Gender in research webinar

Health, demographic change and wellbeing

15 October 2018









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Basic Concepts

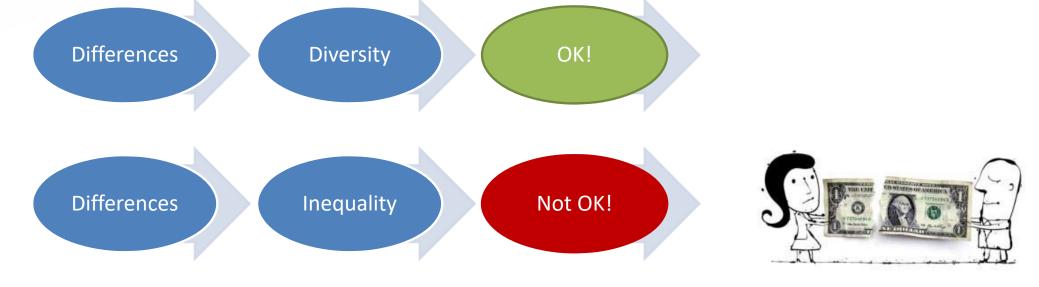
SEX refers to the biologically determined characteristics of men and women in terms of reproductive organs and functions based on chromosomal complement and physiology. As such, sex is globally understood as the classification of living things as male or female.



GENDER refers to the social construction of women and men, of femininity and masculinity, which varies in time and place, and between cultures.

NOTE THAT:

- The problem is not the difference between men and women as such, but the difference in how they are valued
- Certain aspects associated with 'masculinity' still tend to be valued more highly
- The result is inequality of opportunities, segregation & discrimination



GENDER EQUALITY

A situation where individuals of both sexes are free to develop their personal abilities and make choices without the limitations imposed by strict gender roles. The (possibly) different behaviours, aspirations and needs of women and men are considered, valued and favoured equally.

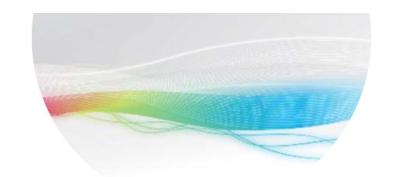


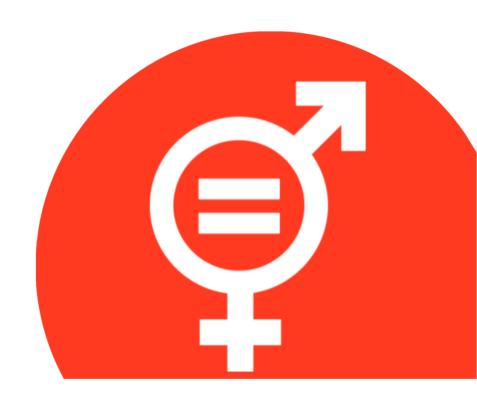


European Commission

Three objectives underpin the European Commission's strategy on gender equality in research and innovation policy:

- Fostering equality in scientific careers
- Ensuring gender balance in decision-making processes and bodies
- Integrating the gender dimension in research and innovation content, i.e. taking into account the biological characteristics and the social features of women and men





Equal
Opportunities
in research at
all levels





Gender and sex variable in the research content

Gender in research

Equal
Opportunities
in research at
all levels

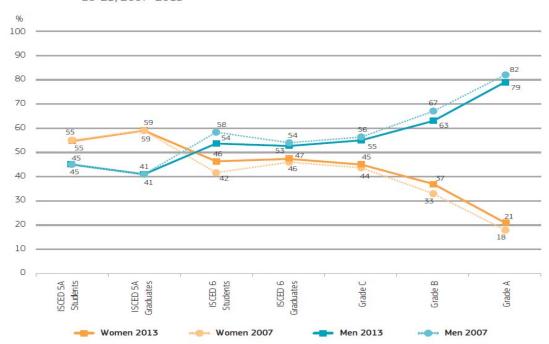
Gender in research

Gender and sex variable in the research content

SHE – figures – 2015: The scissors - diagram

- In only eight out of 28 EU
 Member States did women
 account for more than 40 % of
 researchers.
- Women in the EU have a stronger presence amongst researchers in the higher education and government sectors. In the business enterprise sector, they make up close to one in five researchers (2011)."

Figure 6.1. Proportion of women and men in a typical academic career, students and academic staff, EU-28, 2007–2013



Notes: Reference years Eurostat data: 2007–2012; Reference years for Women in Science (WIS) data: 2007–2013; Exceptions to the reference years (WIS):
AT: 2007–2011; BE (FR), LV, RO: 2010–2013; CY, PT: 2007–2012; DK, LU (Grade A and B, C not available): 2009–2013; ES, IE: 2008–2012; BE (FL), NL, FI: 2011–2013; PL
SK: 2012–2013; FR: 2012; HR: 2014; MT: 2015; EE: 2004 (She Figures 2012); LT: 2007 (She Figures 2012); UK: 2006 (She Figures 2012); Data unavailable for: (Eurostat)
ISCED 5A Students: LU (2007); ISCED 5A Graduates: FR (2012), LU (2007); ISCED 6 Students: DE (2007), LU (2007); ISCED 6 Graduates: FR (2012), LU (2007).

Source: Women in Science database, DG Research and Innovation and Eurostat – Education Statistics (online data code: educ_grad5)

The gap is even bigger if we look at the proportion of women and men in the areas of science and engineering.



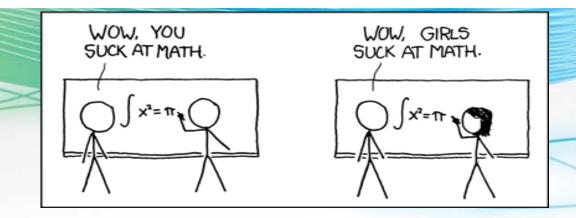


Figure 6.2. Proportions of women and men in a typical academic career in science and engineering, students and academic staff, EU-28, 2007–2013



Notes: Reference year for Eurostat data: 2007–2012; Reference year for WiS data: 2007–2013; Exceptions to the reference years (WiS): AT: 2007–2011; BE (FR): 2010–2013; BE (FL), NL, FI: 2011–2013; CZ: 2007–2008; DK: 2009–2013; IE: 2008–2012; CY, PT: 2007–2012; EL, MK: 2012; PL, SK: 2012–2013; BA, SI: 2013; HR: 2014, LT: 2007 (She Figures 2012); UK: 2006 (She Figures 2012); Data unavailable for: WiS Grade A, B and C: AT, BG, EE, FR, HU, LU, LV, RO; Eurostat: ISCED SA Students: LU (2007), ISCED SA Graduates: FR (2012), LU (2007), LU (2007), LU (2007), NL (2007), ISCED 6 Graduates: FR (2012), IT (2007), LU (2007), PL (2012);

Others: SET fields of education = Science, maths and computing + Engineering, manufacturing and construction; SET fields of science = Engineering and technology + Natural sciences.

Source: Women in Science database, DG Research and Innovation and Eurostat - Education Statistics (online data code: educ_grad5)

Gender Equality Plan

As defined by the European Commission, a gender equality plan consists of a set of actions aiming at:

- Conducting impact assessment / audits of procedures and practices to identify gender bias.
- Identifying and implementing innovative strategies to correct any bias.
- Setting targets and monitoring progress via indicators.



European Commission Communication on 'A Reinforced European Research Area Partnership for Excellence and Growth' (COM(2012) 392 final)

Good practice examples - Areas of intervention:

- Organisational culture:
 - ✓ Organise gender training
- Reconciliation of work and private life:
 - ✓ Measures to facilitate return to work after parental leave
- Recruitment, selection and career progression:
 - ✓ Organise unconscious bias awareness sessions
- Leadership and decision-making:
 - ✓ Gender quota in all decision making bodies
- Sexual and gender-based harassment:
 - √ 'Special Contact Person' for sexual harassment



→ See GEAR tool: http://eige.europa.eu/gender-mainstreaming/toolkits/gear

Equal
Opportunities
in research at
all levels



Gender and sex variable in the research content

Gender in research

Ideas phase

Dissemin ation phase

Gender in research content

Proposal phase

Remember:
Both the variables
sex AND gender can
be relevant

Research phase

- Make 'gender' visible straight away (e.g. in abstract, key words)
- Budget: foresee resources; remember gender training is an eligible cost (budget for training under 'other direct costs')
- Keep 'gender' in mind <u>throughout</u> the proposal preparation and drafting (gender balance in team; management structures; expertise in the consortium; research activities;...)
- → No 'magic formula' or couple of paragraphs
- → No 'excellence' without gender equality!
- → Mobilise expertise

Technical part of the proposal:

1. Excellence:

- 1.1: Objectives: point out relevance; include analysis of sex / gender in relation to the main research topic as objective; explain which knowledge exists already and which are the gaps the research will fill
- 1.2 Relation to the Work Programme: especially when gender is flagged → explain how furthering gender knowledge will help advance the WP objectives

1.3 Concept and Method:

- a) Explain / show the gender expertise in the consortium (interdisciplinary research!), and if missing, say how this will be solved. Refer to existing research on sex/gender in relation to the topic and explain how the project will build on the existing research (if relevant)
- b) Explain the project's approach to sex / gender throughout the research cycle
- 1.4 Ambition: include also a reflection on what the ambition of the project is in relation to gender knowledge

Technical part of the proposal:

2. Impact:

2.1 Expected impacts: include gender! Point out any obstacles or barriers, e.g. missing sex-disaggregated data \rightarrow explain how the project will contribute to solving this obstacle

2.2 Maximise impact

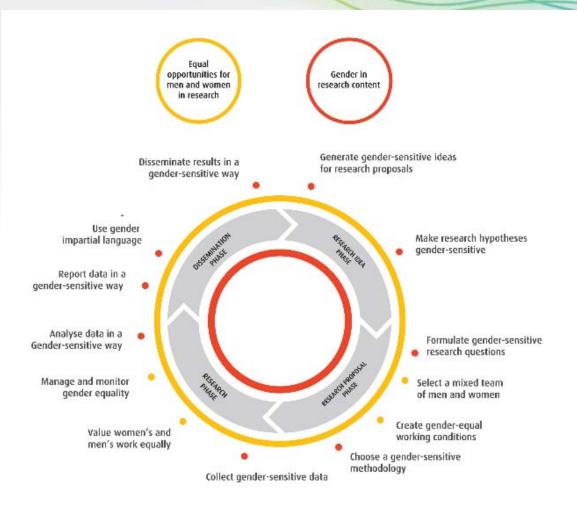
- a) dissemination and exploitation: be consistent and integrate also sex/gender findings in how exploitation is planned; show what the added value will be; how including sex/gender variable will raise the quality of the research
- b) communication: communicate findings! (conference papers; posters; research articles); show how results will be disseminated in a way that makes the sex/gender variable visible

Technical part of the proposal:

3. Implementation

- 3.1: Work Plan: WP's and deliverables: integrate sex / gender throughout; show how the variables sex and/or gender will be taken on board; involve/consult relevant stakeholder groups and experts; consider separate deliverable on gender issues; present Gender Equality Plan in Management work package
- 3.2 Management structures: ensure gender balance in management structures!
- 3.3 Consortium as a whole: ensure and point out gender balance and gender expertise
- 3.4 Resources: gender training to be foreseen; sufficient resources for gender issues in the work plan

Tool: checklist, in https://www.yellowwindow.com/genderinresearch





How to make research gender-sensitive

CHECKLIST FOR GENDER IN RESEARCH

Equal opportunities for women and men in research

	is there a gender balance in the project consortium and team, at all levels and in decision-making posi- tions?
	Do working conditions allow all members of staff to combine work and family life in a satisfactory manner?
	Are there mechanisms in place to manage and monitor gender equality aspects, e.g. workforce statistics, as required by FP7?
	Gender in research content
	Research Ideas phase:
	if the research involves humans as research objects, has the relevance of gender to the research topic been analysed?
	If the research does not directly involve humans, are the possibly differentiated relations of men and women to the research subject sufficiently clear?
	Have you reviewed literature and other sources relating to gender differences in the research field?
	Proposal phase:
	Does the methodology ensure that (possible) gender differences will be investigated: that sex/gender- differentiated data will be collected and analysed throughout the research cycle and will be part of the final publication?
0	Does the proposal explicitly and comprehensively explain how gender issues will be handled (e.g. in a specific work package)?
	Have possibly differentiated outcomes and impacts of the research on women and men been considered?
	Research phase:
	Are questionnalities, surveys, focus groups, etc. designed to unravel potentially relevant sex and/or gender differences in your data?
	Are the groups involved in the project (e.g. samples, testing groups) gender-balanced? Is data analysed according to the sex variable? Are other relevant variables analysed with respect to sex?
	Dissemination phase:
0	Do analyses present statistics, tables, figures and descriptions that focus on the relevant gender differences that came up in the course of the project?
	Are institutions, departments and journals that focus on gender included among the taxest organization, along with mainsteam research magazines?



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Health, demographic change and wellbeing

15 October 2018 Prof. Dr. Vera Regitz-Zagrosek







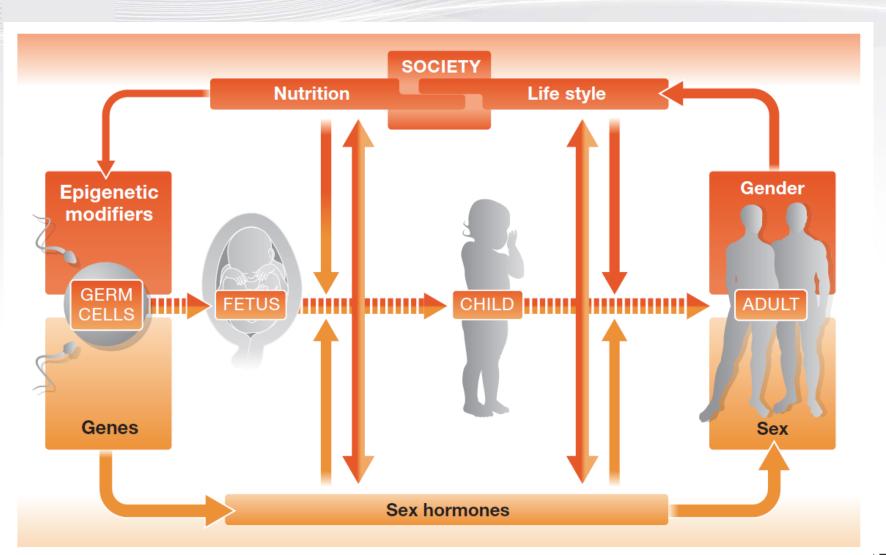
The topics of the call

Call - Better Health and care, economic growth and sustainable health
systems9
1.1 Personalised medicine9
SC1-BHC-01-2019: Understanding causative mechanisms in co- and multimorbidities
combining mental and non-mental disorders10
SC1-BHC-02-2019: Systems approaches for the discovery of combinatorial therapies for
complex disorders11
SC1-BHC-03-2018: Exploiting research outcomes and application potential of the human
microbiome for personalised prediction, prevention and treatment of disease
SC1-BHC-04-2018: Rare Disease European Joint Programme Cofund
SC1-BHC-05-2018: International flagship collaboration with Canada for human data
storage, integration and sharing to enable personalised medicine approaches
SC1-BHC-30-2019: Towards risk-based screening strategies for non-communicable
diseases

Topics of my presentation

- The interaction of sex and gender in biomedicine, its contribution to health and disease across different disease entities – as a first step towards personalized medicine
- How to measure the socicocultural dimension gender as risk factor and predictor in personalized medicine approaches
- Drug development: Sex-related differences in basic research and drug development
- Sex- and gender-related differences in frequent NCD, eg cardiovascular disease, in risk factors and comorbidities,

Sex and gender contribute to disease development



How to measure gender

The generally accepted view (eg CIHR)

Gender roles Behavioral norms men/women in society, expectations

context: institutions (e.g. family), labor force, educational system

Gender identity How we see ourselves as female/male (or as third gender)

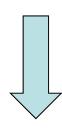
Gender relations How we interact with or are treated by other people

(based on our ascribed gender)

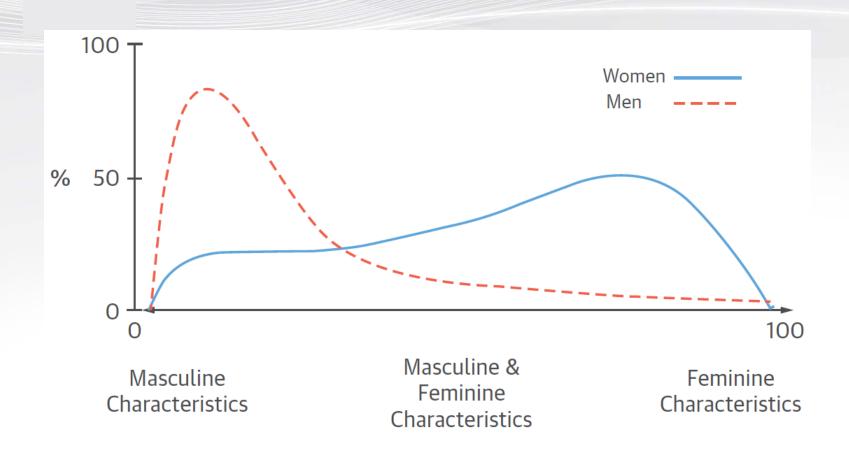
Institutional gender Reflects the distribution of power between men and women in the

political, educational, and social institutions in society. The institutionalized aspect of gender also shapes social norms that define, reproduce, and often justify different expectations and

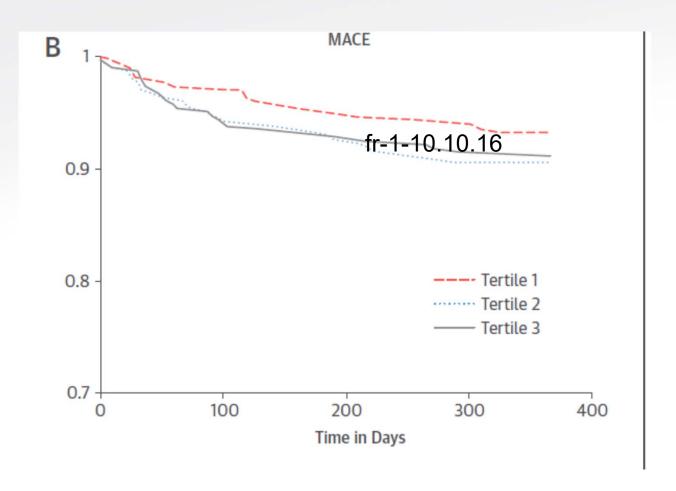
opportunities for men and women.



Gender score distribution in men and women with premature acute coronary syndrome



Gender is a strong predictor of major cardiovascular events after acute coronary syndromes



Some biological risk factors are more closely associated with gender than with sex

Pelletier et al., 2015, JACC 2016



Gender predicts the outcome after acute coronary syndromes better than biological sex

In the genesis Praxy study, 1024 patients with ACS (50 % women) were followed for at least 12 months.

Female gender score was associated with re-events whereas female biological sex was protective.

TABLE 2 Multivariable Cox Proportional Hazards Regressions: Association Between the Gender-Related Score and Recurrent ACS					
	Recurrent ACS HR (05% CI)	p Value			
Gender-related score (from score 0 to 100)	4.50 (1.05-19.27)	0.04			
Female sex	0.50 (0.18-1.40)	0.18			
Age	1.01 (0.94-1.10)	0.77			
GRACE score, for each point increment	1.00 (0.97-1.02)	0.70			
Previous CV event	2.13 (0.94-4.80)	0.07			
Number of CV risk factors	1.02 (0.77-1.35)	0.92			
$\label{eq:acute} \begin{split} \text{ACS} = & \text{acute coronary syndrome; CI} = \text{confidence interval; HR} = \text{hazard ratio; other} \\ & \text{abbreviations as in Table 1.} \end{split}$					

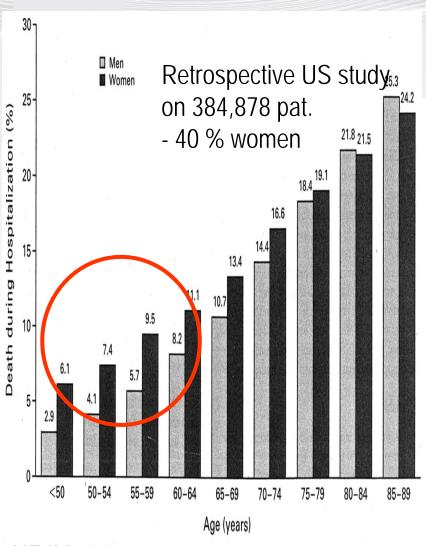


Sex and Gender differences in presentation of myocardial infarction

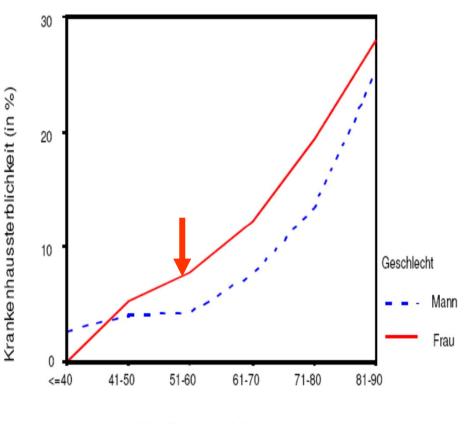


one in four women realize it is a greater th reat than cancer • A woman has a substrahlung bei Anglina pector a man • Of those who survive their first he within a year, vs. 25% of men • 46% of after a heart attack, compared with 22% of men

Sex and Gender in cardiovascular disease - Higher early mortality in women with myocardial infarction



Mortality after MI 2005 in Berlin, 5000 patients



NEJM 1999

Altersklassen in Jahren

Are there differences in cardiovascular disease mechanisms in women and men?

Classical CAD:

Large coronary artery disease

Men

Atherosclerosis

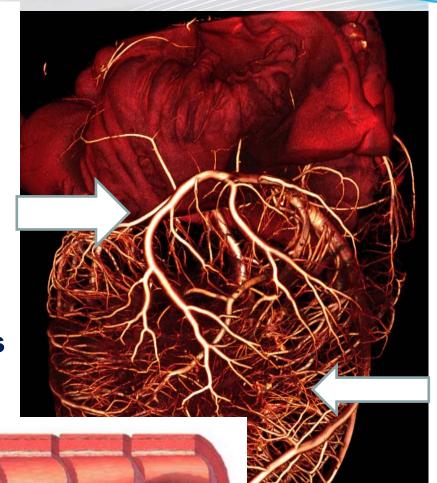
Fatty

streak

Plaques

Normal

vessel



Advanced/vulnerable

plaque

Fibrofatty

plaque

Women

- Spasms
- Dissections
- Dysfunction

Microvascular disease

 Heavily understudied, specific research needed

Gender related problem: Women with myocardial infarction arrive later in hospital than men

	Countr y	Conditio n	Intervall: medical contact – hospital admission	Hospital intervention			
í	France, all	STEMI	Men 200	Men 200 / Women 245			
	Fr, Brittany	STEMI	N	Leurent, 2014			
	Spain	STEMI	N.	M 240 / W 307			
	NL	STEMI	<65 y: Men: 150 min / Women 165	Men=women	Otten, 2013		
	DE	Stemi	M154 / W 189		Ladwig , 2009		
	SW	STEMI	M169 / W 190	Von 51/80 auf 43 / 48 nach intervention	Naegele 2011, rad 2012		
	US	STEMI	M150 / W 195				
	Australia	STEMI	M161 / W 217				



Patient-physician gender concordance and increased mortality among female heart attack patients

Brad N. Greenwood^{a,1}, Seth Carnahan^b, and Laura Huang^c

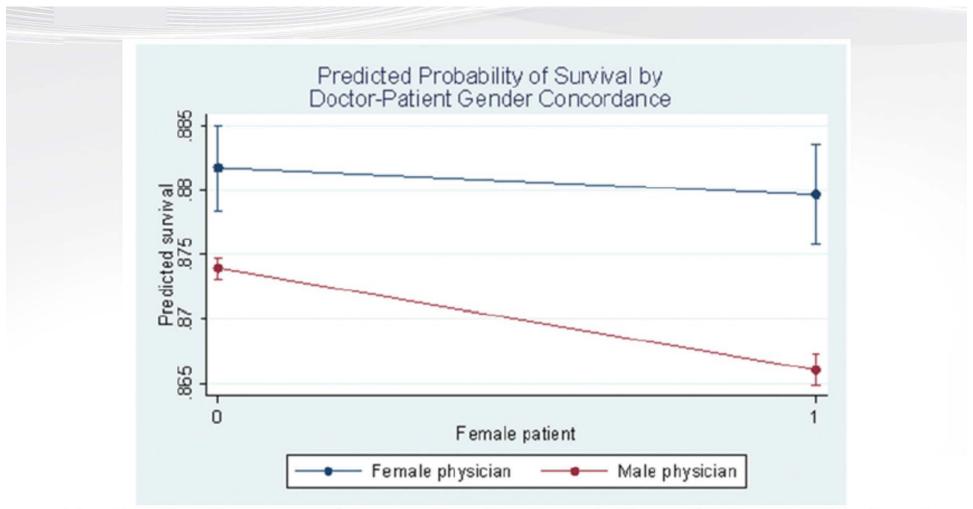


Fig. 1. Gender concordance and patient survival: results from Table 2, column 3, 90% confidence interval displayed. Estimates include controls and hospital quarter fixed effects. Covariates held at sample means. n = 581.797.

Sex and gender differences in the risk factors and comorbidities of NCD

https://gender.charite.de/en/research/projects_with_the_eu/gencad/



FACTSHEET for healthcare professionals GENCAD

Gender differences in coronary artery disease in Europe

- Socioeconomic factors
- Depression and stress
- Rheumatic diseases
- Preeclampsia
- Genetic factors, sudden death
- Erectile dysfunction
- Menopause, Andropause, Hypogonadism

Gender differences in classical risk factors

- Coronary heart disease develops 7-10 years later in women compared to men. Overall, more women are affected, since they live longer and the disease develops in old age.¹
- Hypertension occurs more frequently in men before the age of 50, and in women after the age of 50. Hypertension leads to more strokes and heart failure in women than in men.²³
- Diabetes increases the risk of cardiovascular disease more in women than in men.^{4,5} Women with diabetes and associated risk factors are high risk patients, and need intense management and treatment.⁶
- Dyslipidemia is an equally strong risk factor in women and in men. Lipid lowering therapy should therefore be used in both.¹
- Smoking is a relatively greater risk factor in younger women than in men and smoking rate in women has been reduced less than in men.⁷

- Age
- Hypertension
- Diabetes
- Smoking



Gender differences in non-classical risk factors

- Poor socioeconomic status contributes to gender disparities in cardiovascular health ^{8,9}
- Depression and sustained mental stress occur more often in women than in men and are more important risk factors in women. ^{10,11}
- Autoimmune and rheumatic diseases occur more often in women and are frequently associated with cardiovascular disease.¹²
- Preeclampsia is an increasingly recognized risk factor in women.¹³
 Women who develop preeclampsia have a twofold elevated risk of developing cardiovascular disease later in life.¹⁴⁻¹⁷
- Genetic factors are important in premature heart disease in women and in men 18
- Erectile Dysfunction is associated with general metabolic and cardiovascular health risks in men.¹⁹
- Menopause, polycystic ovary syndrome, andropause and hypogonadismus are associated with increased cardiovascular risk in women and man 2, 13, 19, 20, 21

- Socioeconomic status
- Mental stress
- Autoimmune & rheumatic diseases
- Preeclampsia
- Sex hormones



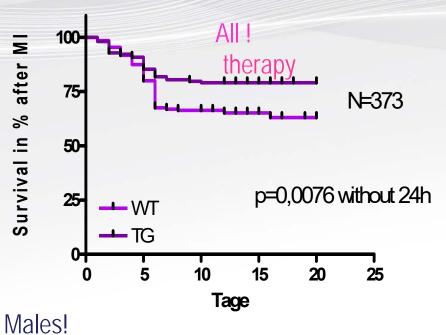
Gender-specific mechanisms of disease

- In men, arteriosclerosis of large coronary arteries is the dominant mechanism leading to myocardial ischemia and infarction.²²
- Middle-aged women frequently have angina pectoris and myocardial ischemia in the presence of normal coronary arteries. The term ischemia hand disease is critically for this form of disease 23-25.
- Large coronary artery arterianglerania
- Ischemic heart disease
- Non-obstructive functional coronary artery disease
- Stress-induced heart dis-

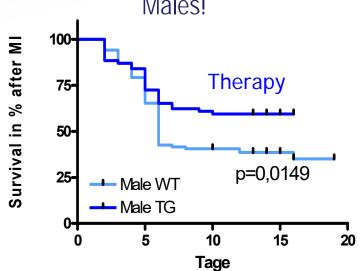
Underestimation of the role of biological sex in basic research and drug development

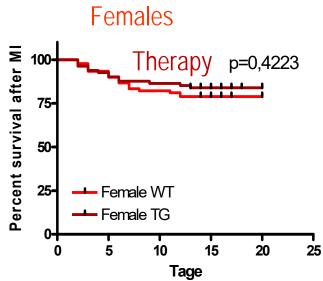


Drug development should be done in male and female mice - example: survival after MI!



Improvement is seen in the whole cohort and in males, not in females.





Sex Bias Is Increasingly Prevalent in Preclinical **Cardiovascular Research: Implications for** Translational Medicine and Health Equity for Women

A Systematic Assessment of Leading Cardiovascular Journals





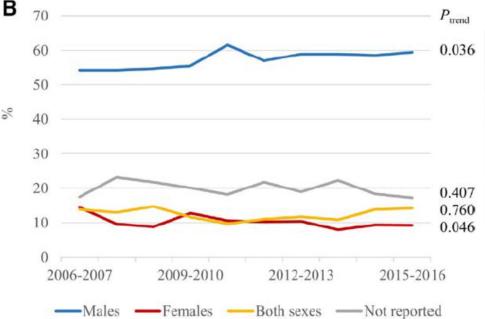


Figure. Temporal patterns in the sex of animals used in preclinical cardiovascular research over a 10-year

Sex and gender differences occur across human diseases

Table 1 | Publications with sex and gender differences in the most frequent clinical entities

Cardiology	Rheumatology Immunology	/ Pneumology	Nephrology	Gastro- enterology/ Hepatology	Neurology	Endo- crinology	Oncology	Haematology
Hypertension (414)	Lupus erythematosus (68)	Asthma (140)	Renal failure (27)	Hepatitis B (22)	Multiple sclerosis (65)	Diabetes mellitus (447)	Skin carcinoma (45)	Anaemia (44)
Myocardial infarction (275)	Rheumatoid arthritis (41)	Lung cancer (116)	Diabetic nephropathy (11)	Hepatitis C (26)	Stroke (129)	Obesity (349)	Gastric cancer (25)	Leukaemia (49)
Heart failure (153)	Systemic sclerosis (3)	Chronic obstructive pulmonary disease (36)	Glomerulone- -nephritis (9)	Hepato-cellular carcinoma (37)	Alzheimer's disease (104)	Osteo- porosis (123)	Renal cell carcinoma (17)	Lymphoma (34)
Atrial fibrillation (38)	Fibromyalgia (15)	Pulmonary hypertension (12)	Polycystic kidney disease (12)	Inflammatory bowel disease (13)	Epilepsy (56)	Hypo- thyreoidsm (33)	Bladder cancer (22)	Thrombo- cytopoenia (6)
Coronary heart disease (207)	Sjögren's syndrome	Pulmonary embolism (110)	Renal artery stenosis (0)	Colorectoral cancer (24)	Parkinson's disease (69)	Hyper- thyreoidsm (16)	Thyroid carcinoma (16)	Purpura (2)
Cardiomyopathy (41)	Ankylosing sponylitis (11)	Sarcoidosis (6)	IgA Nephropathy (2)	Autoimmune Hepatitis (2)	Muscular dystrophy (11)	Morbus Addison/ Cushing disease (5)	Pancreatic carcinoma (10)	Agranulo- cytosis (0)

EMBO reports advance online publication XX Month 2012; doi:10.1038/embor.2012.87

Summary and conclusions

- The inclusion of sex and gender in biomedicine is a first step towards personalized medicine
- The socicocultural dimension gender is a measurable risk factor and predictor of adverse events and should be included in clinical trials
- Sex-related analysis provide significant insights and improvement in basic research and drug development
- Sex- and gender-related differences play a role in NCD, in risk factors and comorbidities,

Thank you for attending this webinar

For background information, some resources, reading list → see the 'hand-out' that you will receive.