Evaluation and Management of Female and Male Infertility



Your Miracle. Our Mission.

Meike L. Uhler, M.D.

Reproductive Endocrinology & Infertility

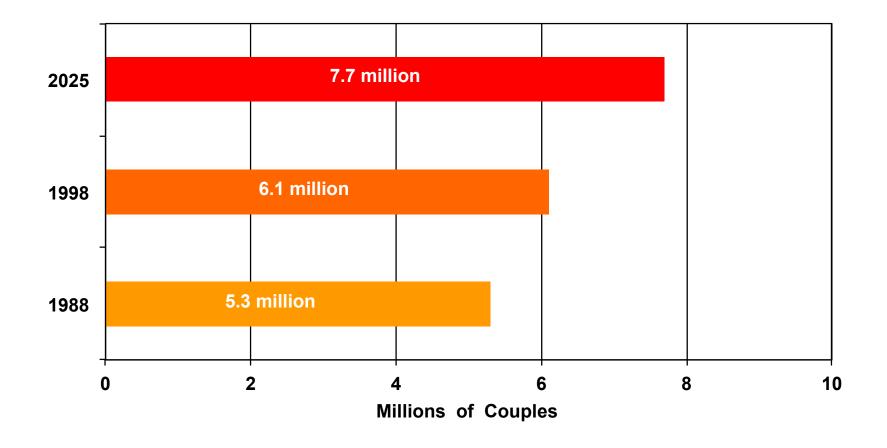
Definition

< 35 years: after 1 year of unprotected intercourse

>35 years: after 6 months of unprotected intercourse



Couples Experiencing Infertility



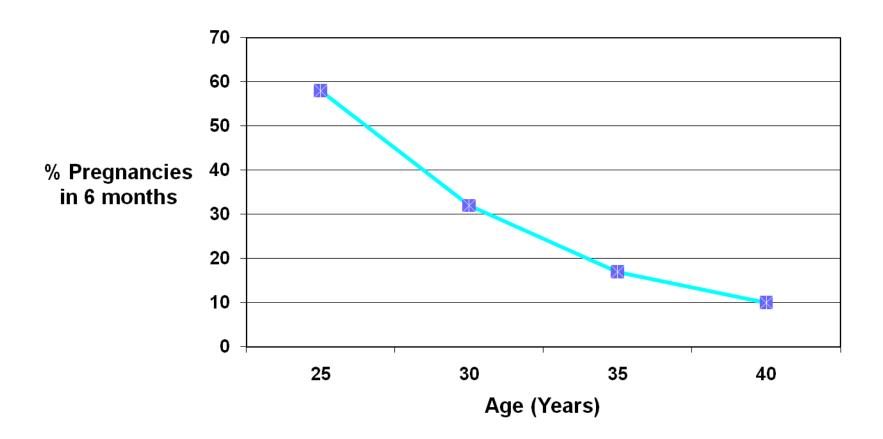


Time Required for Conception

Time of Exposure	Pregnant		
1 month	25%		
3 months	57%		
6 months	72%		
1 year	85%		
2 years	93%		

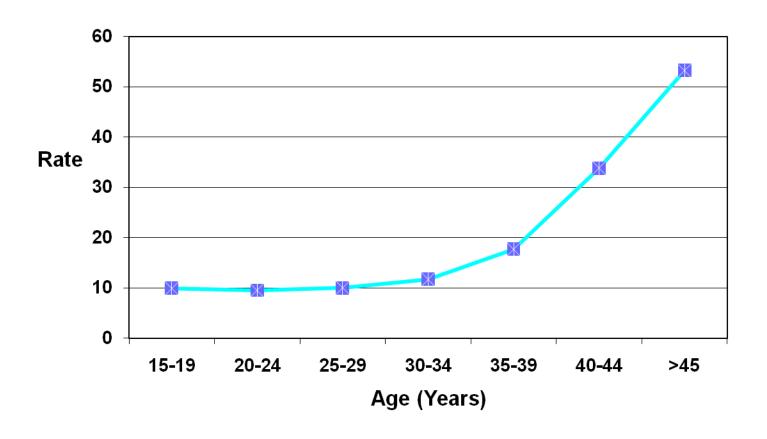


Fertility Decreases with Age





Age and Spontaneous Abortion Rate



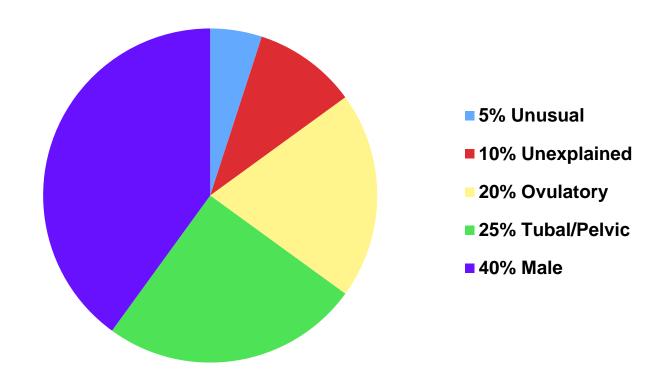


Age and Chromosomal Abnormalities at Birth

	Maternal Age	Maternal Age All Chromosomal Abnormalitie		
	20	1/526		
	30	1/384		
	35	1/204		
	36	1/164		
	37	1/130		
	38	1/103		
	39	1/82		
	40	1/65		
	41	1/51		
	42	1/40		
	43	1/32		
	44	1/25		



Causes of Infertility





History



Prior pregnancy outcome
Detailed menstrual history
STD/PID
PMH, PSH, medications
Social history
FH birth defects, genetic disease



Physical Exam

BMI

Hirsutism, acanthosis nigricans

Thyroid exam

Breast discharge

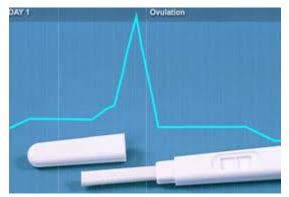
Pelvic exam





Initial Evaluation

D3 FSH, E2, PRL, TSH, LH, AMH Antral follicle count (AFC) Urinary LH and midluteal P4 Hysterosalpingogram or Sonohysterogram Semen Analysis OB prenatal package Comprehensive genetic screening





Additional Diagnostic Tests



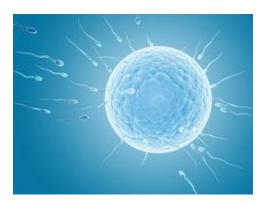
T, DHEAS, 17 OH Progesterone EMB Laparoscopy Hysteroscopy



Male Infertility

Solely responsible in 20% of infertile couples

Contributory in 30-40% of infertile couples





History and Physical



Systemic illnesses

Infection

Previous surgery

Current medication

Substance abuse

Hypospadias

Presence of vas deferens

Testes size

Varicocele



Semen Analysis

Volume **Sperm Concentration** Motility Morphology **WBC**

World Health Organization, 2010 FCI SA Criteria*

 $1.5 - 5 \, \text{mL}$

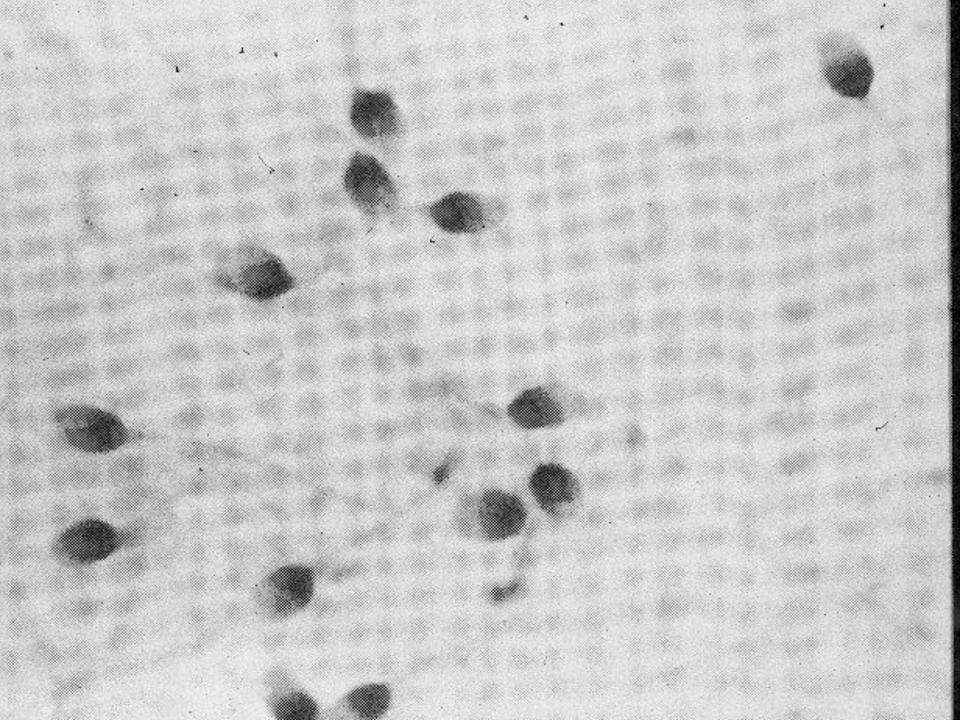
>15 million/mL

>40%

>14%*

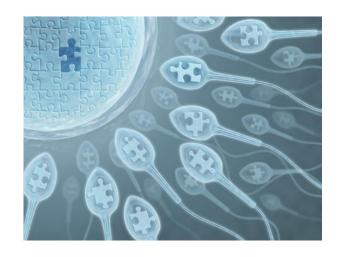
<5/mf



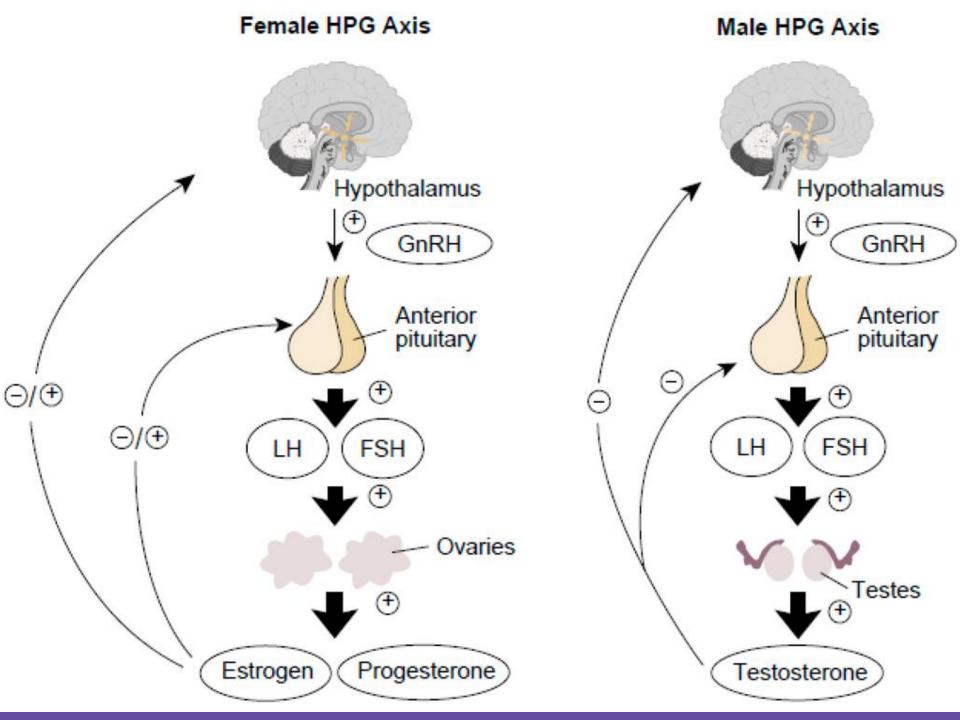


Evaluation

Endocrine testing
Postejaculatory urinalysis
Transrectal or scrotal ultrasound
Specialized tests
Genetic screening







Endocrine Testing

Condition	FSH	LH	Т	PRL
Normal Spermatogenesis	N	N	N	N
Testicular Failure	Н	Н	L	N
Hypogonadotropic/hypogonad	L	L	L	N
Ductal obstruction	N	N	N	N



Specialized Tests

Leukocytes in semen

Direct antisperm antibody test

Sperm viability tests

Postcoital testing

Sperm penetration assay

Computer-aided semen analysis

SCSA

Sperm Aneuploid test





Genetic Screening

Cystic fibrosis gene mutation: Congenital absence of vas deferens

Chromosomal abnormality:

1% of normal men5% of men with oligospermia10-15% of men with azoospermia



Microdeletion of Y-chromosome: AZFa, AZFb, and AZFc



Etiology of Male Infertility

Varicocele	35%
Idiopathic	25%
Primary Testicular Disorders	14%
Genital Tract Obstruction	10%
Sperm Autoimmunity	5%
Hypothalamic-Pituitary Diseases	1%



Clinical Cases

Testosterone supplementation

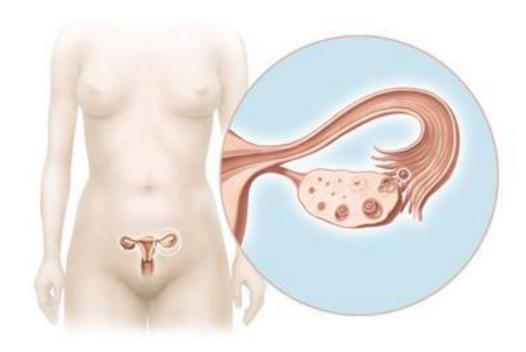
Medical conditions with retrograde ejaculation

Antisperm antibodies

Testosterone



Ovulation Induction





Ovulation Induction: oral medications

Clomiphene citrate

50-150mg x 5 days

Half life=7 days

Hypothalamic receptors

Hot flashed, mood swings



Letrozole

2.5-5mg x 5 days

Half life=2 days

Aromatase inhibitor

Nausea, fatigue





Ovulation Induction: oral medications

Induce ovulation in 80% of patients

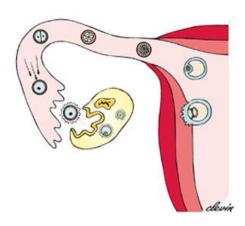
5-10% chance of twins

Rare ovarian hyperstimulation

Possible increased risk of ovarian cancer

May be combined with:

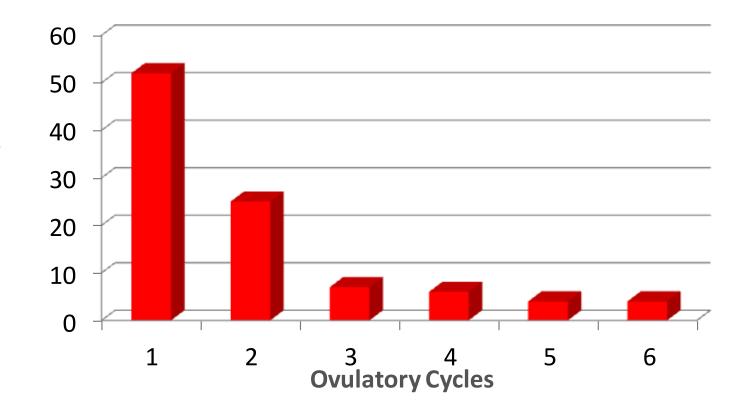
- + Metformin 1500mg
- + Dexamethasone 0.25-0.5mg
- + Bromocriptine 2.5mg
- + Gonadotropins cycle d8-10





Rate of Conception with Clomiphene Citrate

Pregnancy Rate (%)





Ovulation Induction: injectable medications

FSH + LH
GnRH Agonist
GnRH Antagonist
HCG trigger
Progesterone in oil





Ovulation Induction: injectable medications

Induce ovulation in 95% of patients

Breast tenderness, abdominal bloating

Multiple pregnancy rate 20-25%

Ovarian hyperstimulation: mild, moderate, severe

Possible increased risk of ovarian cancer later in life





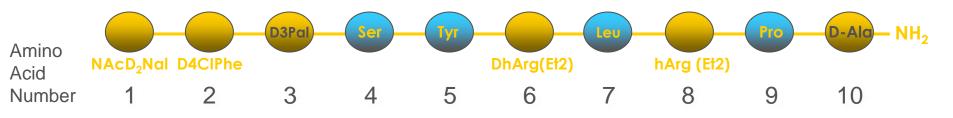
GnRH







AntagonTM





Ovulation Induction: vaginal medications

Progesterone supplementation

Progesterone vaginal suppositories

Crinone 8%

Endometrin 100mg

Estragen supplementation
Estradiol vaginal tablets





Varicocele: Treatment

Palpable varicocele

Couple with documented infertility

Female partner with normal fertility or potentially correctable infertility

One or more abnormal semen parameters

ASRM Practice Bulletin, 2014







Varicocelectomy before assisted reproductive technology: are outcomes improved?

Taylor P. Kohn, M.Phil., Jaden R. Kohn, B.S., and Alexander W. Pastuszak, M.D., Ph.D. b.c

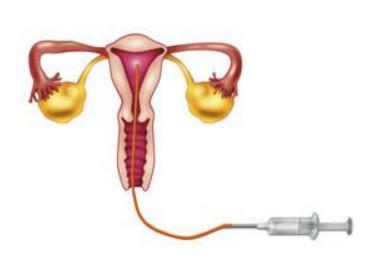
Insufficient evidence surgery improves IUI outcome Possible benefit prior to IVF/ICSI N=6 studies

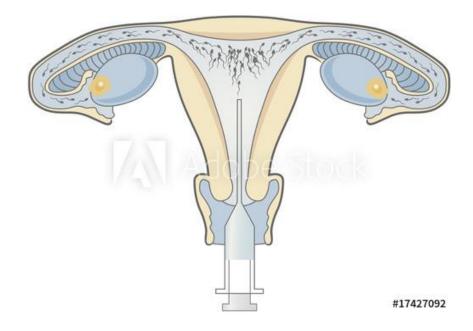
Kohn, et al. Fertil Steril 2017; 108: 385



^a Baylor College of Medicine; ^b Center for Reproductive Medicine, Baylor College of Medicine; and ^c Scott Department of Urology, Baylor College of Medicine, Houston, Texas

Intrauterine Insemination





Percoll Gradient Centrifugation



Treatment Options: Success Rates

Treatment	Success	Rate	(% pe	er cycle)
-----------	---------	------	-------	-----------

Observation 3-4

Natural cycle + IUI 4

Clomiphene + IUI 8-10

Clomiphene + FSH + IUI 10-15

FSH + IUI 15-20

Fertility Centers of Illinois, 2013 N=3033 cycles



Role of tubal surgery in the era of assisted reproductive technology: a committee opinion

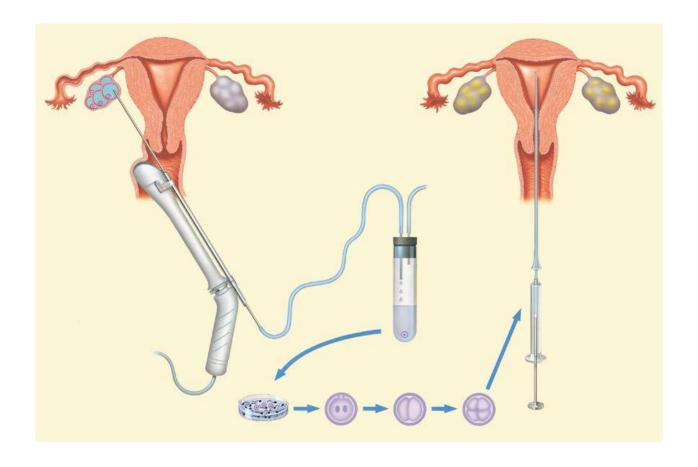
The Practice Committee of the American Society for Reproductive Medicine American Society for Reproductive Medicine, Birmingham, Alabama

Surgical treatment only for mild hydrosalpinges Salpingectomy is recommended prior to IVF

ASRM Practice Committee Document, 2015

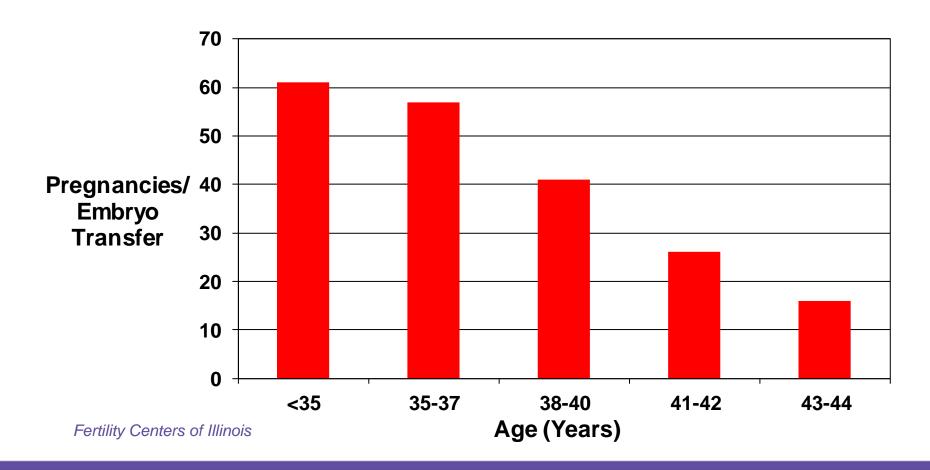


In Vitro Fertilization





IVF Clinical Pregnancy Rates 2017





Preimplantation Genetic Testing

Biopsy: Outer Cell Mass, 5-7 Cells

Error Rate: 1-2%

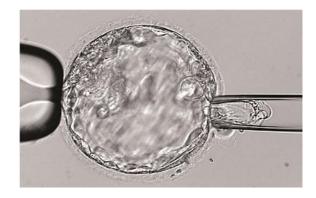
Clinical Error Rate: 5-8%

Platforms: aCGH, SNP Array, NGS, HR NGS

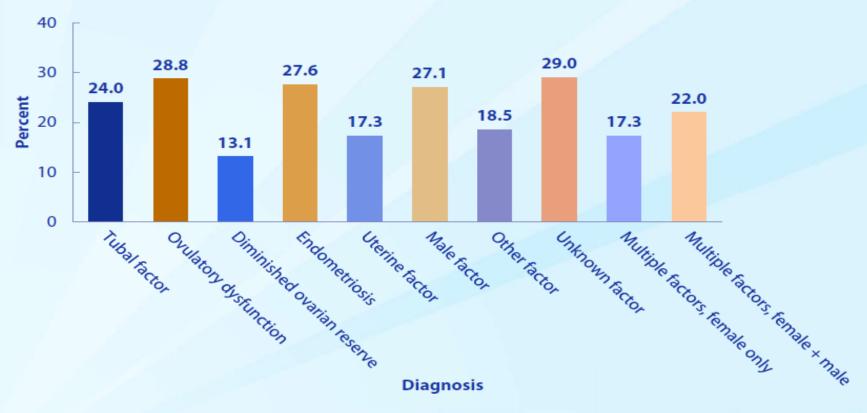
Mosaicism: 20%

Miscarriage Rate: 10%

Munne, S, Fertil Steril 2016, 105-1146



Percentages of ART Cycles Using Fresh Nondonor Eggs or Embryos That Resulted in Live Births, by Type of Infertility Diagnosis, 2015



National Center for Chronic Disease Prevention and Health Promotion

Division of Reproductive Health





Reproductive Options

Donor Oocyte
Donor Sperm
Gestational Surrogacy
Embryo Adoption
Conventional Adoption





Cost of Treatment

Oocyte Vitrification

Procedure and storage \$7,000

Medications \$3,000-5000

IVF

Procedure \$11,500

Medications \$3,000-5000

Freezing and storage \$1,200



PGT

Biopsy \$2,200

Genetic testing \$4,000-5,000



Emotional Aspects of Infertility



Role of stress
Refer to clinical psychologist
Complementary medicine
Support groups
Patient education seminars



Contact



Your Miracle. Our Mission.™

meike.uhler@integramed.com

630-305-7576

630-889-7900

fcionline.com