

### **Male Contraception**

Kirsten M. Vogelsong, PhD

UNDP/UNFPA/WHO/World Bank Special Programme of Research Development and Research Training in Human Reproduction





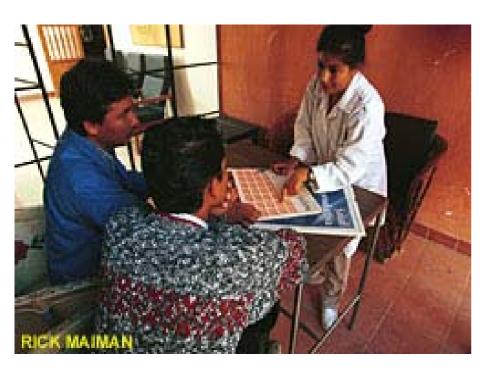
### WHO's Male Reproductive Health Research

- Contraceptive research and development
- Targeted basic science
- Social science
- Men's role in reproductive health
- Norms and Standards in Andrology
- Training workshops





### Why Men in Family Planning?



- Conference on Population and Development, 1994
- Disproportionate reproductive disease burden
- Shared responsibility and gender equity



### Distribution of Contracepitve Use Prevalence

#### World wide contraceptive use

Contraceptive	No. of users (Millions)	Users (%)	First year failure rate (%) - Typical use
Total users <u>Modern methods</u>	550	53	
Female sterilization	163	30	0.5
IUD	110	20	0.8
Oral contraceptives	91	17	5.0
Condom	49	9	14.0
Male sterilization	45	8	0.15
Injectables/implants	11	2	0.3
Vaginal barriers	8	1	20.0
<b>Traditional methods</b>			
Withdrawal	41	7	19.0
Rhythm	32	6	25.0 UNFPA, 1994





# Male Contraception Research and Development

- Use of existing male methods is low, with regional and country differences
- Men are aware of family planning methods
- Men approve of the use of family planning
- Low levels of use may be related to the negative characteristics of existing methods
- Example: In a study conducted in Fiji, Iran, India and Korea, men considered a male pill or injection to be more acceptable than vasectomy





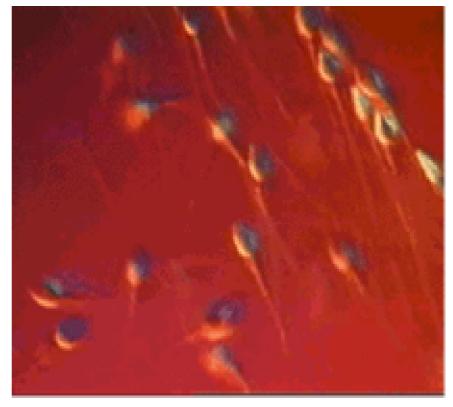
### The Ideal Male Contraceptive

- Safe no harmful side effects
- Effective it works!
- Acceptable to men and their partners
- Affordable to donors, programs, and potential users



### Approaches to Male Contraception: Targeting the sperm

- Inhibit production
- Interrupt transport
- Block deposition
- Disrupt function
- Prevent fertilization







### Inhibiting Sperm Production Hormonal Contraception

T Enanthate Androgen alone

T Undecanoate

T Buciclate

**Pellets** 

**Progestin + Androgen** Norplant

**DMPA** 

**Norethisterone Enanthate** 

**GnRH Agonists** 

**Antagonists** 

Vaccines

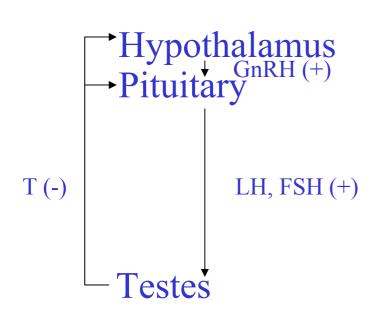
**Antagonists** 

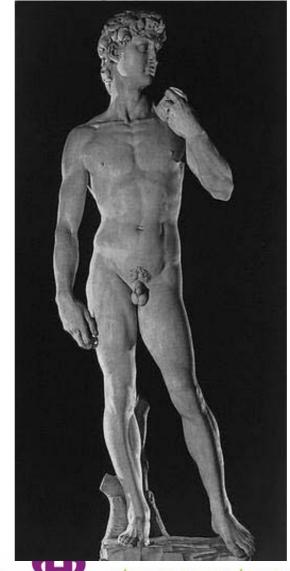
**Vaccines** 

FSH



Hormonal Male Contraception



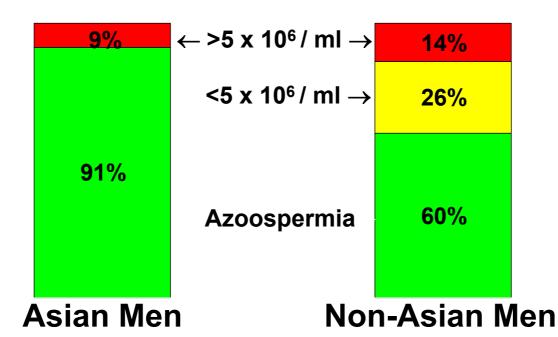




- 1990: 200 mg testosterone enanthate/week will reduce sperm production in some men
- Sperm concentrations consistently below 1 million/ml result in few or zero pregnancies
- All men do not fully suppress
- Requirement for weekly injections and high T concentrations



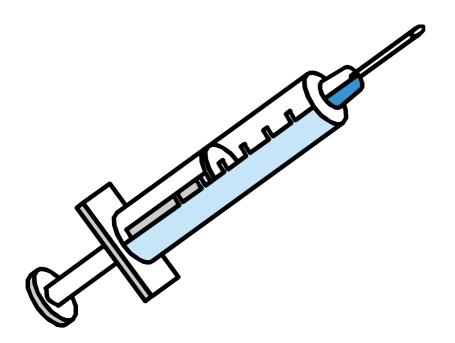




Sperm concentrations following weekly inj. 200 mg T-enanthate







#### Testosterone Enanthate

- Extensive clinical experience
- "Burst" effect
- Short acting
- Weekly injections
- High levels testosterone





# Hormonal Approaches to Male Contraception

#### Androgen alone

#### Testosterone Undecanoate

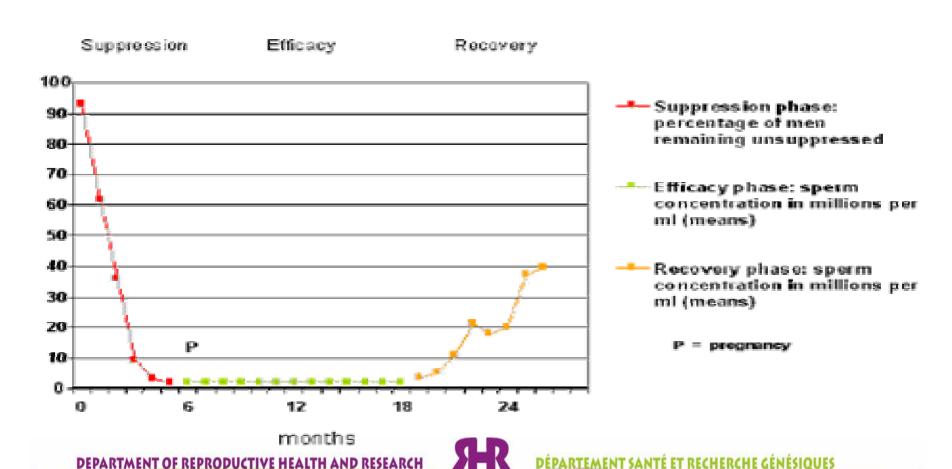
- Oral or injectable
- Longer release profile
- 4-8 week injection intervals may be adequate
- Maintains testosterone in physiological range
- Large dose required

#### Testosterone Buciclate

- No "burst" effect
- Long-acting injectable
- Injections at 3-4 month intervals may be adequate
- High dose required









# Hormonal Approaches to Male Contraception Androgen with Progestin

- More rapid and effective sperm suppression
- Effective in diverse populations
- Reduced overall drug load
- Physiological testosterone levels
- Requires a 2 drug regimen
- Drugs may have different routes or frequencies of administration





## Hormonal Approaches to Male Contraception

### Androgen with Progestin

Progestagen	Androgen		% Oligozoo spermic	Reference
DMPA 250 mg every 6 weeks	19 NT (200 mg every week x 6/7 weeks, then 200 mg/3 or 4 weeks).	67 (W) 98 (A)	92 (W) 99 (A)	Knuth et al (1987)
	TE (200 mg(IM every week x 6/7 weeks, then 200 mg/4 weeks)	59 (W) 96 (A)	91 (W) 96 (A)	WHO (1993)
DMPA 300 mg	T implant (800 mg)	90 (W)	100 (W)	Handelsman et al (1996)





## Hormonal Approaches to Male Contraception

### Androgen with Progestin

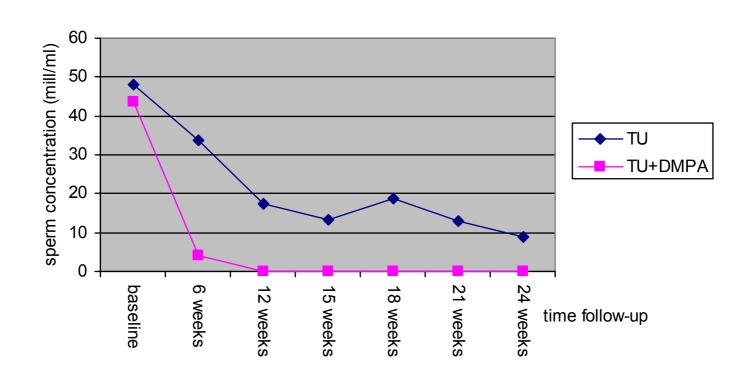
Progestagen	Androgen	_	% Oligozoo- spermic	Reference
Levonorgestrel (o	ral)			
500 μg/day	TE (100 mg/week IM)	67 (W)	94 (W)	Bebb et al (1996)
250 μg/day	TE (100 mg/week IM)	78 (W)	89 (W)	Anawalt et al (1997)
125 μg/day	TE (100 mg/week IM)	61 (W)	94 (W)	(1007)
Desogestrel (oral)				
300 μg/day	TE (100 mg/week IM)	81 (W)	94 (W)	Wu et al (1998)
150 μg/day	TE (50 mg/week IM)	73 (W)	100 (W)	

W=White, A=Asian, DMPA=depotmedroxyprogesterone acetate, TE=testosterone enanthate 19 NT= 19 nortestosterone hexyloxyplenylpropionate





# Hormonal Approaches to Male Contraception Androgen with Progestin







## Hormonal Approaches to Male Contraception

#### Other Approaches

- Androgen with anti-androgen (*cyproterone* acetate)
  - Progestin with anti-androgen properties
  - May block the activity of any residual T in the testis
- Androgen with GnRH Analogue
  - Effective suppression of gonadotrophins
  - High cost; frequent application





### Interrupting Sperm Transport

#### Vasectomy/Sterilization

### Worldwide, 42-60 million men (5-8% of married couples) rely on vasectomy

- New Zealand 23%
- United States 11%
- The Netherlands 11%
- South Korea 11%
- Australia 10%

- China 8%
- India 7%
- Nepal 5%
- Thailand 5.3%
- Sri Lanka 4.6%
- Brazil 1.6%

All other developing countries have <1% prevalence of use





### Interrupting Sperm Transport

### Vasectomy/Sterilization

Contraceptive Prevalence by Methods Used (China 1982, 1988, and 1992 National Family Planning Surveys)

Method		Year of Surv	ey
	1982	1988	1992
	(n=172,788)	(n=406,387)	(n=73,946)
Sterilization	24.5%	35.0%	46.1%
Female	17.6%	27.2%	35.9%
Male	6.9%	<b>7.8%</b>	<b>10.2%</b>
<u>IUD</u>	34.9%	29.5%	33.1%
Others*	10.1%	6.6%	5.2%
Non-users	30.5%	28.9%	15.4%
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<sup>\*</sup> The oral pill, condom, spermicide and other methods.





### Interrupting Sperm Transport Vasectomy/Sterilization

#### Conventional vasectomy

- highly effective and safe
- incision required
- permanent

#### Percutaneous vas occlusion

- many compounds evaluated
- lower efficacy rates
- some additional complications

#### No-scalpel vasectomy

- highly effective
- Somewhat more acceptable
- lower complication rates





# Methods of Vasectomy Success of Reversal

Method	Follow-up (no and %)	Sperm (no and %)	Normal (no and %)	Pregnancy (no and %)
No-scalpel	19/23	16/19	13/19	15/19
Vasectomy	(82.6)	(84.2)	(68.4)	(78.9)
Chemical Vas occlusion	26/31	18/26	12/26	13/26
	(83.9)	(69.2)	(46.2)	(50.0)
MPU	31/34	10/31	10/31	9/31
Vas occlusion	(91.2)	(32.3)	(32.3)	(29.0)





### Blocking sperm deposition





## Blocking sperm deposition Male Condoms

- Condoms are effective at preventing pregnancy and STI/ HIV
- Condom use is low even in countries with high prevalence of HIV/AIDS
- How can we increase condom use?







## Blocking sperm deposition Male Condoms

#### Condom studies

- Randomized comparative studies of "standard" and "new" condoms
  - Acceptability and preference
  - Contraceptive efficacy
  - Prevention of STI
- Reasons for use and non-use of condoms



# Disrupting Sperm Function and Preventing Fertilization

• Targeted basic science research on testicular, epididymal or vas approaches:

- Some promising targets:
  - functional development, i.e. motility
  - structural development, i.e. organelles
  - structure and function, i.e. membrane integrity
     and intracellular pathways

### Acceptability/Sociobehavioral

- Current use of male methods
- Preferences for new methods
- Characteristics of new methods
- Continuation and discontinuation of trial

- Effects on mood
- Effects on behavior
- Effects on cognition
- Partner's views on mood and behavior



**Studies** 



### Acceptability/Sociobehavioral Studies

#### Reports from 25 Swedish men participating in TE trial

#### **Expectations**

- Freedom and security
- Problems with female methods
- Desire for more satisfying sex life
- Need for male control
- Fear of negative side effects

#### Satisfaction

- Greater freedom
- More ease in sex life
- Would recommend method to others
- Trouble with injections
- Fear of problems with aggressiveness
- Dermatological problems



# Acceptability/Sociobehavioral Studies

	Very important	Somewhat important	Not important
Men should share responsibility for contraception	41.2	51.0	7.8
Contributing to solving the population problem	41.6	48.7	9.7
I felt I was doing a good thing for my country	36.7	52.9	7.9
I like to be involved in new things	25.0	56.8	18.2
I felt pride in contributing to scientific advancement	26.9	51.6	21.4
Pioneer of a new method of contraception	24.4	46.1	29.5
My wife wanted me to take responsibility	23.1	44.8	32.1
I joined for getting the financial compensation	12.7	28.6	58.8



# Acceptability/Sociobehavioral Studies

	Month 4	Month 8
	%	%
Reasons for perceived inconvenience	(n = 78)	(n = 117)
Have to come to clinic	23.1	9.3
Once a month too frequent	70.5	76.3
Wait at the clinic	1.3	5.1
Other	5.1	9.3
Total	100.0	100.0
Reasons for dissatisfaction	(n = 87)	(n = 117)
Side effect	11.5	6.0
Inconvenience	54.0	48.7
Injection pain	21.8	12.0
Others	12.6	33.3
Total	100.0	100.0





### Men's Role in Reproductive Health

#### Men can:

- Inhibit access to and use of FP
- Expose women and themselves to disease including HIV
- Act as barriers to women's reproductive health

OR

Facilitate use of contraception

OR

 Protect themselves and their partners from infection

OR

 Act as partners in promoting reproductive rights and care for all

