# NEXT GENERATION SEX AND GENDER SCIENCE AND EDUCATION IN MEDICINE

Johannes Gutenberg University, Mainz

February 23, 2023

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She/her/elle Professor of Medicine Université de Montréal, Canada



### I'M GOING TO INVITE YOU TO REFLECT WITH ME ON:

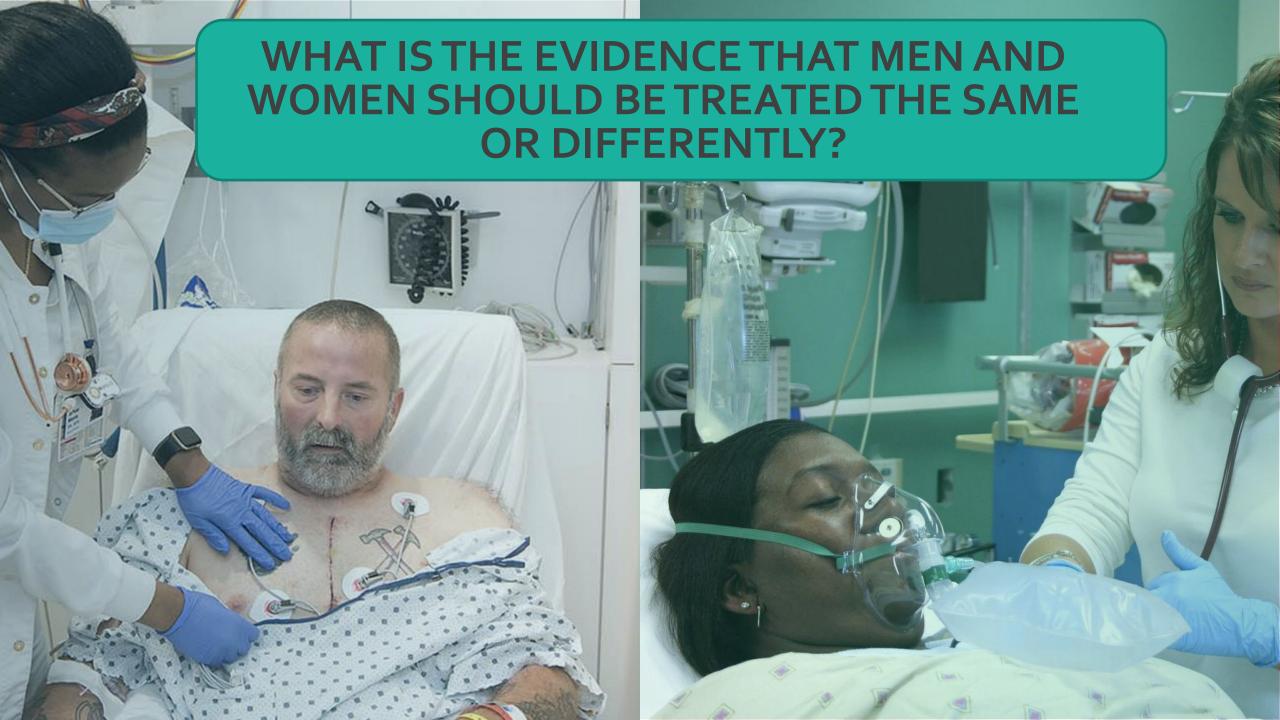
A vision for the next generation of sex and gender science

Why it is important to consider sex and gender in medicine

What actions YOU can take to make research, education and practice more rigorous, more inclusive and more impactful

### SOME ASSUMPTIONS





# ASSUMPTION 2 YOU KNOW HOW TO DISTINGUISH BETWEEN SEX AND GENDER

### SEX

Biological attributes of humans and animals, including physical features, chromosomes, gene expression, hormones and anatomy.



### **GENDER**

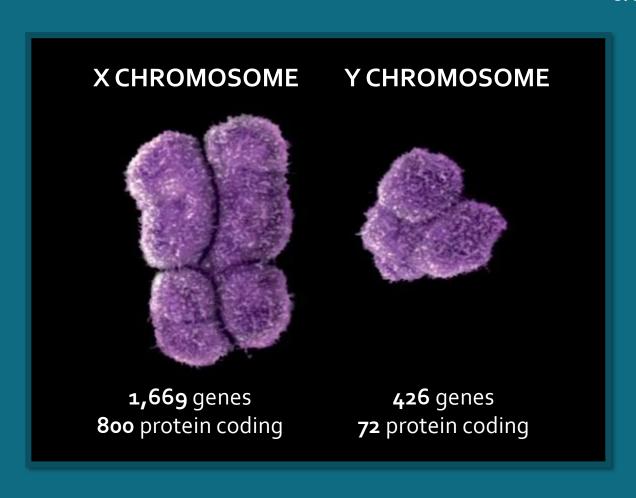
Socially-constructed roles, behaviours, expressions and identities of girls, women, boys, men and gender diverse people.



### **GENETIC SEX IS IMPORTANT**

Only 32.5% of genome-wide association studies included the sex chromosomes.

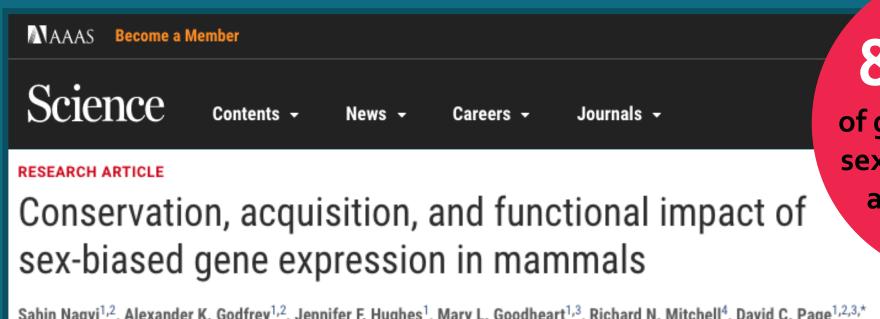
Wise et al. Am. J. Hum Genet. 2013; 92(5): 643-647.



Incomplete X chromosome inactivation affects at least 23% of X-chromosomal genes.

Leads to a gene dosage imbalance between sexes.

### AUTOSOMES ARE IMPORTANT PLAYERS IN SEX-BIASED GENE EXPRESSION



85-90%

of genes exhibiting sex bias in humans are autosomal.

Sahin Naqvi<sup>1,2</sup>, Alexander K. Godfrey<sup>1,2</sup>, Jennifer F. Hughes<sup>1</sup>, Mary L. Goodheart<sup>1,3</sup>, Richard N. Mitchell<sup>4</sup>, David C. Page<sup>1,2,3,\*</sup>

+ See all authors and affiliations

Science 19 Jul 2019: Vol. 365, Issue 6450, eaaw7317

#### EVEN PROTEINS CAN BE INFLUENCED BY SEX

Cell

**Article** 

### Disentangling Genetic and Environmental Effects on the Proteotypes of Individuals

Natalie Romanov, 1,7 Michael Kuhn, 1 Ruedi Aebersold, 2,3 Alessandro Ori, 4 Martin Beck, 1,5,7,\* and Peer Bork 1,6,8,\*

Romanov et al., 2019, Cell 177, 1308-1318

Sex can explain
13% of differences
in protein complex
abundance and
stoichiometry.

<sup>&</sup>lt;sup>1</sup>Structural and Computational Biology Unit, European Molecular Biology Laboratory, Heidelberg, Germany

<sup>&</sup>lt;sup>2</sup>Department of Biology, Institute of Molecular Systems Biology, ETH Zurich, Switzerland

<sup>&</sup>lt;sup>3</sup>Faculty of Science, University of Zurich, Zurich, Switzerland

<sup>&</sup>lt;sup>4</sup>Leibniz Institute on Aging - Fritz Lipmann Institute, Jena, Germany

<sup>&</sup>lt;sup>5</sup>Cell Biology and Biophysics Unit, European Molecular Biology Laboratory, Heidelberg, Germany

<sup>&</sup>lt;sup>6</sup>Max Delbrück Center for Molecular Medicine, Berlin, Germany

<sup>&</sup>lt;sup>7</sup>Present address: Max Planck Institute of Biophysics, Max-von-Laue-Straße 3, Frankfurt am Main, Germany

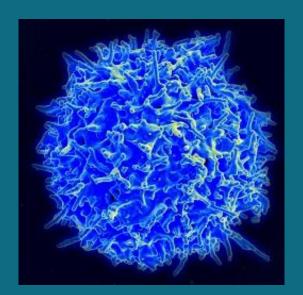
<sup>8</sup>Lead Contact

<sup>\*</sup>Correspondence: martin.beck@embl.de (M.B.), bork@embl.de (P.B.) https://doi.org/10.1016/j.cell.2019.03.015

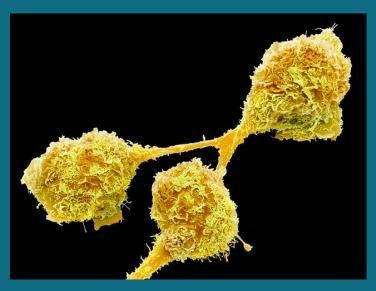
# INCLUDING MALES AND FEMALES LEADS TO NEW DISCOVERIES

## Chronic pain mechanisms are mediated by different immune cells

Female Mice - T Cells



Male mice – Microglial Cells



Sorge et al. Different immune cells mediate mechanical pain hypersensitivity in male and female mice. Nat Neuroscience 2015; 31:1165-1167.

### Historical representation of the research pipeline

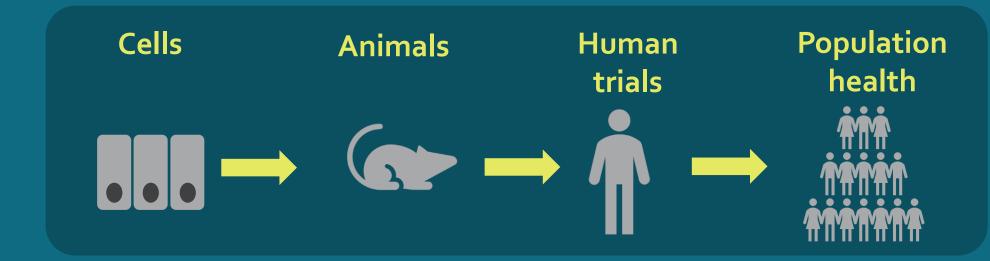
**76%** MALE/NOT REPORTED

**75% MALE** 

67% MEN

51% WOMEN

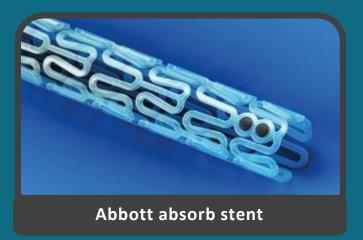
WOMEN > MEN

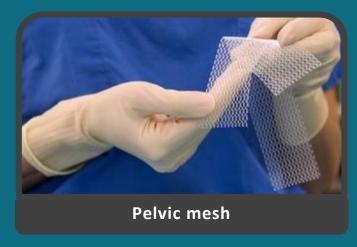


Clinical care

#### WHY IS THIS IMPORTANT?

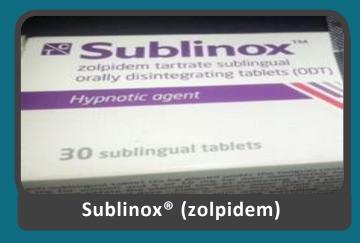
If sex is not accounted for at both the pre-clinical and testing phases for drugs, biologics and medical devices, then the benefits may not be equal. At worst, we may be introducing harm

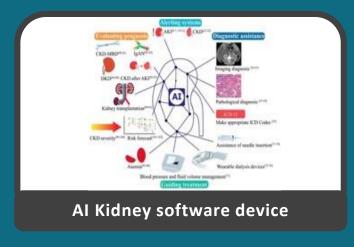




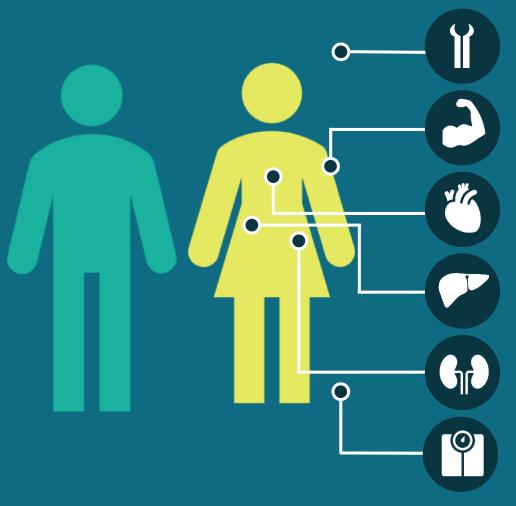








### Women process medicines differently than men.



Women's immune system is more active, mounting twice as many antibodies to many vaccines as men.

Women usually have more body fat than men, which means fatsoluble medicines might linger.

Women's **heart electrophysiology** is slightly different from men's, so medicines might have different risks or results.

Women have 77 genes for medicine metabolizing enzymes or transporters in the liver that are expressed differently due to sex. Estrogen affects how medicines are metabolized by the liver.

Women's kidneys are smaller, eliminating medicines from the body at a 15-20% slower rate.

Women tend to be smaller than men, so the same dosage might have a smaller volume of distribution & a greater effect.

### SEX/GENDER MEDICINE

When research is done "properly" in males and females, we move towards safer personalized medicine: the right dose at the right time for the right person

Test 2 doses of the medicine for women to see if they can have equal benefit but lower side effects from lower dose.

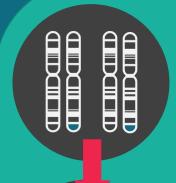






Women: 25 ug dose

### Women showed higher desmopressin sensitivity than men.



Desmopressin is an AVP analog drug that binds to the arginine vasopressin (AVP) receptor in the kidney to prevent excess urination. The AVP receptor is coded by the AVPR2 gene, which lies on the X chromosome.



Normally in females, one of the two X chromosomes gets randomly inactivated. The random inactivation of one of the X chromosomes in female cells prevents differences in X chromosome gene dosage between male and female cells for most genes.



However, some X-linked genes **escape** inactivation in females (12%–20% in humans), and therefore those genes are expressed at higher levels in females than in males.



Some women may have an overexpression of the AVPR2 gene.



More receptors on the cells



Higher effect of AVP, more water retained, higher risk of hyponatremia

Relative female/ male sensitivity to the antidiuretic effect of desmopressin in nocturia is 2.7, corresponding to significantly higher desmopressin sensitivity in women.

# 10 medicines were withdrawn from the U.S. market between 1997 and 2000.



of these medicines posed greater health risks for women than men.

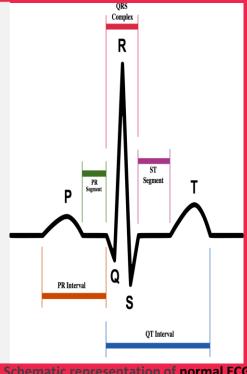


### Sex AND gender played a role

### **Biological factors**

Seldane®, Propulsid®

cause fatal arrythmias in women only because of sex differences in the heart



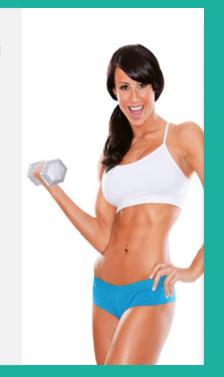
Schematic representation of normal ECG

### Social factors

Redux ®, Fen-Fen appetite suppressants

cause valvular heart disease equally

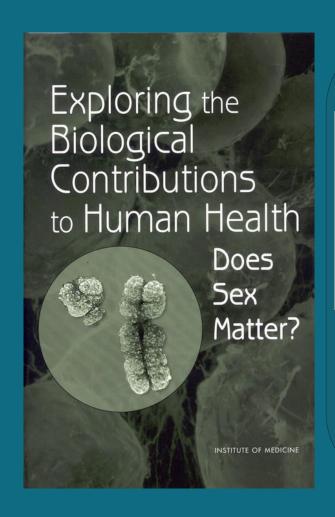
but more women take them



### **ASSUMPTION 3**

# YOU MAY VIEW SEX AND GENDER AS BINARY CONCEPTS

# IN 2001, THE U.S. INSTITUTE OF MEDICINE DEFINED SEX AND GENDER



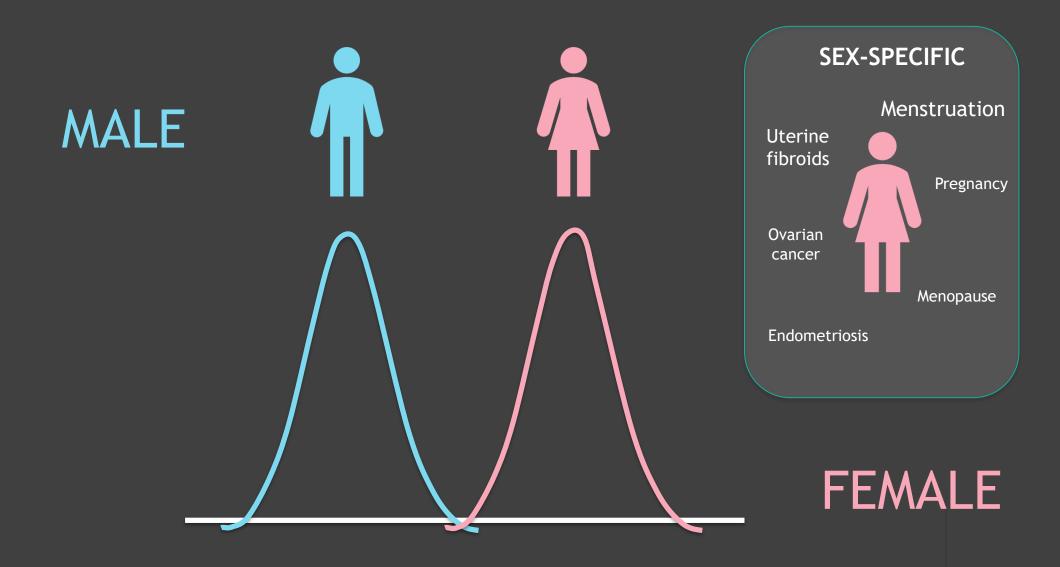
#### BOX 1-1 Definitions

**Sex:** The classification of living things, generally as male or female according to their reproductive organs and functions assigned by chromosomal complement.

**Gender:** A person's self-representation as male or female, or how that person is responded to by social institutions based on the individual's gender presentation. Gender is rooted in biology and shaped by environment and experience.

Institute of Medicine (US) Committee on Understanding the Biology of Sex and Gender Differences; Editors: Theresa M. Wizemann and Mary-Lou Pardue. Washington (DC): National Academies Press (US); 2001.

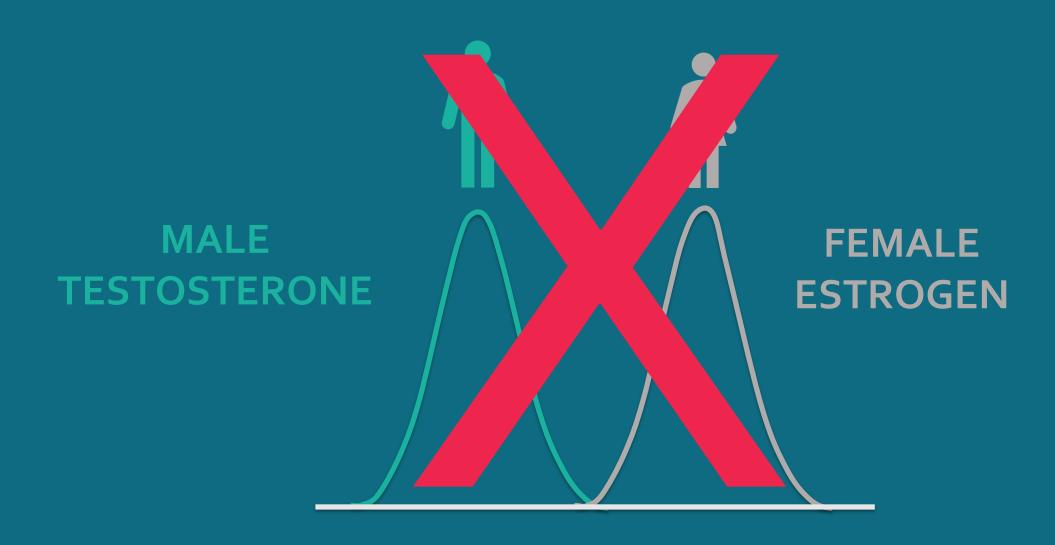
### BINARY/DICHOTOMOUS TRAITS



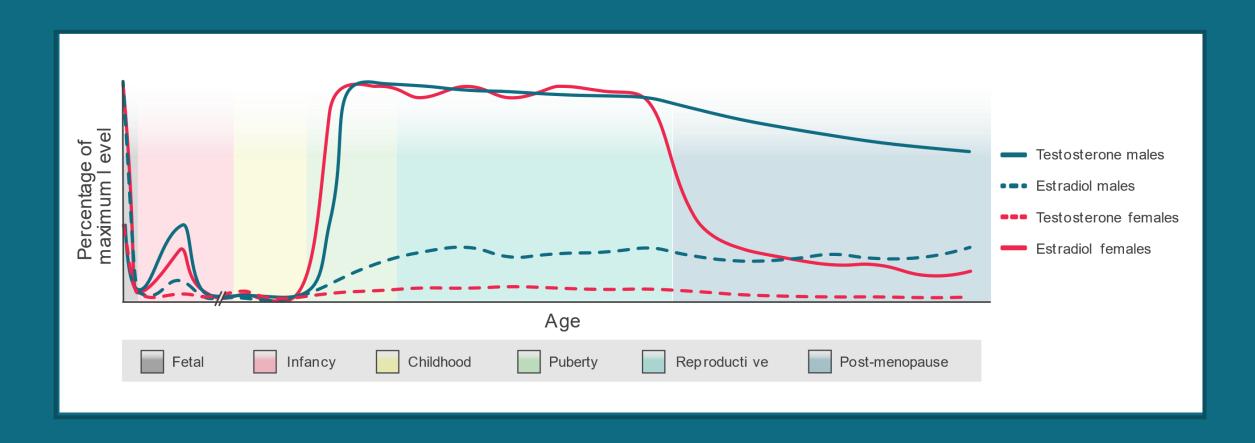
# SEX CAN COME IN MANY DIFFERENT SHAPES AND SIZES



# THINK ABOUT SEX HORMONES. IS IT APPROPRIATE TO REFER TO:



# ESTROGEN IS NOT A "FEMALE" HORMONE AND TESTOSTERONE IS NOT A "MALE" HORMONE



### SEX EXISTS ON A SPECTRUM...

SEX

Male Intersex Female

### ... JUST LIKE GENDER

**GENDER IDENTITY** 

Man/boy

Transgender / Genderqueer / Two-spirit / Etc.

Woman/girl

**GENDER EXPRESSION** 

### **REAL-LIFE HARM & CONFUSION**



#### Sex, gender, and sports

New regulations challenge the evidence based, benevolent ethos that underlies medical practice

BMJ 2019; 364 doi: https://doi.org/10.1136/bmj.l1120 (Published 20 March 2019)

Cite this as: BMJ 2019;364:I1120

Cara Tannenbaum professor of medicine and pharmacy<sup>12</sup>, Sheree Bekker prize research fellow<sup>3</sup>

### MULTIDIMENSIONALITY OF GENDER

#### **IDENTITY**

- ✓ Inner sense of self as masculine, feminine, both, or neither
- ✓ Cisgender, transgender or gender-nonconforming

#### ROLES

- ✓ Societal expectations
- ✓ Influences on everyday actions
- ✓ Career you pursue, role in family, etc.

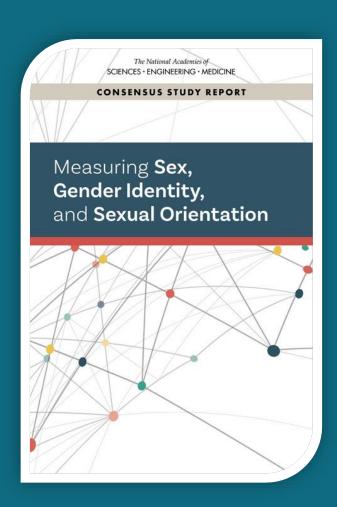
#### **RELATIONS**

✓ Interpersonal interactions E.g., family, workplace dynamics

#### **INSTITUTIONALIZED**

- Distribution of power in political, educational, social institutions in society
- ✓ Shapes social norms

# IN 2022, THE U.S. NATIONAL ACADEMIES OF SCIENCES, ENGINEERING AND MEDICINE EXPANDS DEFINITIONS OF SEX AND GENDER



#### **TERMINOLOGY**

**Sex** is a multidimensional construct based on anatomical and physiological traits (sex traits) that include external genitalia, secondary sex characteristics, gonads, chromosomes, and hormones. *Intersex* refers to people whose sex traits do not all correspond to the same sex.

Gender is a multidimensional construct that links gender identity, which is a core element of a person's individual identity; gender expression, which is how a person communicates their gender to others; and social and cultural expectations about status, characteristics, and behavior that are associated with sex traits. Transgender refers to people whose current gender identity is different from the sex they were assigned at birth. Transgender experience refers to all people who can be classified as transgender and transgender identity refers to people who identify as transgender. Cisgender refers to people whose current gender identity corresponds to the sex they were assigned at birth. Nonbinary is an umbrella term for gender identities that lie outside the gender binary.

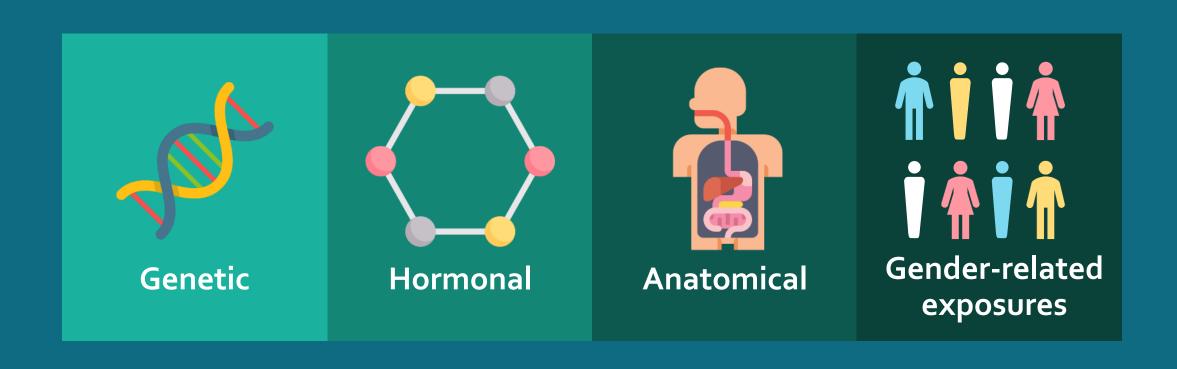
National Academies of Sciences, Engineering and Medicine. Measuring Sex, Gender Identity and Sexual Orientations. 2022

# THE BIOLOGICAL AND THE SOCIAL DO NOT EXIST IN ISOLATION



Sex Gender

# DETERMINING THE SOURCE OF VARIABILITY IN LUNG DISEASE WITH SEX AND GENDER SCIENCE



## COMBINING SEX AND GENDER ANALYSIS: ASTHMA AS AN EXAMPLE

Asthma is more common in males before puberty, but more common in females after puberty.

Is this due to SEX (BIOLOGICAL) or GENDER (SOCIOCULTURAL/BEHAVIOURAL) factors?

#### Sex-related hypotheses

- Sex differences in lung development
- Sex differences in hormone levels
- Sex differences in atopy and immune responses to allergens

#### Gender-related hypotheses

- Differences in gendered play and occupation
- Differences in smoking habits
- Differences in beliefs about adherence to medication, especially during pregnancy

Gendered factors can influence our biology. For example, boys may be more likely to engage in outdoor play, which influences the types of microbes and pollutants they are exposed to. These exposures can influence epigenetic processes, such as DNA methylation, histone modification, which can in turn modify gene expression between sexes.

Sex versus gender contributions are difficult to separate. Both should be factored in study designs involving humans whenever possible.

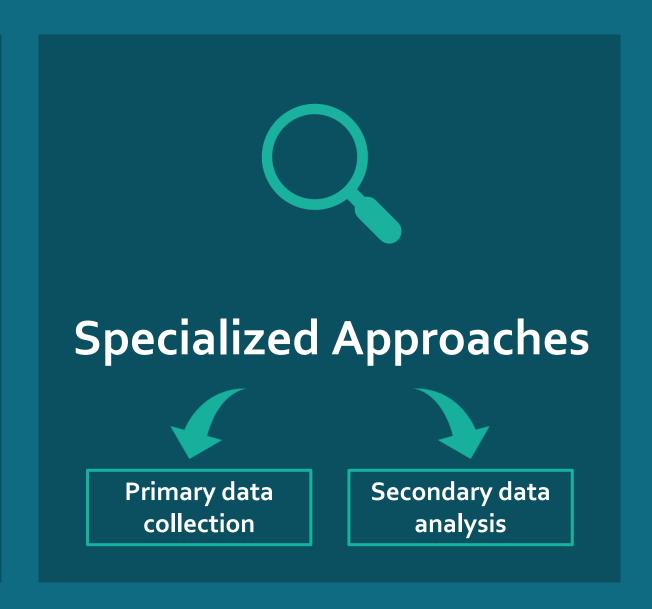
WHAT ACTIONS CAN YOU TAKE TO APPLY A SEX AND GENDER SCIENCE LENS TO RESEARCH, **EDUCATION AND** PRACTICE?



### IT'S NOT A ONE-SIZE-FITS-ALL APPROACH



General Research
Principles



### STEP

### Know your literature!

What is currently known about sex and/or gender differences in the incidence, manifestations, mechanisms, progression and **response to treatment** of the condition under study?

#### **nature reviews** immunology

Perspective | Published: 11 June 2020

Considering how biological sex impacts immune responses and COVID-19 outcomes

Eileen P. Scully, Jenna Haverfield, Rebecca L. Ursin, Cara Tannenbaum & Sabra L. Klein S

Nature Reviews Immunology 20, 442-447(2020) | Cite this article



The influence of sex and gender domains on COVID-19 cases and mortality

Christina P. Tadiri, Teresa Gisinger, Alexandra Kautzky-Willer, Karolina Kublickiene, Maria Trinidad Herrero, Valeria Raparelli, Louise Pilote and Colleen M. Norris; for the GOING-FWD Consortium

CMAJ September 08, 2020 192 (36) E1041-E1045; DOI: https://doi.org/10.1503/cmaj.200971



Springer Canadian Journal of Public Health

Can J Public Health. 2020 Dec; 111(6): 980-

983.

Published online 2020 Sep 29.

doi: 10.17269/s41997-020-00417-z

Language: English | French

PMCID: PMC7523484

PMID: 32990927

Sex, gender and COVID-19: a call to action

Sofia B. Ahmed<sup>™1,2,3</sup> and Sandra M. Dumanski<sup>1,2,3</sup>

# AS A GENERAL RULE...

To integrate sex and/or gender as variables in your research, there should be **theoretical** or **empirical** *rationale*.

You should integrate **SEX** into your research if there is biological plausibility to do so, for example:

- Does the phenomenon, condition or disease under study affect the sexes differently?
- Can the model system being used be differentiated by sex?
- Are there preliminary data suggesting a unique influence of sex?

You should integrate **GENDER** into your research if there is sociocultural plausibility to do so, for example:

- Does the study involve humans?
- Are there preliminary data suggesting a unique influence of gender?

### STEP 2

#### Include males and females

Report the sex of your cells, tissues, animals, human participants in your experiments when studying models of disease that affect all sexes.



## STEP 3

# Determine which aspect(s) of gender is relevant to your research.

# PRIMARY DATA COLLECTION

Design the study to measure the aspect(s) of gender based on theoretical or empirical rationale in the human population under study.

# SECONDARY DATA COLLECTION

Determine if data on gender identity, or other gender-related variables, are available in the exiting data set being analysed.

#### TWO STEP QUESTION ON SEX AND GENDER IDENTITY

Q1. What sex were you assigned at bin on your original birth certificate?	
Q2. Which best describes your current identity?    Man/Boy   Woman/Girl   Gender-diverse   Non-binary   Prefer to self-describe (please	The two-step question on sex and gender identity can also be used when seeing patients

# EXAMPLES OF OTHER GENDER-RELATED VARIABLES THAT MAY BE RELEVANT

#### **ROLES**



Housework responsibilities



Childcare responsibilities



Weekly work

#### **RELATIONS**



Civil status



Spousal support



Relationships with others

#### **INSTITUTIONALIZED**



Personal income



Education level



Status in country & community

### STEP 4

#### Ensure appropriate sample size.

If there is theoretical or empirical rationale, the sample size should be statistically powered to allow for sex and/or gender specific sub-group analyses.

Depending on statistical power, there are three types of analyses you could do.

High statistical power

Low statistical power

Effect modification & interaction by sex

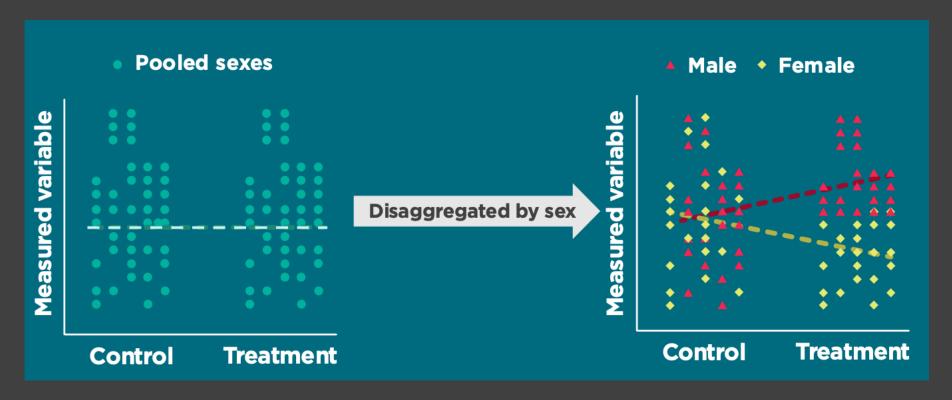
Detect main effects within sex strata

Disaggregate by sex

If underpowered, subgroup analyses heighten the risk for type II error, or false negatives.



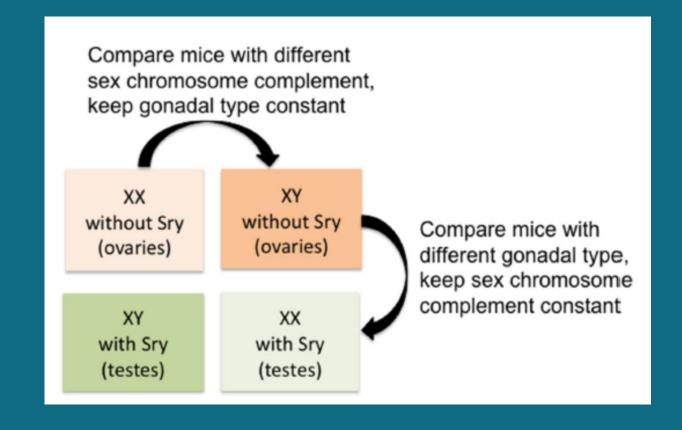
#### Pitfalls of not disaggregating by sex



#### **Basic Science**



#### If you find a sex difference, ask why



# NEW METHODS THE 4-CORE GENOTYPES MODEL

Arnold et al. Front Neuroendocrinol. Jan 2009 Clayton JA. FASEB J. 2016

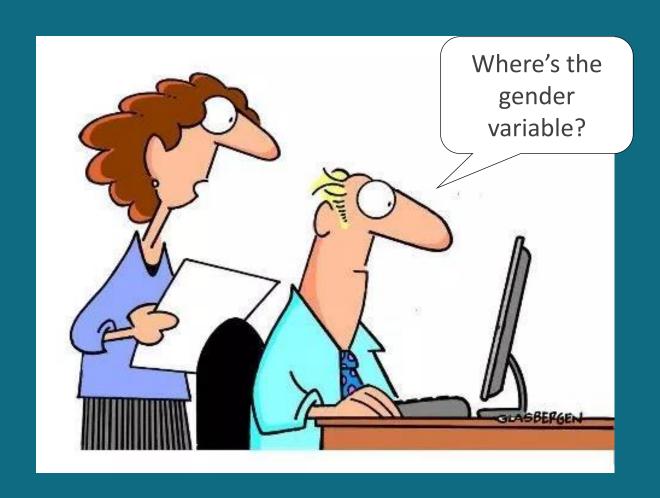
# HUMAN STUDIES: DISAGGREGATE YOUR DATA BY SEX AND GENDER

Stratify all results by sex & gender or you risk missing important differences

You can't

control for women!

# COULD A SEX DIFFERENCE BE EXPLAINED BY A GENDER-RELATED VARIABLE?



Income?

Housework responsibilities?

Marital status?

Education level?

Number of hours spent caregiving per week?

Work hours per week?

# BEST PRACTICES FOR INCORPORATING SMALL SUBGROUPS IN QUANTITATIVE ANALYSES



For example, for sex-skewed diseases or non-binary individuals.



If possible, oversample the underrepresented subgroup so that the sample size will have enough power to draw statistically valid results.



Where privacy standards allow, disaggregate the raw data and report for future meta analyses.

GENERAL RULE Do not exclude data even if the sample size is low - include and present the raw data, if possible.

#### SEX AND GENDER ONLINE TRAINING MODULES

SEX AND GENDER IN BIOMEDICAL RESEARCH



SEX AND GENDER IN PRIMARY
DATA COLLECTION WITH HUMANS



SEX AND GENDER IN THE ANALYSIS OF DATA FROM HUMAN PARTICIPANTS



http://www.discoversexandgender.ca/

#### FOCUS ON METHODS – The "why" and the "how"

CHECK OUT OUR MEET THE METHODS SERIES

Issue 1 | August 2020

#### **MEET THE METHODS SERIES:**

#### MEASURING AND MANIPULATING SEX HORMONES IN LABORATORY ANIMALS



In basic science, sex hormones can be measured and manipulated in mice and rats using a variety of techniques. The CIHR Institute of Gender and Health asked Margaret McCarthy, Ph.D., Professor of Pharmacology at the University of Maryland, about her views on how to best integrate these methods in her research. Margaret McCarthy's research focuses on the influence of sex hormones on the developing brain, with a special emphasis on understanding the cellular mechanisms that establish sex differences. Here are Margaret McCarthy's recommendations:



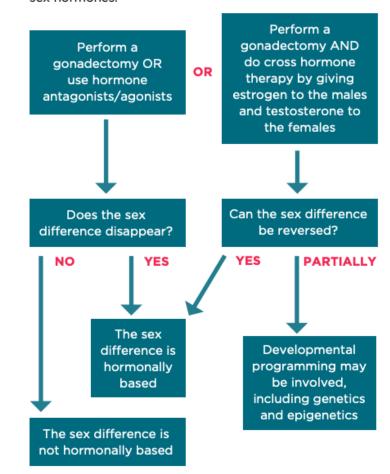
For **males**, seminal vesicle weight can be a good indicator of androgen levels.
The higher the weight, the higher the androgen levels.



For **females**, uterine weight can be used as an indicator of estrogens in non-pregnant mice or rats. The higher the weight, the higher the estrogen levels.

#### What methods do you recommend to manipulate sex hormones in male and female mice and rats?

Once a sex difference is known, there are several methods to determine if the effect is driven by sex hormones:



https://cihr-irsc.gc.ca/e/52107.html

#### HOW MIGHT WE INTEGRATE SEX AND GENDER INTO

# MEDICAL EDUCATION



# CLINICAL PRACTICE





### Alberta medical students explore LGBTQ health issues

'We're talking about how to have an inclusive practice for LGBTQ patients and sexual and gender minorities'

CBC News · Posted: Mar 11, 2017 10:05 PM MT | Last Updated: March 12, 2017



Derek Fehr, U of A medical student who helped organize the conference "We initially started the University of Alberta Inclusive Health Conference because we felt that our medical school curriculum didn't really address any of these issues that we felt were super keen to address these health disparities."



#### University of Alberta Inclusive Health Conference

- Started in 2010
- Led to curriculum changes
- Accredited

#### Yale school of medicine

Dr. Njeri Thande worked with 1st and 2nd year students to change the curriculum of Yale School of Medicine

"To get the best outcomes, we need to practice individualized medicine, which must include consideration of sex and gender."

#### New Opportunities in Medical Education

February 27, 2018



Dr. Njeri Thande (right) is working with WHRY and students such as Nisha Dalvie (left) to integrate a focus on sex and gender into the Yale School of Medicine curriculum.

#### YALE MEDICAL STUDENTS

audited 548 lectures with a survey to track the content of each class, and to critically appraise each topic and research study presented

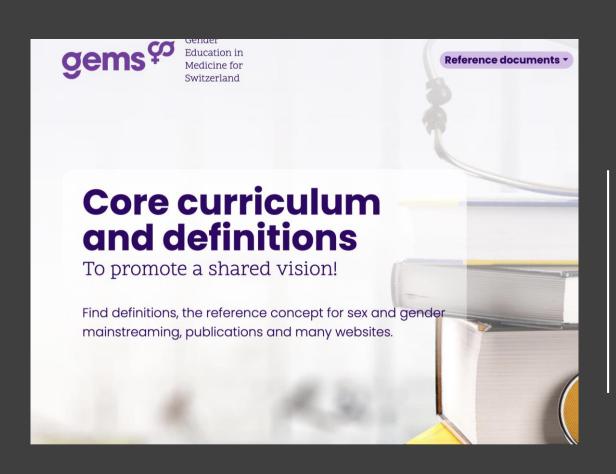
- 1. Is there any mention of the sex or gender of the participants in each topic or study?
- 2. Are the study's results analyzed by sex or gender?
- 3. Does the discussion underscore gaps in sex-specific knowledge?
- 4. Does the discussion mention sex or gender considerations in the presentation or treatment of the condition?

of all sessions raised the topic of sex or gender influences on physiology and pathophysiology or the experience of the patient in the health care environment.

The results informed recommendation s for improving the inclusion of data on sex- and gender-based content.

Thande et al. The Influence of Sex and Gender on Health: How Much Is Being Taught in Medical School Curricula? J Womens Health (Larchmt) 2019 Dec;28(12):1748-1754.

# In Switzerland, universities are working together to integrate sex and gender in the Swiss Medical Curriculum



#### Core curriculum

Reference document for Sex and Gender integration in the Swiss medical curriculum

Working group of the project Sex and Gender integration in the Swiss Medical Curriculum

October 2022

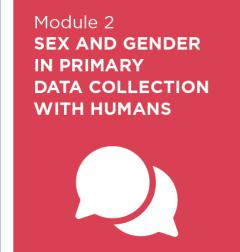
#### Mandatory training for students and faculty

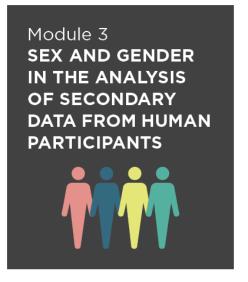
# Advancing next gen science

#### **Training Modules**

Three online sex and gender training modules that many academics credit with building the capacity of the health research enterprise have now been completed by almost 30,000 users. 95% of course completers report an intent to improve their behaviour with respect to sex and gender in health research.







"Everyone in our Faculty of Medicine's graduate school is now required to do one of the IGH sex and gender training modules as part of their professional skills course. This wasn't something that the Faculty of Medicine paid much attention to before."

-IAN GRAHAM, PROFESSOR, UNIVERSITY OF OTTAWA

# RESEARCH PROJECT AWARDS AND RECOGNITION



Best integration of sex or gender

# EXPLORE BIASES, PAY ATTENTION TO LANGUAGE

#### **GUIDELINES FOR INCLUSIVE LANGUAGE FOR SEX AND GENDER DIVERSITY**

#### **CURRENT TERMS**



Mother/Father
Wife/Husband
Boyfriend/Girlfriend
Vagina/Penis
Breastfeeding



"Do you have a husband here or other family I can call?"



"Are you and your boyfriend using protection? Will your husband be here for the baby's ultrasound?"

#### **INCLUSIVE ALTERNATIVES**



Parent
Partner
Significant other
Genitals
Nursing or chest feeding



"Do you have a loved one here or a significant other who I can call?"



"Do you regularly practice safe sex? Will your birthing partner be coming to the ultrasound?"

As recommended by the Canadian Society of Obstetrics and Gynecology Canada



# 6 COGNITIVE STEPS to integrate sex and

gender into clinical

practice

Identify patient sex and gender.

2 Recognize sex differences in clinical manifestation of disease.

E.g., Females more likely to have coronary microvascular disease and to present without substernal chest pain

3 Recognize potential limitations in diagnostic testing.

E.g., Variable prognosis of exercise treadmill test in males versus females.

Use sex-specific thresholds for biomarkers/laboratory value references.

E.g., Troponin, sex-specific EKG criteria for ST-elevation MI

When available, dose medications based upon sex-specific evidence.

E.g., Sex-based dosing of sedative medications or anticoagulants for treatment of myocardial infarction.

6 Mitigate gender and racial bias.

# MITIGATING GENDER BIAS: A SYSTEMATIC APPROACH TO DIAGNOSING & TREATING HEART ATTACKS

Huded et al. (2018) JACC

HISTORIC CONTROL GROUP 490 men 233 women

Usual care for heart attacks allows diagnostic bias among physicians

INTERVENTION GROUP 378 men 171 women

#### **New 4-step protocol**

- 1. ER Criteria do not require cardiology consult
- 2. Guideline based drug therapy despite diagnostic uncertainty
- 3. Immediate transfer to the catheterization lab
- 4. Initial radial approach for vascular access

#### Men vs Women

77% vs 69% guideline therapy 104 vs 112 min door to balloon time

30-day mortality 6.1% higher in women

#### Men vs Women

84% vs 80% guideline therapy 89 vs 91 min door to balloon time

30-day mortality reduced to 3.2% higher in women

# CLASSIFYING LUNG FUNCTION ACCORDING TO GENDER IDENTITY CAN BE IMPRECISE

#### TRANSGENDER WOMAN

SPIROMETRY WITH FEMALE SEX DESIGNATION

range (91.5% of predicted).

Overestimates patient's lung function.



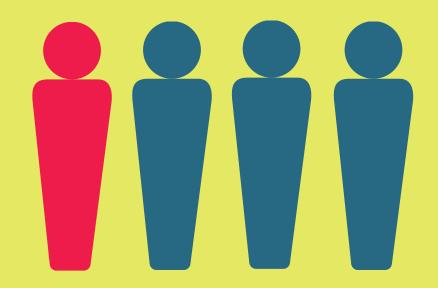
SPIROMETRY WITH MALE SEX DESIGNATION

FEV1 considered **low** (78% of predicted).

Better estimation of patient's true pulmonary function.

Females have lower predicted flow rates and volumes in spirometry. Clinicians must balance appropriate classification of disease severity yet maintain respect in honoring a patient's reassigned gender.

## APPLYING A SEX AND GENDER SCIENCE APPROACH MIGHT HELP US TO INTERVENE AND UNDERSTAND WHY...



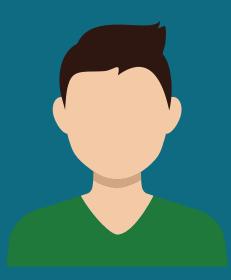
# MEN ACCOUNT FOR THREE OUT OF FOUR DEATHS BY SUICIDE.

# EVERYONE HAS A ROLE TO PLAY IN CHANGING CULTURE

STUDENT OR POST-DOC



FACULTY OR PRINCIPAL INVESTIGATOR



HEALTHCARE PROFESSIONAL



MEDICAL SCHOOL DEAN



Classes & student poster competitions

Thesis defenses

Teaching & conferences

Patient care

Ensure sex and gender content in courses and exams

Culture starts shifting when we change the "norm"



# ACTIONS YOU CAN TAKE NOW

- If you are a student, pay attention and point out if sex and gender are not addressed in lectures, assignments or clinical cases
- If you are a researcher, call out your peers during research presentations if data on sex and gender are not reported and if incorrect terminology is used
- If you are an educator, include sex and gender in your courses and ask if the course curriculum includes a focus on sex and gender science
- If you are a clinician, systematically work through the 6 cognitive steps to account for sex and gender in clinical care to prevent bias.



# ACTIONS YOU CAN TAKE NOW

- If you review for funders or journals, evaluate the appropriate consideration of sex and gender in research design, methods, analysis
- If you are a member of a professional society, ask if the clinical practice guidelines committees pay explicit attention to sex and gender in their recommendations
- Ask questions during a thesis defense or at conferences if data are not disaggregated by sex or take gender into account
- Start a working group or student network in your area for introducing new methods in sex and gender science!

# I INVITE YOU TO BE BOLD AND PUSH BOUNDARIES

# TO BE THE LEADERS OTHERS FOLLOW



