Hopelessly Infertile: Are There Any Miracles?

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When you absolutely cannot have children, it's called sterility.

When it seems to be taking an awfully long time, but you still hope, that's called infertility.

Infertility is worse.

Katherine Bouton Ms. Magazine April/97

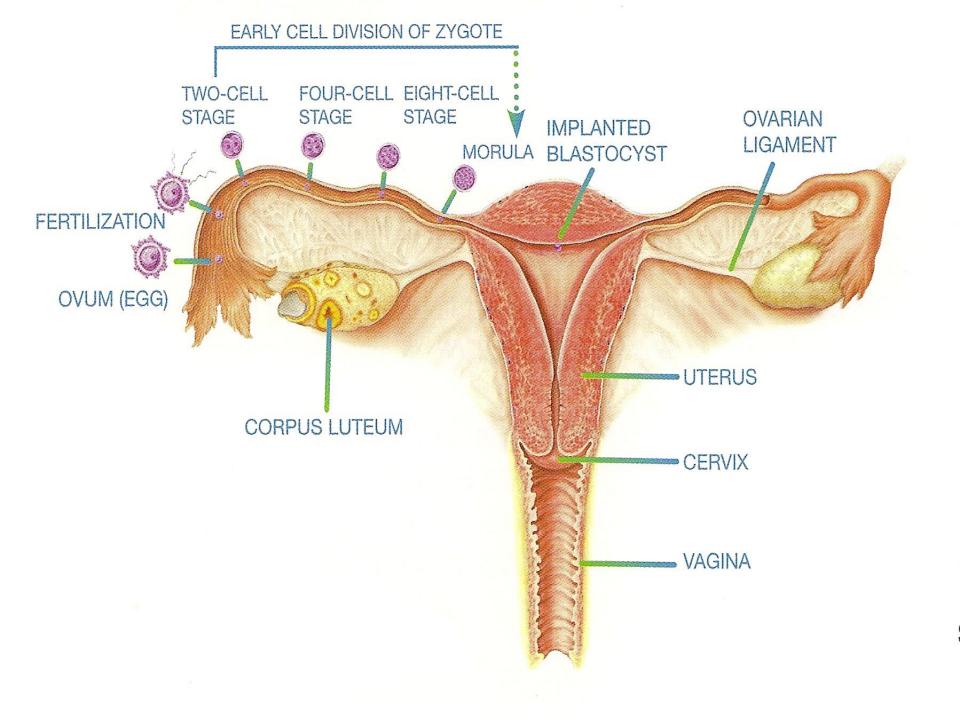


Assisted Reproductive Technology (ART)

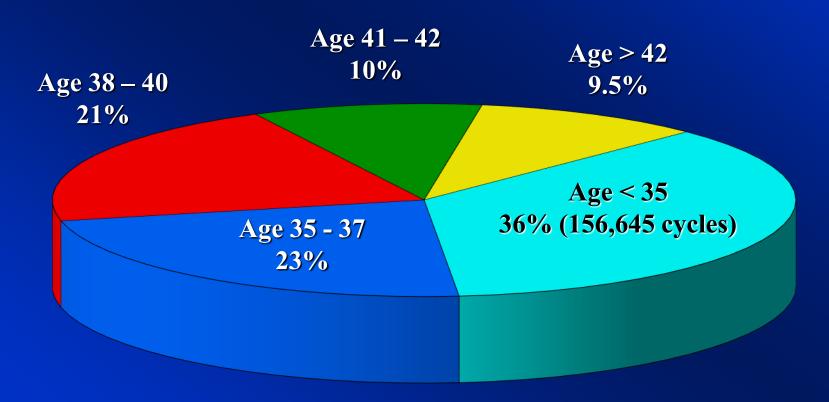
"Techniques that combine egg and sperm outside the female reproductive tract to enhance the probability of pregnancy."

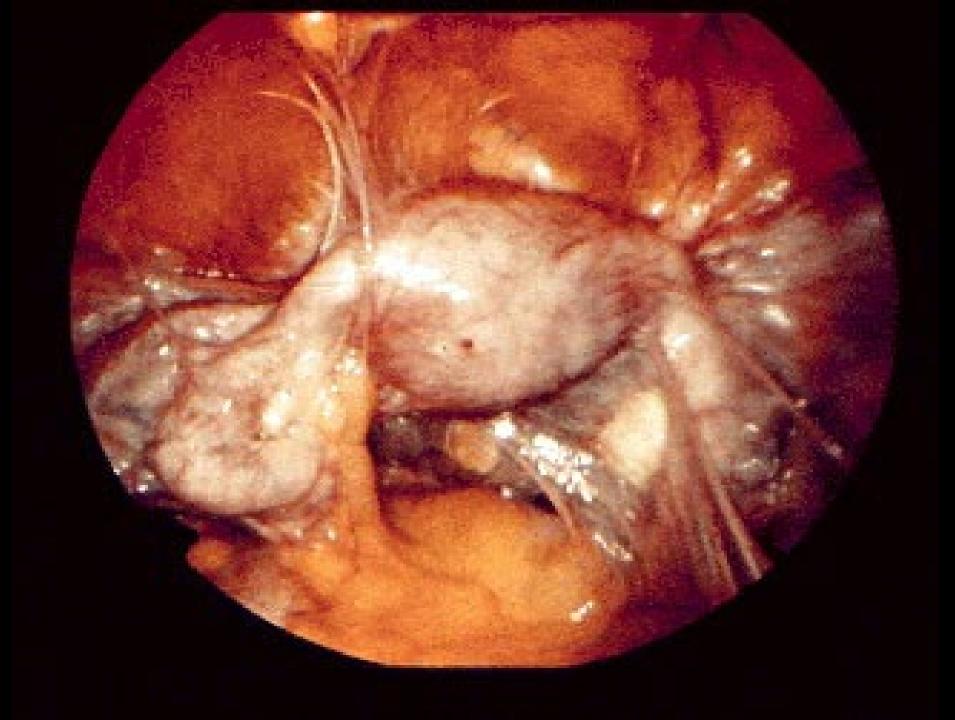
In Vitro Fertilization (IVF) Technique

- Controlled ovarian hyperstimulation (superovulation)
- Oocyte retrieval
- In vitro fertilization
- Embryo culture
- Transcervical embryo transfer

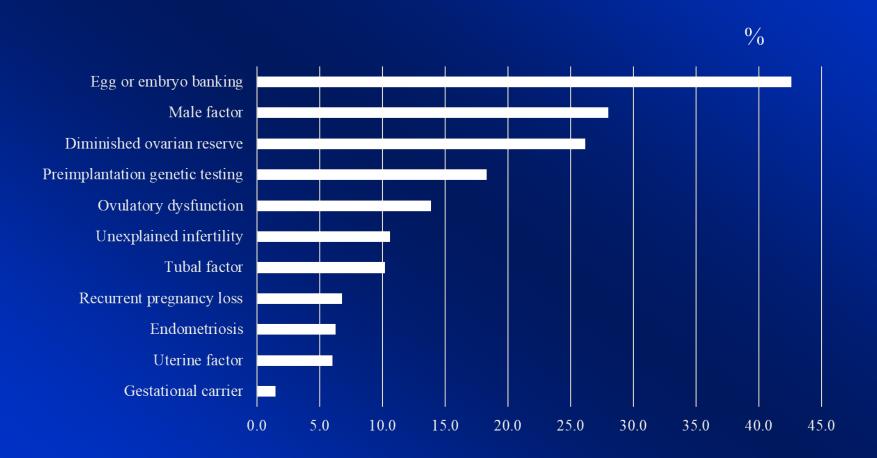


ART Use By Age Group - United States, 2022

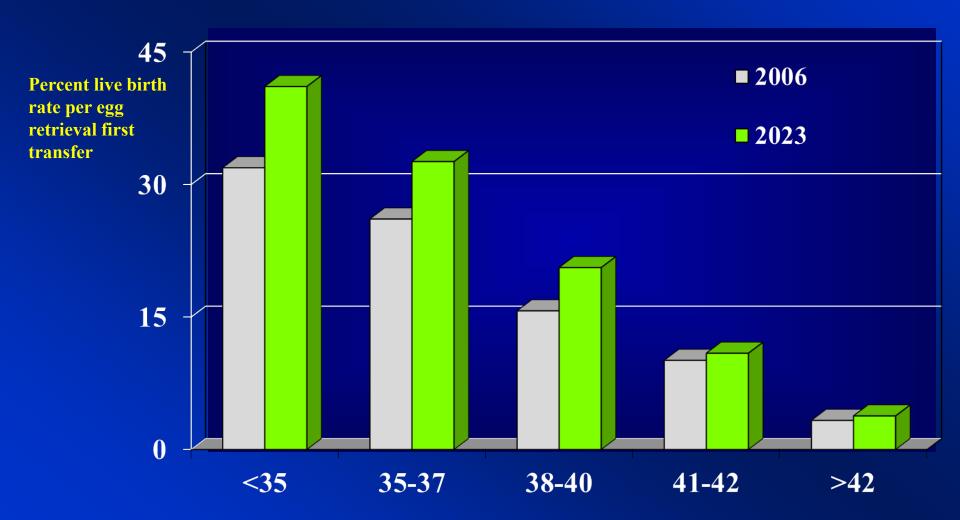




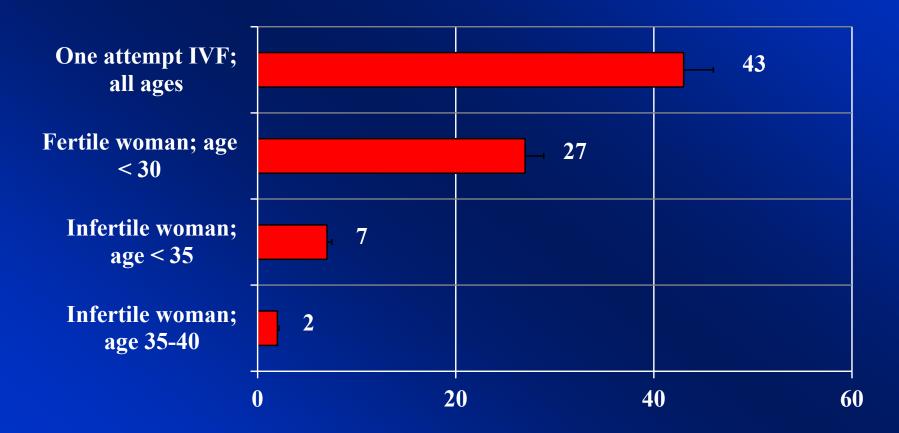
Reasons for using ART – United States, 2022



IVF Live Birth Rates by Age Groups SART 2023



Monthly Fecundity Rate with Live Birth



NHS, 2003 CDC/SART 2022

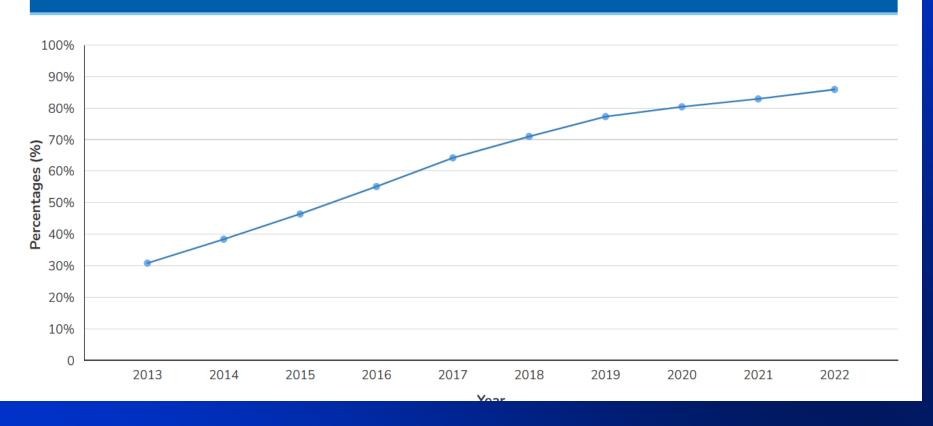
ART Concerns in 2006

- Exposure to superovulation
 - discomfort
 - ovarian hyperstimulation syndrome
 - ?? risk of neoplasm
- Risk of multiple pregnancy
- Risk to fetus
 - ? increased incidence of low birthweight and preterm delivery
- Cost

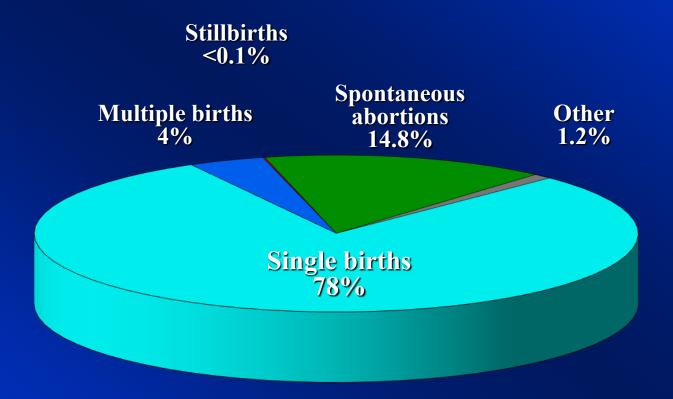


Single Embryo Transfer

Percentage of Embryo Transfer Cycles in Which a Single Embryo Was Transferred, 2013–2022



Outcomes of Pregnancies From ART 2022 CDC



Cryopreservation: Vitrification

7.5% ethylene glycol 7.5% DMSO 20% SSS in M199-H

Equilibration Solution

15% ethylene glycol 15% DMSO 0.5 M sucrose 20% SSS in M199-H

Liquid Nitrogen Plunge

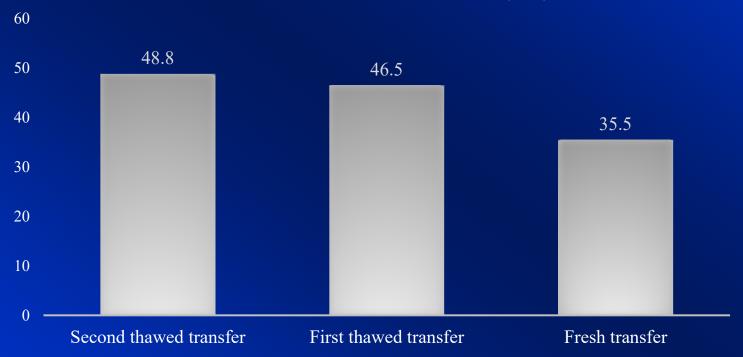
Expose Time 50 to 60 Sec.

Vitrification Solution



Fresh vs Thawed Embryo Transfer

LIVE BIRTH RATE PER RETRIEVAL/THAW (%)

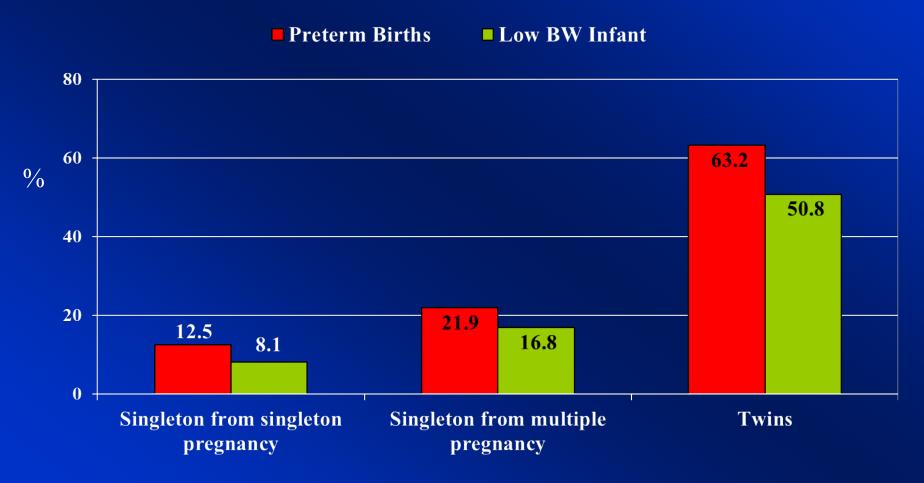


Based on 53,088 egg retrievals: under 35, non-donor oocytes https://www.sartcorsonline.com/EmbryoOutcome/PublicSARTOutcomeTables

Thawed Embryo Transfer

- Only 12% of egg retrievals result in fresh embryo transfer
- Vitrification of embryos preserves embryonic competence
- Controlled ovarian hyperstimulation may adversely affect uterine receptivity
- Allows for widespread pre-implantation genetic testing
- Further reduces risk of ovarian hyperstimulation syndrome

Risk of Pre-term Birth and Low Birthweight in Infants Conceived with ART – CDC 2022



Risk of Congenital Anomalies

	Absolute risk per 1000 ART Births	Odds Ratio
Major birth defects	43	1.5
Hypospadias	7.6	<i>3.1</i>
Epigenetic or Imprinting Disorder Beckwith Wiedemann Angelmann	0.3	17.8
ICSI-Sex chromosome aneuploidy	6	3.0

ART Trends in 2025

- Pre-implantation genetic testing
- Fertility preservation
- Cost

Preimplantation Genetic Testing

- Identifies embryonic genotype
 - Aneuploidy PGT-A
 - Single gene mutation PGT-M
 - Chromosomal rearrangements PGT-SR
 - Sex selection
 - Polygenic conditions?
- Avoids pregnancy termination
- Opportunity for gene therapy

PGT-A

- Implantation rate improved with PGT-A in all age groups
- Miscarriage rate was lower with PGT-A in all age groups
- Livebirth rate improved in age > 37yo

Chang J et al. Fertil Steril 2016;105:394

- Number needed to treat for one additional livebirth:
 - 21 cycles or 35 embryo transfers

The NEW ENGLAND JOURNAL of MEDICINE

ORIGINAL ARTICLE

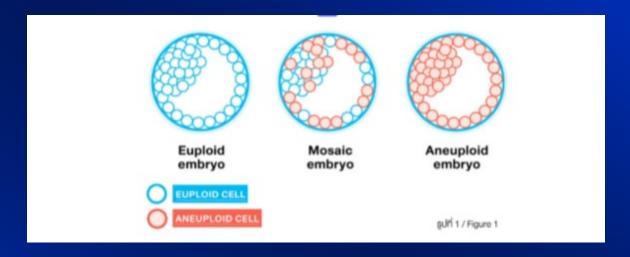
Live Birth with or without Preimplantation Genetic Testing for Aneuploidy

J. Yan, Y. Qin, H. Zhao, Y. Sun, F. Gong, R. Li, X. Sun, X. Ling, H. Li, C. Hao, J. Tan, J. Yang, Y. Zhu, F. Liu, D. Chen, D. Wei, J. Lu, T. Ni, W. Zhou, K. Wu, Y. Gao, Y. Shi, Y. Lu, T. Zhang, W. Wu, X. Ma, H. Ma, J. Fu, J. Zhang, Q. Meng, H. Zhang, R.S. Legro, and Z.-J. Chen

ABSTRACT

- 1212 good-prognosis patients randomly assigned to PGT-A or no PGT-A
- 69.8% of all biopsied embryos were euploid
- No difference in cumulative live birth rate (77.2% vs 81.8%)
- Pregnancy loss was decreased with PGT-A (8.7% vs 12.6%)

Embryonic Mosaicism



- 18 transfers of mosaic embryo in women with no available euploid embryos
- Results in six euploid healthy livebirths and two miscarriages

 Greco E et al. NEJM 2015, 373: 2089
- False positive rate of 0-30% with next gen sequencing (NGS)

Goodrich D et al. J Assist Reprod Genet. 2016; 33:1473-1480



Our Tests

Who We Serve

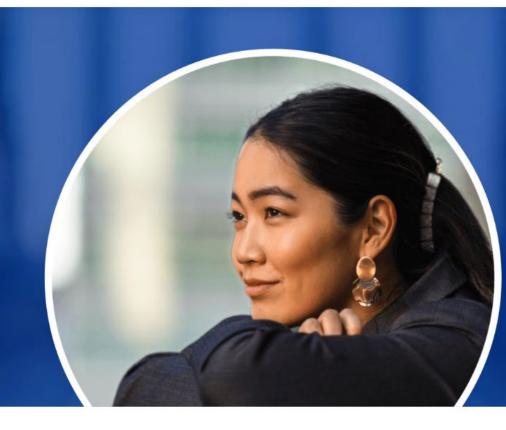
Our Approach

About Us

Resources

Genomic Analysis for Personalized Healthcare

Using whole genome sequencing and state-of-theart bioinformatics, MyOme analyzes a single patient sample to produce comprehensive and clinically actionable reports that allow patients and their providers to make personalized lifestyle, prevention, and treatment choices.



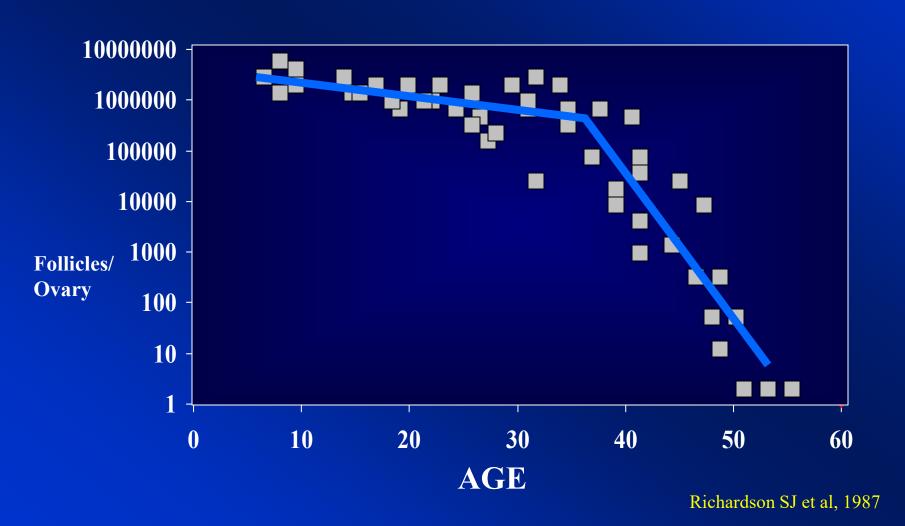
Fertility Preservation Mature Oocyte Cryopreservation

- Prior to gonadotoxic treatment or gonadectomy in reproductive age females
- Excessive available oocytes during IVF
- Unavailable male partner
- Elective fertility preservation (EFP)

American Society of Clinical Oncology (ASCO) Guidelines

- As part of informed consent prior to therapy, oncologists should address the possibility of infertility with patients
- Referrals for interested patients to reproductive specialists
- Discussion at the earliest possible opportunity
- Sperm, oocyte and embryo cryopreservation for centers with expertise are standard practice

Age-Related Changes in Ovarian Reserve



Elective Fertility Preservation(EFP)

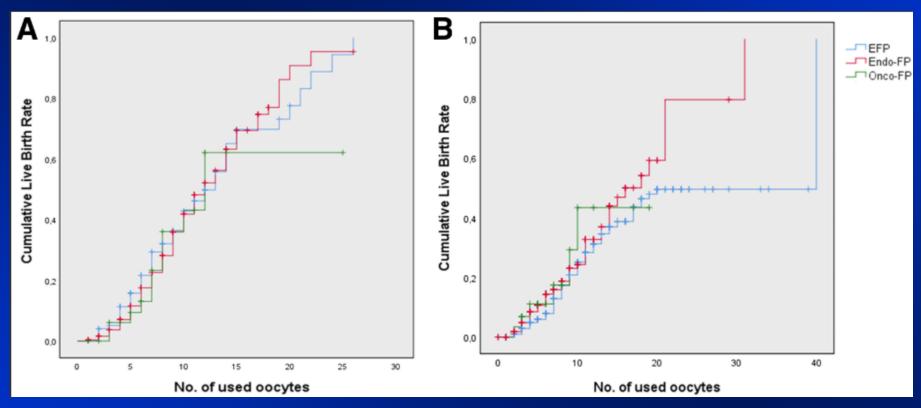
- 1468 patients
 - mean age: 37.2
 - mean no. vitrified oocytes per retrieval: 9.8
- 137 patients returned for IVF
 - Mean age at return: 39.2
 - 85.2% survival rate
 - 27% ongoing preg rate per patient

Cobo A, et al. Fertil Steril 2016:755

Fertility Preservation

 $\overline{\text{Age}} < /= 35$

Age > 35



EFP: elective fertility preservation

Endo-FP: endometriosis Onco-FP: oncologic

