 women undergoing 454 egg retrievals is displayed in categories defined by the associated Egg Retrieval Score (ERS): reduced, fair, good, and excellent. Within each category, the column height represents the percent of retrievals resulting in 0 to 4 eggs, 5 to 11 eggs, or 12 or more eggs <sup>1,2</sup>.

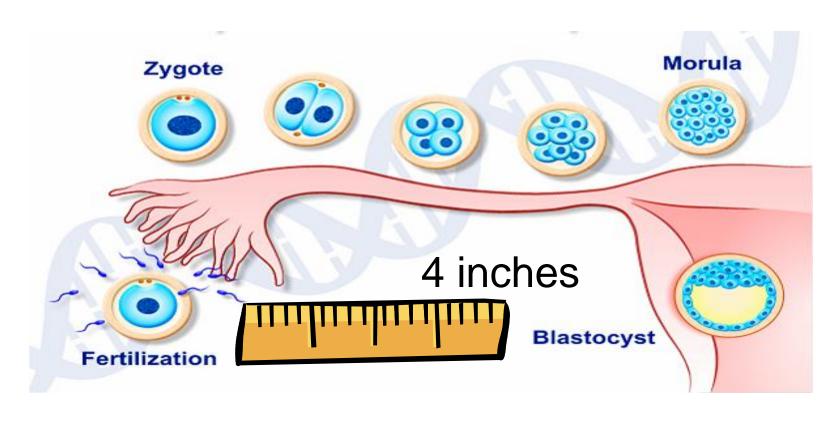
#### YOUR EGG RETRIEVAL SCORE™ (ERS) COMPARED TO OTHERS



AGE IN YEARS

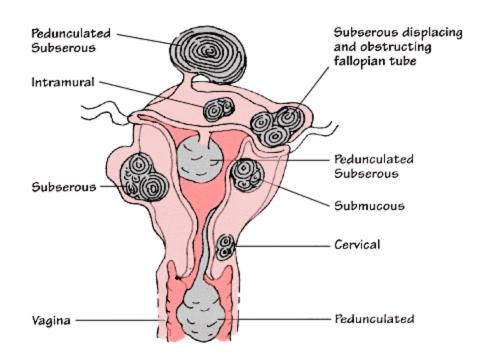
The graph to the left shows your ERS relative to other women being treated at fertility centers. Bars show two standard deviations above and below a rolling 5 year average ERS. Data is currently available for ages 22 through 42 years of age. Chance of good egg supply is indicated by color based upon the ERS: ERS 0 to 5 is red, ERS 6 to 10 is yellow, ERS 11 to 15 is light green, and ERS 16 to 20 is green.

### Q3: Can the sperm and egg meet?



\$ 600,000 to \$ 1,000,000

### **Q4**: Is there a safe place for the baby to grow?

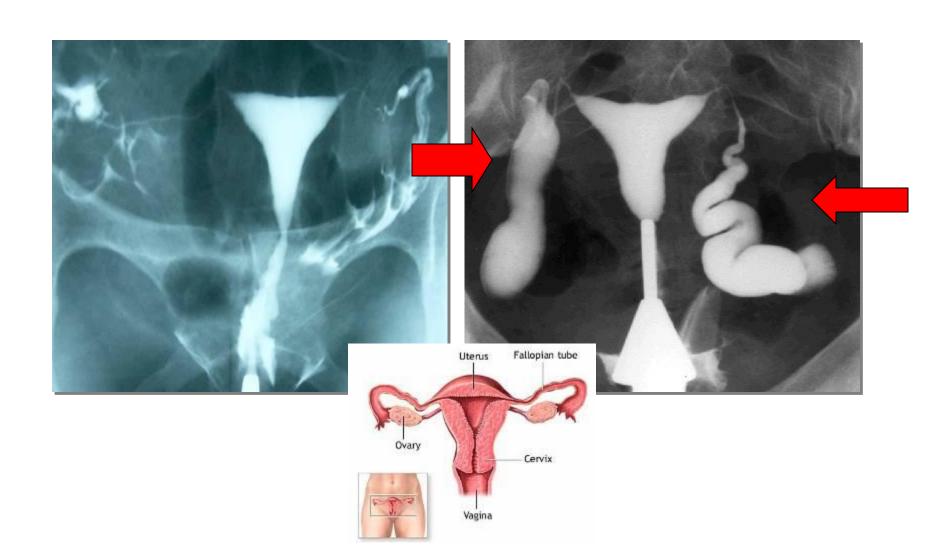




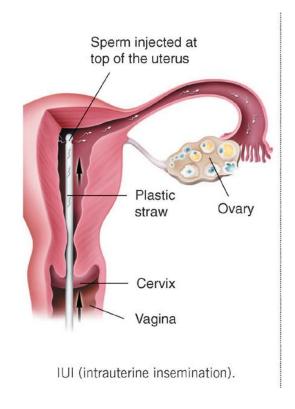
**Abnormal** 

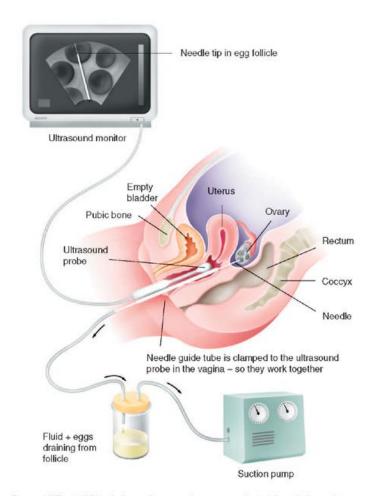
**Normal** 

### Hysterosalpingogram:(HSG) "Dye Study"



### Infertility Treatments & Outcomes

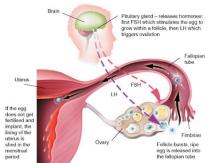




For most IVF and ICSI techniques, the woman's eggs are collected through the vagina using an ultrasound scanner to guide a needle into the follicles and draw off the fluid and eggs.

### Treatments for Infertility

- Timed Intercourse
- Clomid therapy, aka Ovulation Induction (can be used with injectable medications also, e.g., Follistim™)
- Intrauterine Insemination (IUI), aka Artificial Insemination
- Surgery for Tubal Reversal
- 5. In Vitro Fertilization (IVF) "Test Tube Baby"



The menstrual cycle is typically 28 days; for the first 14 days the egg matures in the ovan before being released into the fallopian tube at ovulation. If the ripe egg is unfertilised, the lining of the uterus is shed in the menstrual period about 14 days after ovulation.

### Timed Intercourse: "natural"

Women: Egg, natural state





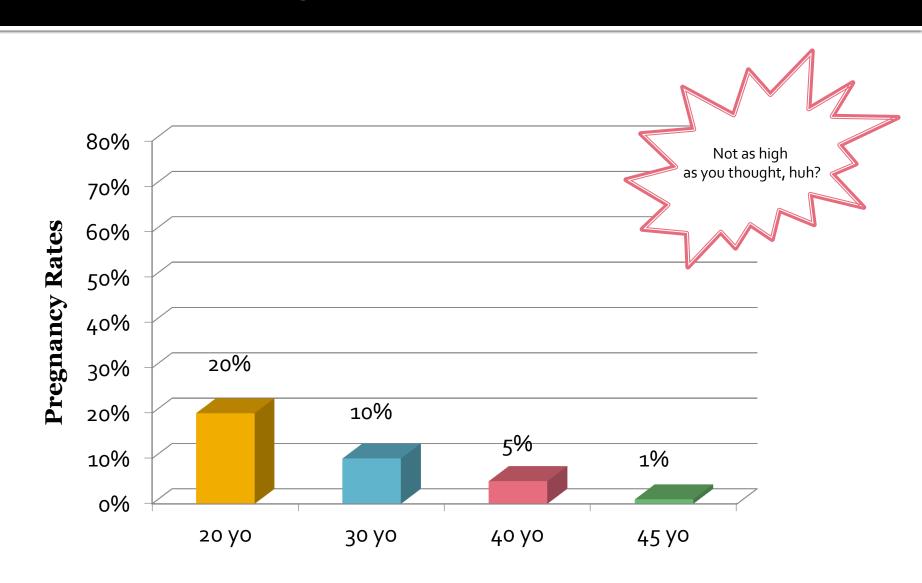
Men: Sperm, natural state

or





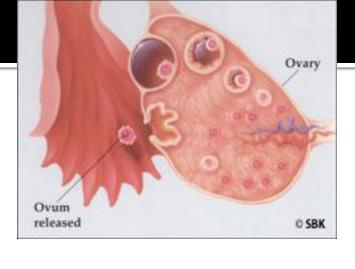
### Outcomes: Timed Intercourse or Sex Pregnancy Rates per Month



### **Ovulation Induction: Eggs**





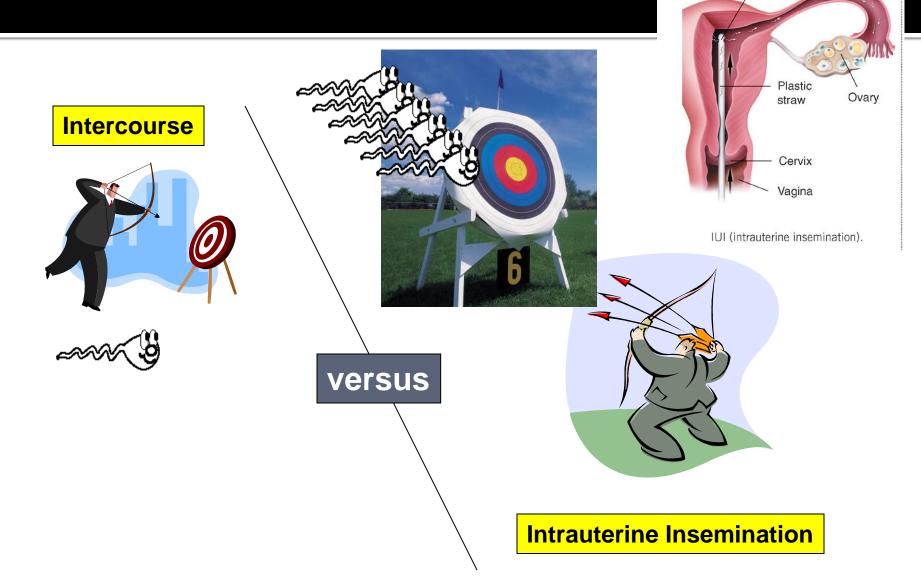


or



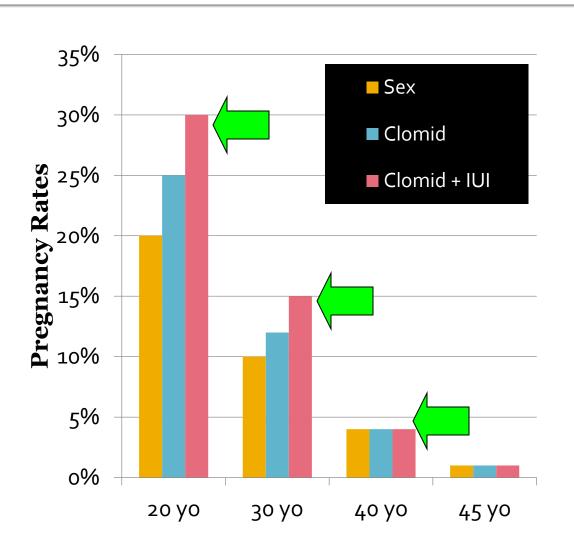


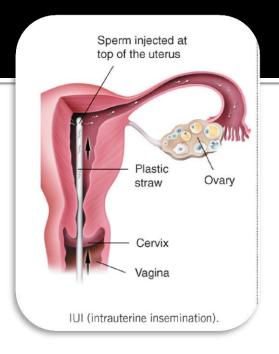
### Intrauterine Insemination (IUI)

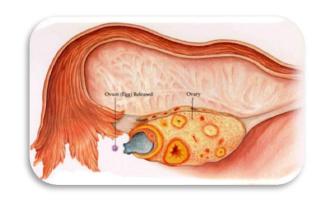


Sperm injected at top of the uterus

# Outcomes: Clomid + IUI Pregnancy Rates per Month

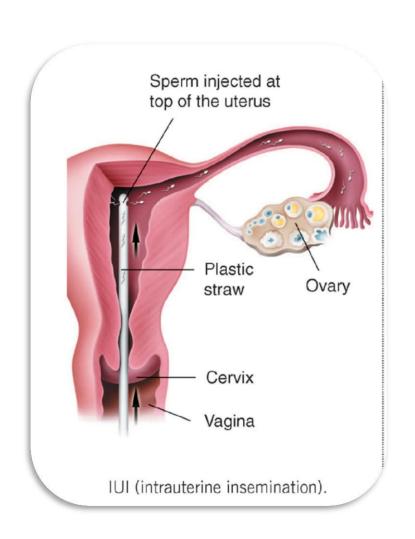






# Should we try IUI first?

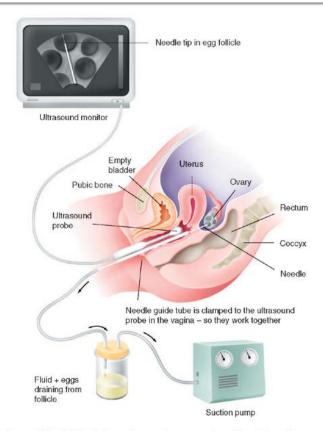
- Yes
- Yes
- Yes
- Maybe???



### Sex and Surgery Cross-section of Abdomen Gas-filled Cavity 20% Your Practice On line **Pregnancy Rates** 15% Sex 10% Surgery 5% 0% 20 yo 30 Ao 40 yo

# Test Tube Babies = IVF





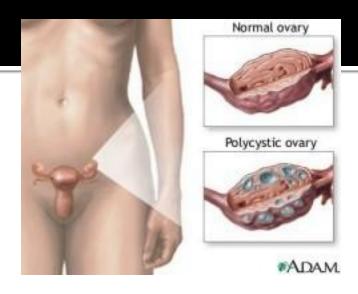
For most IVF and ICSI techniques, the woman's eggs are collected through the vagina using an ultrasound scanner to guide a needle into the follicles and draw off the fluid and eggs.

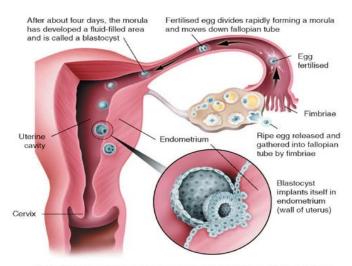
In Vitro means "In Glass" which is another name for "In the Lab"

# IVF

### Goals

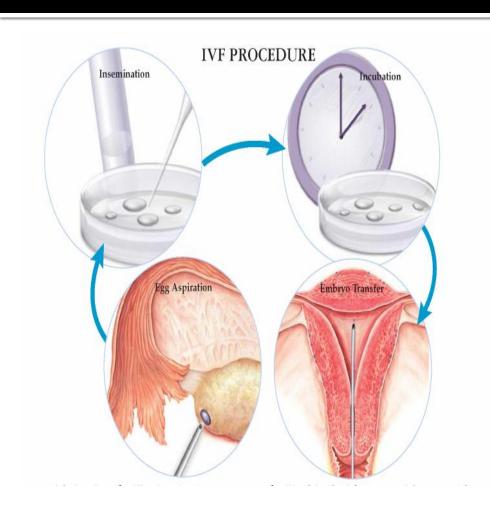
- "Save" eggs
- Grow eggs from both ovaries
- Get <u>largest number</u> of "mature" i.e., fertilizable eggs are developed
- Improve selection process of eggs, sperm, and embryos to enhance the pregnancy rates to levels up to 10 times the natural rates
- "Cure" male factor infertility
- "Cure" tubal factor infertility

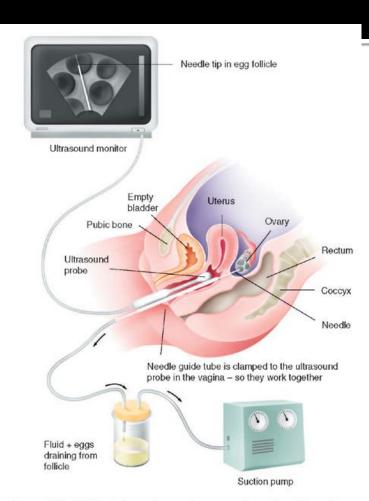




The fertilised egg divides rapidly, first to form a morula and then a blastocyst. After about six to seven days, the blastocyst reaches the uterine cavity where it implants in the endometrium.

## In Vitro Fertilization





For most IVF and ICSI techniques, the woman's eggs are collected through the vagina using an ultrasound scanner to guide a needle into the follicles and draw off the fluid and eggs.

# IVF – In Vitro Fertilization: Eggs



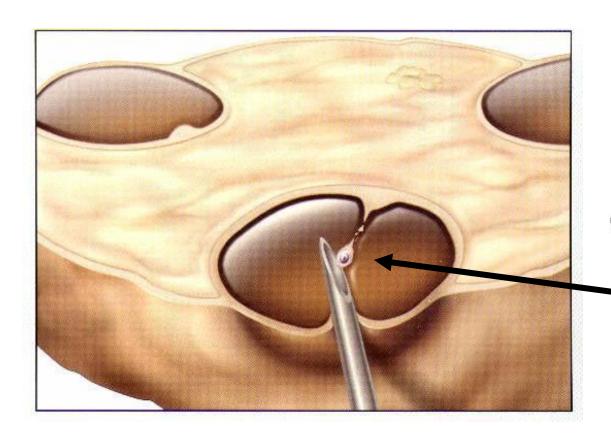
versus

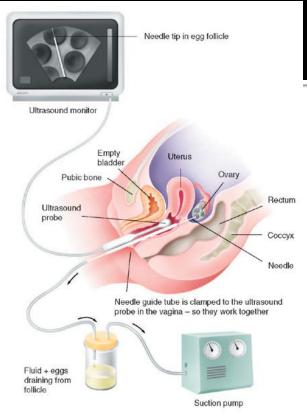
or





# Egg Aspiration

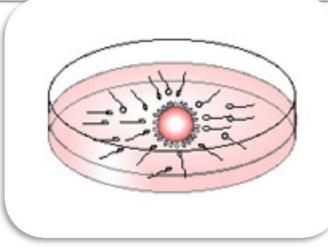




For most IVF and ICSI techniques, the woman's eggs are collected through the vagina using an ultrasound scanner to guide a needle into the follicles and draw off the fluid and eggs.

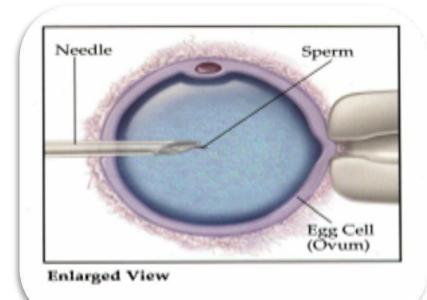
Egg

# IVF - Fertilization



OLD

NEW - ICSI



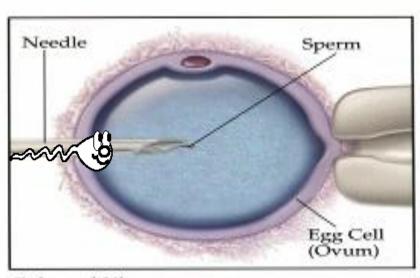
## ICSI: Intracytoplasmic Sperm Injection

### Intercourse



versus





**Enlarged View** 





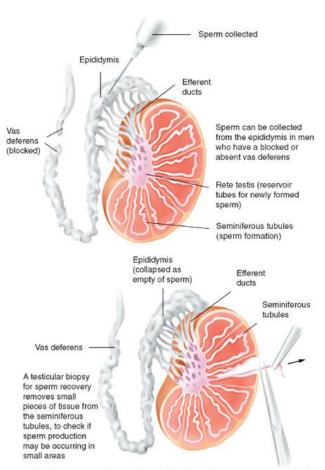
# In Vitro Fertilization - Lab



# What if there are no sperm?

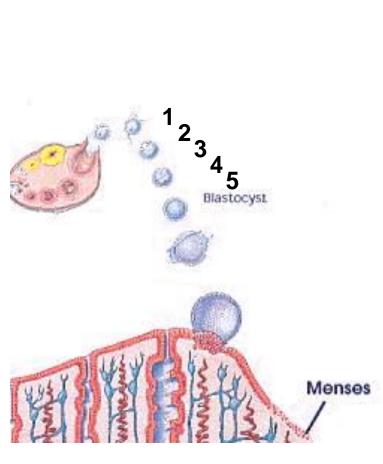
### **Testicular Biopsy**

- Biopsy of Testicular Tissue
  - TESE = Testicular Sperm Extraction
  - PESA, MESA
- Goal
  - Get sperm
    - If Vasectomy then > 98% sperm
    - If born without sperm in ejaculate then > 70% chance of sperm



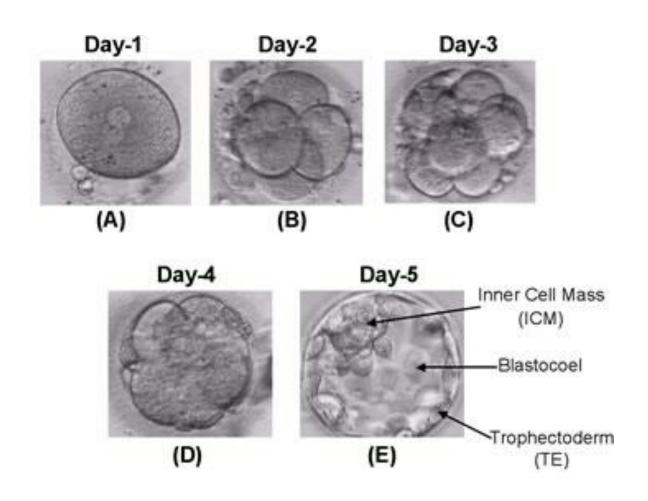
If the man is releasing no sperm in his ejaculate, surgical sperm recovery techniques may be necessary.

# Incubate Fertilized Eggs for 3 to 5 days

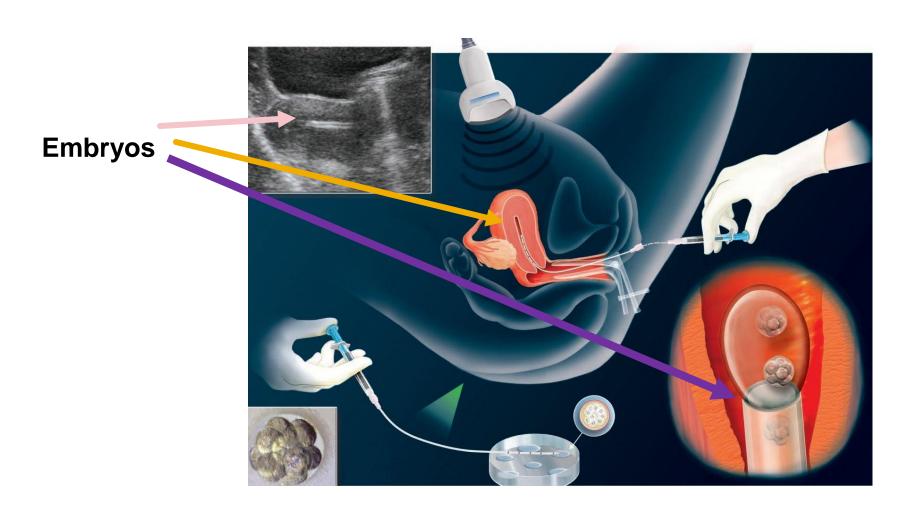




# In Vitro Embryonic Development

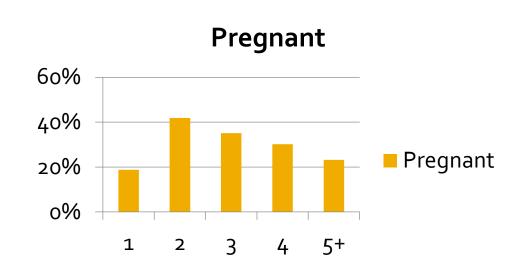


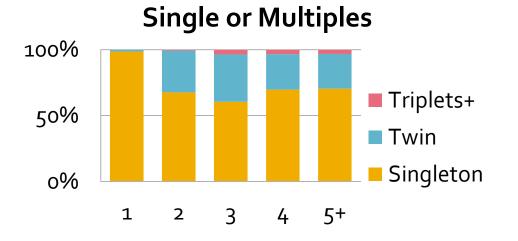
# IVF last step: Embryo Transfer



### How Many Embryos should I ask for?

- 20 yo
  - One or two
    - Each embryo has a 29% chance of "sticking"
- 30 Ao
  - Two or three
    - Each embryo has a 15% chance
- 40 yo
  - Four or more
    - Each embryo has a 4% chance
- > 40
  - ???????





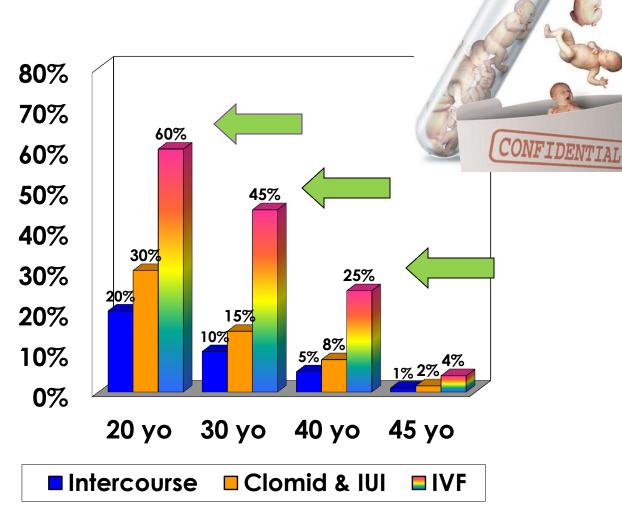
# God has a sense of humor!



### **Outcomes: IVF at RMFC**

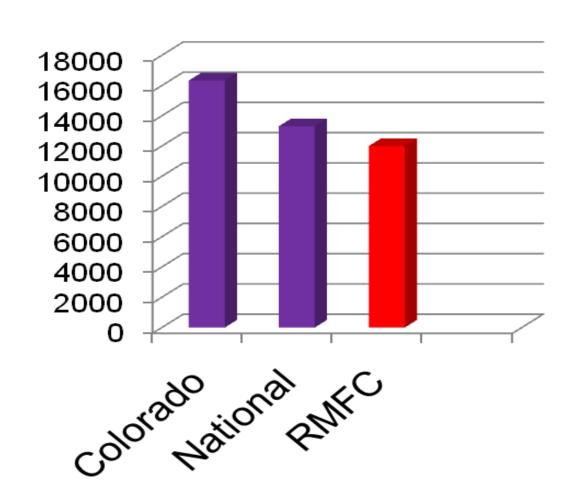
**Pregnancy Rates per Attempt** 







# Average IVF Cost in Colorado is 17,335 + meds, at RMFC \$ 12000 + meds



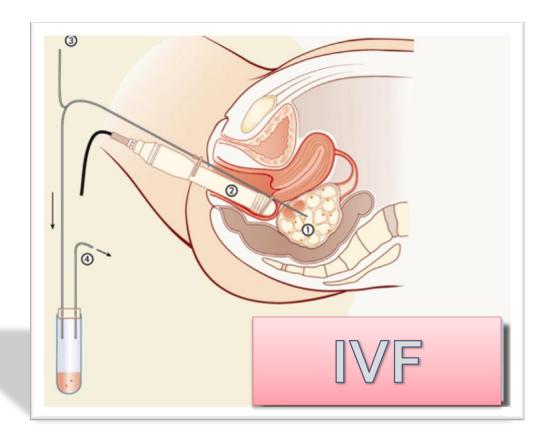
Military
Discount
\$ 500



## Do I have to do IVF?

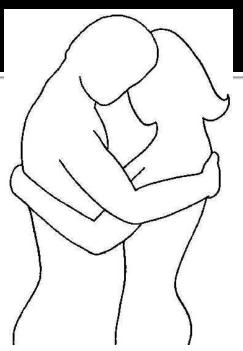
- NO
- NO
- NO
- Maybe!!!





# Summary

- Know the Facts
  - Sperm, Egg, Age, Other causes of infertility
- Know the Tests
  - Semen analysis, SDFA, OAR, HSG
- Know the Treatments
  - TCM, Ovulation Induction, Insemination, Surgery and IVF
- Know the Outcomes
  - What are the realistic expectations for getting pregnant using YOUR egg and YOUR sperm!



# A Gift of Progeny for you!

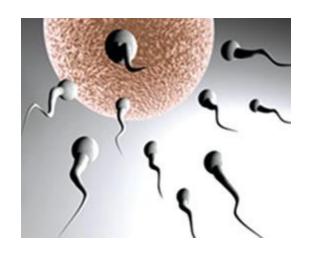


As a way to thank you for coming tonight, RMFC is please to provide you with a coupon for one thousand dollars (\$1,000) off the cost of IVF





# Relax, Enjoy, Breath, Recharge!







# What about Synergy?

Can adding TCM in the form of Acupuncture help me get pregnant?



Paul C Magarelli, M.D., Ph.D.
Board Certified
Reproductive Endocrinologist &
Infertility Specialist



Reproductive Medicine & Fertility Centers 265 Parkside Drive, Suite 200 Colorado Springs, CO 80910



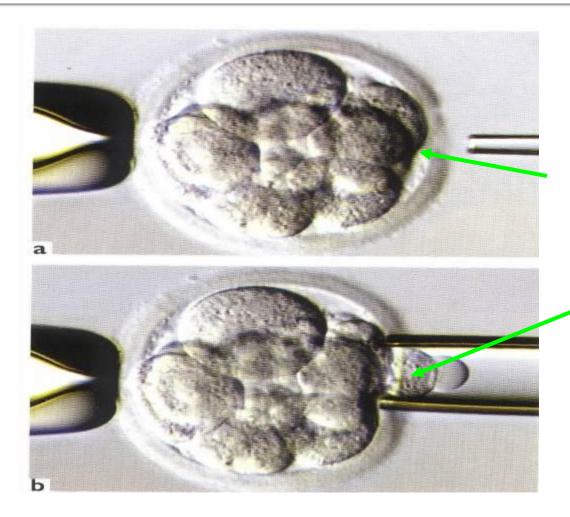
719-475-2229 info@RMFCFertility.com

### **PGD**

### Pre-implantation Genetic Diagnosis

- Diagnoses genetic disease in embryo so that only "normal" embryos are transferred – currently there are over 59 genetic markers that they can test
- Help to "Balance Families"
- Ovarian tissue freezing
- Oocyte freezing

## **Preimplantation Genetic Diagnosis**



Embryo at 8 cell stage Hole made with enzymes Pipette used to suck out Blastomere

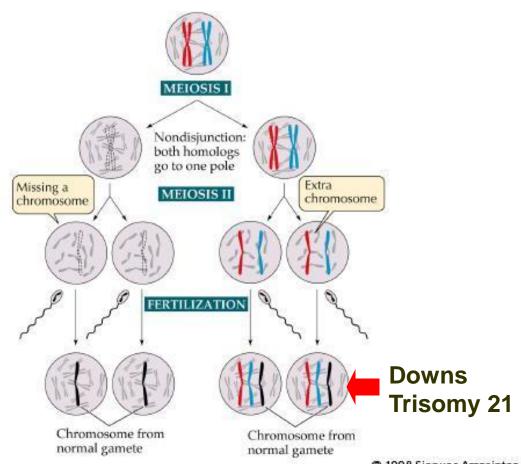
## **Preimplantation Genetic Diagnosis**



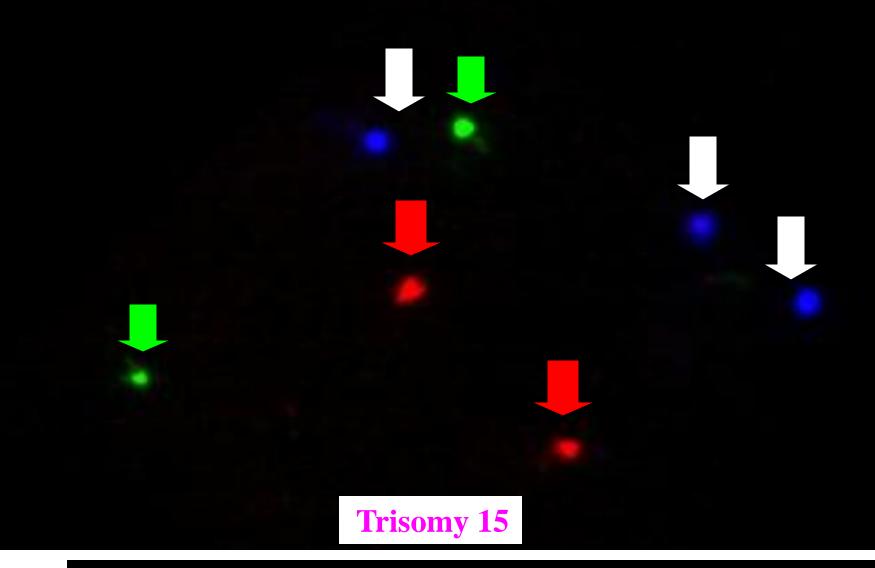


- Blastomere being remove
- Two Blastomere with full chromosomal complement

# Aneuploidy or Woops!

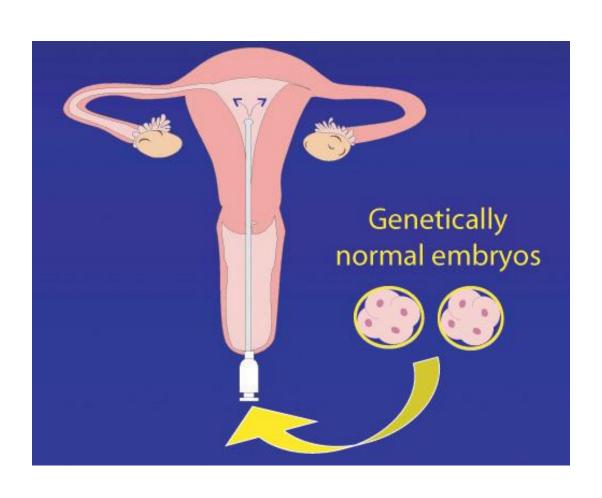


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# PGD last step: Embryo Transfer





# Family Balancing

Corona Institute for Reproductive Medicine & Fertility has joined leading national IVF centers by offering <u>families</u> the opportunity to <u>balance</u> the sex of their future <u>additional</u> offspring.

#### Family Balancing:

- Married
- Have at least one child
- •Desire a child of the less represented sex of children in the family



#### Issues:

- Not for everyone
- May be medically indicated
- •Can reduce risks of genetic disease in offspring

Please consult your religious advisors regarding these treatments. It is certainly not for everyone, but some couples may need our help and we will be here for you.