# Caring for the Transgender Patient

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#### No Financial Disclosures

Medications Discussed are Off-Labeled Use

I am NOT final authority on Transgender Health

### Objectives

Review Transgender Terminology & Demographics

Discuss Transgender Health Disparities

Develop Methods to Foster a Gender-Affirming Environment

Brief Review of Medical & Surgical Options for Transitioning

Effects of Hormone Therapy on Labs & Imaging



### World Gender Customs<sup>1</sup>



### Two Spirit

Two Spirit people held positions of great respect

- Medicine Men/Women
- Shamans/Visionaries/Mystics
- Keepers of oral traditions
- Cooks
- Matchmakers/Marriage counselors
- Singers/Artists

Osh-Tisch<sup>2</sup> (Finds Them and Kills Them) - pictured on the left

- Badé of the Crow Tribe
- Afforded distinctive social and ceremonial status
- Served as a scout and earned a reputation for bravery after the Battle of the Rosebud, June 17, 1876.





# Terminology<sup>3</sup>

#### **Transgender**

A person whose gender identity and assigned sex at birth do not align

- Transgender Man: Female-To-Male
- Transgender Woman: Male-To-Female

#### **Non-binary**

A person whose gender identity falls outside of the traditional gender binary structure

#### **Cisgender**

A person whose gender identity and assigned sex at birth correspond (i.e., a person who is not transgender).

#### **Gender Fluid**

A person whose gender identity is not fixed.

# Terminology<sup>3</sup>

### Intersex/Differences of Sexual Development (DSD)

Group of rare conditions where the reproductive organs and genitals do not develop as expected secondary to variations in chromosomal, hormonal, gonadal or anatomical development

#### **Transsexual**

Sometimes used in medical literature or by some transgender people to describe those who have transitioned through medical interventions

#### **Transition/Gender Affirmation Process**

Period when a person makes social, legal, and/or medical changes

- Gender Affirming Hormone Therapy
- Gender Affirming Surgery

#### **Gender Dysphoria**

Experienced by some whose gender identity does not correspond with their assigned sex at birth. Manifests itself as clinically significant distress or impairment

# Terminology<sup>3</sup>

#### **Problematic**

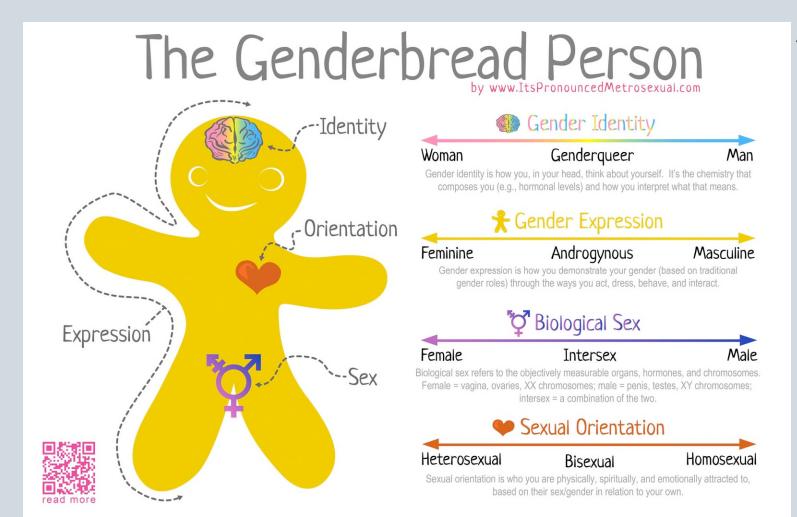
- Transgenders, A transgender
- Transgendered
- Transgenderism
- Sex change, Pre-op, Post-op
- Biologically male/female, genetically male/female, born a man/woman
- Passing/Stealth

#### **Preferred**

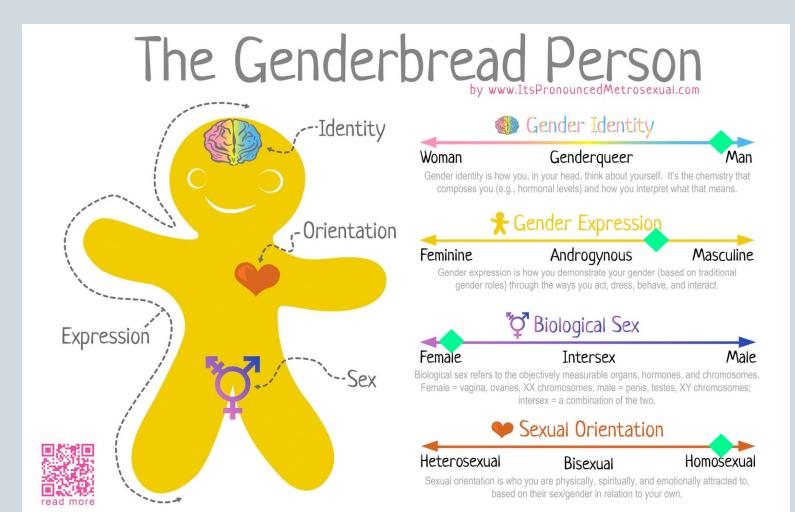
- Transgender people, A transgender person
- Transgender
- Being Transgender
- Transition
- Assigned male/female at birth, Designated male/female at birth
- Visibly/Not Visibly Transgender

Defamatory: Tranny, She-male, He/She, It, Shim









#### Gender Identity Development<sup>5</sup>

#### Establishment of gender identity varies

For some children, gender identity may be fairly firm when they are as young as two or three years. For others it may be fluid until adolescence and occasionally later

#### Around 2-years-old

- Children become conscious of the physical differences between sexes

#### Around 3-years-old

- Most children are easily able to label themselves as either a boy or a girl

#### Around 4-years-old

- Most children have a stable sense of their gender identity
- Same time of life children learn gender role behavior

### US Transgender Demographics

#### 2017 - Williams Institute<sup>6</sup>

State	Age										
	13 to 17		18 to 24		25 to 64		65 and older		All Adults (ages 18+)		
	Number	Percent	Number	Percent	Number	Percent	Number	Percent	Number	Percent	
United States	149,750	0.73%	205,850	0.66%	967,100	0.58%	217,050	0.50%	1,397,150	0.58%	
Missouri	2,500	0.63%	3,600	0.60%	17,000	0.54%	4,400	0.50%	25,050	0.54%	

Epilepsy 0.66%<sup>7</sup>

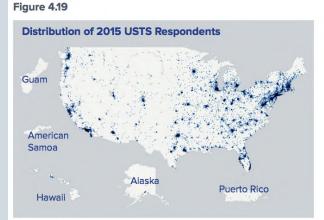
Down's Syndrome 0.15%8

Type 1 DM 0.34%<sup>9</sup>



# U.S. Transgender Survey (2015)<sup>10</sup>

- 27,715 respondents from all 50 states, D.C, American Samoa, Guam, Puerto Rico and US military bases overseas
- Anonymous online survey for transgender adults (18 and older)
  - English and Spanish
- Explored categories related to education, employment, family life, health, housing and interactions with the criminal justice system





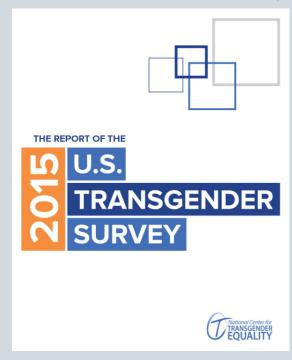
Each dot on the maps represents the number of people in a zip code. Every dot corresponds to at least one person, and the size of each dot increases in accordance with the number of people in each zip code.

# Health Disparities<sup>10</sup>

Disparities stem from structural/legal factors, social discrimination, and lack of culturally competent

health care

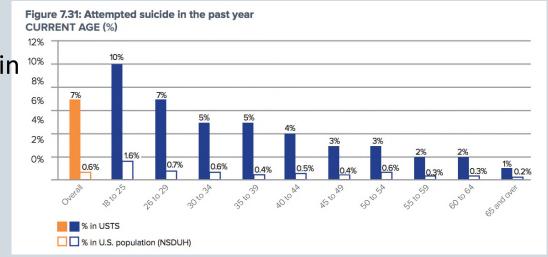
- Poverty/Homelessness
- Access to Healthcare
- Unemployment
- Substance Abuse
- Bullying & Physical/Sexual Assault
- Sex Work
- HIV/STDs
- Obesity
- Eating and Body Image Disorders
- Depression/Anxiety
- Suicidality



# Suicidality<sup>10</sup>

40% of respondents have attempted suicide in their lifetime, nearly nine times the rate reported in the general U.S. population (4.6%)

Seven percent (7%) of all respondents attempted suicide in the past year, nearly twelve times the rate of attempted suicide in the U.S. population (0.6%).



# Health Disparities<sup>10</sup>

- Refusal/Denial of Insurance Coverage
- Discrimination within the medical setting
- Delay in seeking health care
- Lack of population specific competent care



# Health Disparities<sup>7</sup>



33% had at least one negative experience with a doctor or other health care provider related to being transgender over the past year - Verbal harassment, refusal of treatment, or having to teach the health care provider about transgender health

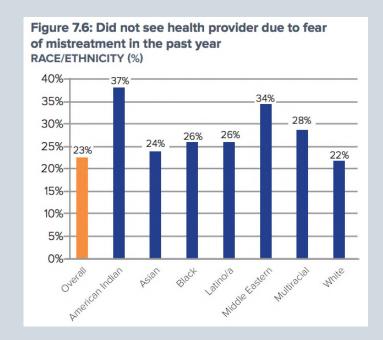
# Health Disparities<sup>10</sup>

23% did not see a doctor when they needed to because of fear of being mistreated as a transgender person

24% had to teach the provider about transgender health in order to receive appropriate care

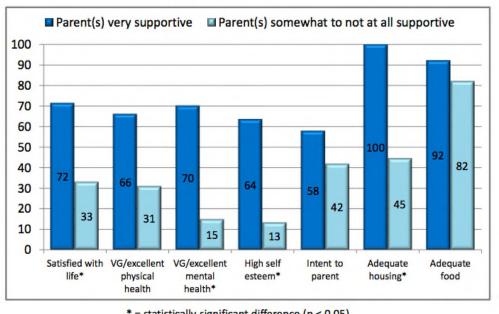
8% were refused transition-related health care

6% were harassed in a health care setting



# Societal Support<sup>11</sup>

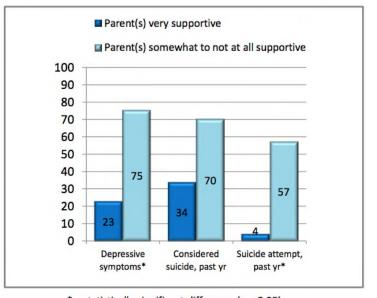
Figure 1. Proportion of trans youth age 16-24 years in Ontario experiencing positive health and life conditions, by level of parental support



\* = statistically significant difference (p < 0.05)

# Societal Support<sup>11</sup>

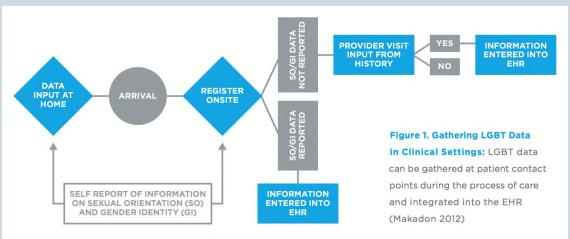
Figure 2. Proportion of trans youth age 16-24 years in Ontario experiencing negative health and life conditions, by level of parental support



\* = statistically significant difference (p < 0.05)

#### Creating a Gender Affirming Environment<sup>12</sup>

- Simple changes in forms, signage, and office practices can go far in making LGBT individuals feel more welcome
- The Institute of Medicine recommends inclusion of structured data fields to obtain information on sexual orientation and gender identity (SOGI) as part of electronic health records.



### Creating a Gender Affirming Environment<sup>12</sup>

1. Which of the categories best describes your current annual income? Please check the correct category:    <\$10,000	2. Employment Status:  □ Employed full time □ Employed part time □ Student full time □ Student part time □ Retired □ Other	3. Racial Group(s):  African American/Black  Asian  Caucasian  Multi racial  Native American/Alaskan Native/Inuit  Pacific Islander  Other	4. Ethnicity:    Hispanic/Latino/Latina   Not Hispanic/Latino/Latina    5. Country of Birth:   USA   Other
6. Language(s):  □ English □ Español □ Français □ Portugês □ Русский	7. Do you think of yourself as:  Lesbian, gay, or homosexual Straight or heterosexual Bisexual Something Else Don't know	8. Marital Status:	1. Referral Source:  Self Friend or Family Member Health Provider Emergency Room Ad/Internet/Media/ Outreach Worker/School

#### Creating a Gender Affirming Environment<sup>12</sup>

1.	What is your current gender identity? (Check an/or circle ALL that apply)
	☐ Male
	☐ Female
	☐ Transgender Male/Trans Man/FTM
	☐ Transgender Female/Trans Woman/MTF
	☐ Genderqueer
	☐ Additional category (please specify):
	□ Decline to answer
2.	What sex were you assigned at birth? (Check one)
	☐ Male
	☐ Female
	☐ Decline to answer
3.	What pronouns do you prefer (e.g., he/him, she/her)?

• Develop and display non-discrimination policies that include sexual orientation and gender

identity

 Educational brochures on LGBT health topics can be made available where other patient information materials are displayed

· Gender-neutral facilities

Provide diversity training to all staff including receptionists, medical assistants,
nurses, and physicians, to treat all LGBT
patients with respect including using patients' preferred names and pronouns



#### Front-line Staff

- Front desk staff, nursing staff, lab and x-ray staff, etc.
- Critical Role
- Address people without using terms indicating gender
  - Sir/Ma'am, Mr./Mrs./Miss/Ms.
- If unsure of name or pronouns, ask...
  - What name and pronouns would you like me/us to use?"
  - Never ask a person what their "real" name is.

### Name/Pronoun Awareness



We should respect the pronouns a person uses

- She/Her/Hers
- He/Him/His
- They/Them/Theirs
- Others

Avoid the phrase "preferred pronouns"

- "What pronouns do you use?"

#### Making Mistakes

- Many providers are uncomfortable discussing gender identity with patients due to fear of making a mistake or upsetting a patient
- It is okay to make mistakes, as long as you are considerate towards the person you are addressing
- If you do make a mistake, simply apologize a thoughtful apology can go a long way in changing their experience, even beyond your interaction

Continue to educate yourself about LGBT health topics

- Glossary of LGBT Terms https://www.lgbthealtheducation.org/publication/lgbt-glossary/
- Gender Spectrum https://www.genderspectrum.org/
- UCSF Center of Excellence for Transgender Health http://transhealth.ucsf.edu
- TransLine http://project-health.org/transline/
- TransYouth Family Allies http://www.imatyfa.org/healthcare.html
- Endocrine Society Guidelines Endocrine Treatment of Gender-Dysphoric/Gender-Incongruent Persons www.endo-society.org/guidelines
- WPATH Standards of Care http://www.wpath.org/





#### **GnRH Agonist**

Available at Tanner Stage 2-5

Suppresses pubertal progression

Time to explore gender/development and desired outcomes

Can stop & proceed with endogenous puberty

Prevents development of undesired sexual characteristics that are difficult or impossible to erase

Decrease need for various surgical procedures

May allow for lower dosages of hormone replacement therapy

#### **GnRH Agonist**

Subcutaneous Implants (Histrelin)

Effective for 12+ months

Intramuscular Injection (Leuprolide/Triptore

Every 1, 3 or 6 month preparations



#### **GnRH Agonist & Fertility**

Treating early puberty temporarily impairs spermatogenesis and oocyte maturation

Delaying or temporarily discontinuing blockers to promote gamete maturation is an option

Not often preferred because mature sperm production is associated with later stages of puberty and significant secondary sex characteristic development

Reproductive Endocrine Gynecologist can counsel before hormone treatment or surgery regarding potential fertility options

Early suppression of trans-females may lead to insufficient penile tissue for vaginoplasty



70 68

66

64

#### **GnRH Agonist & Brain Development**

Pubertal suppression with GnRH agonists is not associated with a detrimental effect on higher order cognitive processing (ToL performance scores - reaction time or accuracy)<sup>14</sup>

Psychological support and puberty suppression were both associated with an improved global psychosocial functioning in

GD adolescents<sup>15</sup>

eligible GD adolescents

\*, P<0.05

\*\*, P<0.01

\*\*\*, P<0.001

\*\*\*, P<0.001

\*\*\*\*, P<0.001

\*\*\*\*, P<0.001

Figure 2 Gender dysphoria adolescents' psychosocial functioning (CGAS) at baseline, after psychological support, and after puberty suppression

CGAS, Children's Global Assessment Scale; Time 0, baseline; Time 1, 6 months from baseline (after 6 months of psychological support); Time 2, 12 months from baseline (delayed eligible gender dysphoria [GD] adolescents, after 12 months of psychological support; immediately eligible GD adolescents, after 12 months of psychological support + 6 months of puberty suppression); Time 3, 18 months from baseline (delayed eligible GD adolescents, after 18 months of psychological support + 12 months of puberty suppression)

—All participant 15

Delayed eligible

**GD** adolescents

Immediately

#### **GnRH Analogues & Bone Health**

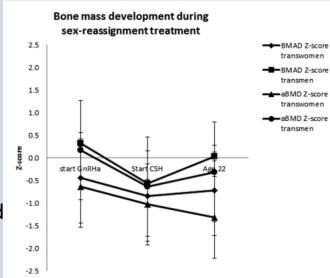
BMD below pretreatment potential suggests delayed attainment of peak bone mass or attenuation of peak bone mass<sup>16</sup>

FTM individuals - sex steroid therapy does not seem to be associated with significant changes in BMD<sup>17</sup>

MTF individuals - sex steroid therapy appears to be associated with increased BMD<sup>17</sup>

The impact of these BMD changes on patient important outcomes such as fracture risk remains uncertain.

Delayed growth plate closure



**Figure 1.** Longitudinal z-score (mean  $\pm$  SD) development of the LS from start medical treatment until the age of 22 years in transmen and transwomen.

#### **Gender Affirming Hormones**

Exogenous 17-beta-Estrogen or Testosterone

Develop desired secondary sex characteristics and suppress/minimize undesired secondary sex characteristics

Age availability varies between institutions - 13.5 to 16 years for initiation with parental consent

If suppressed, extensive delay could impact bone health and psychosocial development



## Hormonal Options for Transitioning

### **Adjunctive Therapies**

Bicalutamide: androgen receptor blocker

- Liver disease

Spironolactone: androgen receptor blocker & testosterone synthesis suppression

- Diuretic effect; Hyperkalemia; Negative effects of well being, energy or mood

5-alpha reductase inhibitors: blocks conversion of testosterone to dihydrotestosterone

Oxandrolone: weak androgen; escapes aromatization into estrogen

Progesterone: gonadotropin suppression; possible improved breast development;

cessations of menses

## Hormonal Options for Transitioning

#### **Testosterone**

Injection, Topical, Pellet Implant

SubQ vs IM route: subQ route equal efficacy and improved patient satisfaction, less pain, smaller needle, more gradual absorption, avoid IM fibrosis from long term (possibly > 50 years) IM therapy  $^{18}$ 

#### **Side Effects**

Hair loss

Unwanted body hair

Acne

Vaginal atrophy

Effects on mood/energy

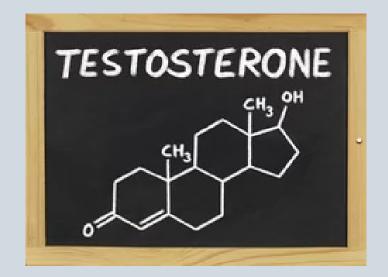
Weight/Appetite changes

Secondary exposure

Impaired fertility

Abnormal fetal development

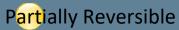
Changes in sex drive



#### **Expected Onset**<sup>B</sup> **Expected Maximum Effect**<sup>8</sup> **Effect** Skin oiliness/acne 1-6 months 1-2 years Facial/body hair growth 3-6 months 3-5 years Scalp hair loss >12 months<sup>C</sup> variable Increased muscle mass/strength 6-12 months 2-5 years<sup>D</sup> Body fat redistribution 3-6 months 2-5 years Cessation of menses 2-6 months n/a Clitoral enlargement 3-6 months 1-2 years Vaginal atrophy 3-6 months 1-2 years Deepened voice 3-12 months 1-2 years

TABLE 1A: EFFECTS AND EXPECTED TIME COURSE OF MASCULINIZING HORMONES A







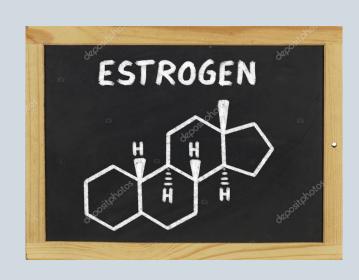


A Adapted with permission from Hembree et al.(2009). *Copyright 2009, The Endocrine Society*. Estimates represent published and unpublished clinical observations.

C Highly dependent on age and inheritance; may be minimal.

D Significantly dependent on amount of exercise.

# Hormonal Options for Transitioning



### 17-beta Estradiol

Injection, Topical, Oral/Sublingual

Ethinyl estradiol/Premarin not recommended

Increased adverse health effects & unable to measure in serum

#### Side effects

Exacerbation of migraines/seizure
Effects on mood
Hot flashes
Weight changes
Impaired fertility
Changes in energy
Decreased libido and erectile dysfunction

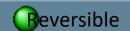
#### **Effect** Expected Onset<sup>8</sup> Expected Maximum Effect<sup>B</sup> Body fat redistribution 3-6 months 2-5 years Decreased muscle mass/ 1-2 years<sup>C</sup> 3-6 months strength Softening of skin/decreased 3-6 months unknown oiliness Decreased libido 1-3 months 1-2 years Decreased spontaneous 3-6 months 1-3 months erections Male sexual dysfunction variable variable Breast growth 3-6 months 2-3 years Decreased testicular volume 3-6 months 2-3 years Decreased sperm production variable variable Thinning and slowed growth of > 3 years<sup>D</sup> 6-12 months body and facial hair No regrowth, loss

stops 1-3 months

1-2 years

TABLE 1B: EFFECTS AND EXPECTED TIME COURSE OF FEMINIZING HORMONES A

Male pattern baldness









A Adapted with permission from Hembree et al. (2009). Copyright 2009, The Endocrine Society.

<sup>&</sup>lt;sup>B</sup> Estimates represent published and unpublished clinical observations.

<sup>&</sup>lt;sup>C</sup> Significantly dependent on amount of exercise.

D Complete removal of male facial and body hair requires electrolysis, laser treatment, or both.

#### Thromboembolic Disease

Additional increase risk in >40 years of age, tobacco users, sedentary and obese

Risk decreased with transdermal/IM route of estradiol

Highest risk during the first year of therapy

### Cardiovascular/Cerebrovascular Disease

Increases rates of VTE and ischemic stroke but not myocardial infarctions

Higher rates of tobacco use, obesity, age >40 years, diabetes and lipid disorders, and reduced

physical activity

Transdermal route has lower risk

### Epilepsy/Seizure Disorder

Lowers seizure threshold and may exacerbate underlying seizure disorders

### Migraines

Exacerbate migraine events



#### Cancer

No identified difference in general cancer rate in transgender patients on hormone therapy compared to sex-assigned controls.

Insufficient evidence of changes in risk factor for organ-specific cancer risk

**Breast Cancer** 

Transwomen on estrogen have developed breast cancer but degree of risk compared to cisgender female peers poorly studied

Duration of estrogen exposure, family history of breast cancer, obesity and use of progestins likely influence the level of risk

If a patient has a particular organ, screening should occur regardless of hormone use

### Hyperprolactinemia

Increased risk during the first year of therapy

Unlikely to develop after the first year

May promote the clinical appearance of pre-existing but clinically unapparent prolactinoma

Expectant management only in absence of visual disturbance, galactorrhea or new onset headaches

### Liver Disease

May have transient liver enzyme elevations and, rarely, clinical hepatotoxicity

### Peri-operative Use

Many surgeons prefer that estrogen be discontinued for at least 2 weeks before and after any procedure

Can have profound impact to patient

No evidence suggests that transgender women who lack risk factors (personal/ family history, excessive use of estrogen, smoking) must cease peri-operatively

Appropriate use of prophylaxis (heparin or compression devices)

Informed consent of pros and cons

#### Alternatives:

Lower estrogen dose

Convert to transdermal route

## Risks of Masculinizing Hormone Therapy

### Polycythemia/Erythrocytosis

Hgb & Hct levels should be interpreted in terms of dosing and menstruation status

- Physiologic male range testosterone and amenorrhea: expect male range Hgb/Hct

### Mental Health Conditions/Aggression

No clear evidence of direct association between testosterone and mental health status

May see some influence when on higher doses or supra-physiologic blood levels

#### **Hair Loss**

Unpredictable nature, extent and time course

Managed with 5-alpha reductase inhibitors

## Risks of Masculinizing Hormone Therapy

### Metabolic Syndrome/PCOS

Not contraindicated but require monitoring for dyslipidemias and diabetes

### Liver Disease

May have transient liver enzyme elevations

#### Cancer

No clear increased risk for breast, cervical, ovarian or endometrial cancers

#### Cardiovascular

Evidence suggests that risk is unchanged among transgender men using testosterone compared with non-transgender women

# Surgical Interventions

### Transgender women

**Top Surgery** 

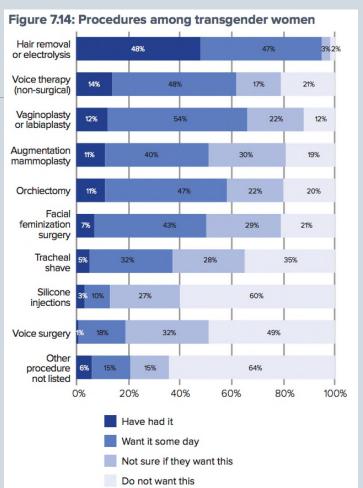
- Augmentation mammoplasty

#### **Bottom Surgery**

- Vaginoplasty
- Orchiectomy

#### Other

- Facial feminization surgery
- Reduction thyrochondroplasty (tracheal cartilage shave)
- Voice surgery
- Facial/Body Hair Removal



Washington University in St. Louis
School of Medicine

## Surgical Interventions

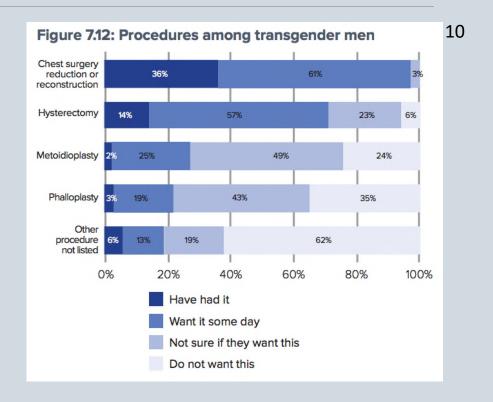
### Transgender men

**Top Surgery** 

- Masculinizing Chest Surgery (Mastectomy)

**Bottom Surgery** 

- Phalloplasty/Metoidioplasty
- Hysterectomy



## Other Special Considerations

### **Chest Binding**

Provides a flat chest contour

Tight fitting sport bras, shirts, ace bandages,

Prolonged use can cause breast pain, skin irritation, fungal infections, rib damage



### **Genital Tucking**

Provides a flat groin contour

Testicles moved into inguinal canal

Penis and scrotum moved posteriorly to the perineum

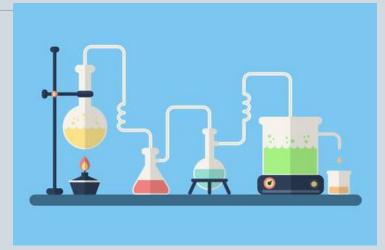
Tight fitting underwear or special undergarment (gaffe)

Some use adhesives or duct tape

Prolonged use can cause skin irritation, hernias, genitourinary tract trauma/infection, testicular pain/torsion/trauma

# Laboratory Sciences

- Transgender appropriate reference interval studies are virtually absent
- Not all patients that have name or gender marker changed are on gender-affirming hormone therapy
- Little to no guidance on when to transition sex-specific normal ranges
  - Many transition after 6 months of hormone therapy
- Lab supplied references ranges may not be appropriate
- Consult with lab to obtain reference ranges for both 'male' and 'female' norms then apply the most appropriate range when interpreting results



## Laboratory Sciences

Alkaline phosphatase, hemoglobin and hematocrit, and creatinine may vary depending on the patient's current sex hormone configuration

- Several factors contribute to these differences
  - Bone mass
  - Muscle mass
  - Number of myocytes
  - Changes on menstruation
  - Erythropoetic effects
  - Potential for pulsatile undetected androgen activity in those with retained gonads
- No empirical or published evidence of which eGFR equation is better suited for transgender individuals

## Laboratory Sciences - Feminizing Therapy

Table 3. Lower and upper limits of normal to use when interpreting selected lab tests in transgender women using feminizing hormone therapy

Lab measure	Lower Limit of normal	Upper Limit of normal
Creatinine	Not defined	Male value
Hemoglobin/Hematocrit	Female value	Male value
Alkaline Phosphatase	Not defined	Male value

20

### Laboratory Sciences - Masculinizing Therapy

Table 3. Lower and upper limits of normal to use when interpreting selected lab tests in transgender men using masculinizing hormone therapy

Lab measure	Lower Limit of normal	Upper Limit of normal
Creatinine	Not defined	Male value
Hemoglobin/Hematocrit	Male value if amenorrheic*	Male value
Alkaline Phosphatase	Not defined	Male value

<sup>\*</sup> If menstruating regularly, consider using female lower limit of normal.

## Laboratory Sciences

### **Bone Health**

- Growth Plate Maturation & Bone Mineralization
- Standards for interpretation are based on age matched male or female standards
- No guidelines on how to interpret standards in transgender patients undergoing hormonal therapy
  - GnRH agonist vs Gender-Affirming Hormones vs Other hormonal options
- When to compare to cis-gender peers matched for sex assigned at birth or affirmed gender

# Questions?



Compassion is the basis of all morality.

Arthur Shopenhaur



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