## A framework for the Analysis of Gender Bias in research and medical care

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## "Normality is believing that what happens is normal"

 (El Roto, Spanish humorist)LA NORMALIDAD CONSISTE EN CREER QUE LO


## Definitions of Gender Bias

## Medical practice:

"The differential medical treatment of men and women, the impact of which may be positive, negative or neutral."

Lenhart Sh. Gender discrimination: A health and career development problem for women physicians. J Am Med Women Assoc 1993; 48: 155-9.

## Research purposes:

"A systematic, erroneous gender dependent approach related to social construct, which erroneously regards women and men as similar or different, in the exposure to risk factors and the natural history of disease."

## Coronary Angiographs between women and men

Harvard Hospital (30,000 patients) and New Haven Hospital (30,000 patients)
Ayanian IZ, Epstein AM. N Engl J Med 1991; 325: 221-5
№ of coronary angiographs per 100 patient admissions
Men
Women
Whites
18.8 \%
9.4\%
Non Whites
14 \%
8.6\%

## 2011 10Q Report:

Advancing Women's Heart Health through Improved Research, Diagnosis and Treatment


June 2011 • Washington, DC

## Presentation Focus

1. Gender bias in research is due to an incorrect assumption of:

Equality between women and men
Exposure to risk factors and their consequences.

Early (and even later symptoms) of diseases suffered by both sexes.

Differences between women and men
Responses to treatment

Outcomes/Prognosis
2. Gender bias in research specifically related to women's health problems.

MT Ruiz-Cantero, LM Verbrugge. A two way view of gender bias in Medicine. J Epidemiol Comm Health 1997; 51: 106-9 MT Ruiz-Cantero et al. A framework to analyse gender bias in epidemiological research. J. Epidemiol Comm Health 2007; 61: 46-53

## Incorrect assumption of Equality between women and men in research

|  | Epidemiological bias relating to gender insensitivity |  |
| :--- | :--- | :--- |
| Research context | Selection Bias | Measurement Bias |
| Clinical | Under-representation of women <br> in clinical trials. See slide 8 |  |
| Social |  |  |

## Participation by gender in the published clinical trials of specific pharmaceuticals

Antiretrovirals ${ }^{1}$

${ }^{1}$ Ruiz Cantero MT, Pardo MA. European Medicines Agency policies for clinical trials leave women unprotected. J Epidemiol Comm Health 2006;60:911-3.

## Etoricoxib ${ }^{2}$


${ }^{2}$ Chilet-Rosell E, Ruiz-Cantero MT, Horga JF. Women's health and gender-based clinical trials on etoricoxib:methodological gender bias. J Public Health (Oxf) 2009; 31: 434-45

## Incorrect assumption of Equality between women and men in research

|  | Epidemiological bias relating to gender insensitivity: |  |
| :--- | :--- | :--- |
| Research context | Selection Bias | Measurement Bias |
| Clinical | Under-representation of women <br> in clinical trials. See slide 8 | Lack of sex stratification in <br> clinical trials. <br> Why? Overall sample size <br> stratified by sex is too small <br> to produce valid results |
| Social | Under-representation of women in <br> research on exposure to <br> chemical risks | Familism Bias |

## Incorrect assumption of Differences between women and men in research

|  | Epidemiological bias relating to gender insensitivity |  |
| :--- | :--- | :--- |
| Research context | Selection Bias | Measurement Bias |
| Clinical | Example: Lower Ferritin values <br> for women incorrectly <br> accepted as normal <br> See slide 11 |  |
| Social |  |  |

Reference Population values vs. Accepted Normal values of iron storage protein Ferritin: 50-200 ng $/ \mathrm{ml}$.

Table 1 Reference values of ferritin in public and private health centres in Spain

| Public and private health centres | Reference values <br> $(\mathrm{ng} / \mathrm{ml})$ | Women | Men |
| :--- | :---: | :---: | :---: |
| Cliniall Laboratory, Cornellá-Earcelona, Catalonia | $30-300$ |  |  |
| Basurto Hospital, Vasco Country | $15-150$ |  |  |
| Laboratory Echevame, Granada, Andalusia | $10-204$ |  |  |
| San Juan de Dios Hospital, Barcelona, Catalonia | $10-120$ |  |  |
| La Plana Hospital, Vila-Real Valencian Community | $5-150$ | $5-150$ | $20-200$ |
| Unilabs, Barcelona, Catalonia |  | $5-140$ | $29-280$ |
| Hermanos Miralles Primary Care Centre, Madrid |  | $14-150$ | $40-340$ |
| Folguera Laboratory, Barcelona, Catalonia |  | $15-150$ | $30-400$ |
| Fornells, Oló, Crespo Laboratory, Barcelona, Catalonia | $20-200$ | $20-450$ |  |

1. Inconsistency of the prevalence of anemia across the Spanish regions
2. Unequal access to treatment based on the patient's geographical location

## Incorrect assumption of Differences between women and men in research

$\left.\begin{array}{|l|l|l|}\hline & \text { Epidemiological bias relating to gender insensitivity } \\ \hline \text { Research context } & \text { Selection Bias } & \text { Measurement Bias } \\ \hline \text { Clinical } & \begin{array}{c}\text { Example: Lower Ferritin values } \\ \text { for women incorrectly } \\ \text { accepted as normal }\end{array} & \begin{array}{c}\text { Example: When some studies } \\ \text { wrongly attribute the blame } \\ \text { to woman patients for the } \\ \text { delay in receiving diagnostic } \\ \text { and treatment }\end{array} \\ \text { See slide 13 }\end{array}\right\}$

## Delay Timeline in receiving diagnostic and treatment



## Incorrect assumption of Differences between women and men in research

|  | Epidemiological bias relating to gender insensitivity: |  |
| :---: | :---: | :---: |
| Research context | Selection Bias | Measurement Bias |
| Clinical | Example: Lower Ferritin values for women incorrectly accepted as normal <br> See slide 11 | Example: Some studies wrongly attribute the blame to woman patients for the delay in receiving diagnostic and treatment <br> See slide 13 |
| Social | Example: Studies on eating disorders, which tend to exclusively select women for participation in such research | Example: The myth of a higher prevalence of mental health problems in women than in men |

## Gender bias in research and medical care specifically related to women's health problems

1. Hormone Replacement Therapy (HRT)
2. "Ectomies": Hysterectomies, preventive mastectomies, episiotomy, oophorectomies
3. Aggressive treatments of breast cancer
4. Disease Mongering more frequently aimed at women than men

## Hormonal Replacement Therapy and Related Diseases

| Estrogen <br> Endometrial Cancer <br> $N$ Engl J Med | New restrictive <br> Recommendation from <br> the Spanish Drug |
| :--- | :--- |
| Administration |  |

Commercial Success elixir of youth!

Women's Health Initiative. JAMA:
Excess incidences of cancer and cardiovascular diseases in women

## CONCLUDING REMARKS RECOMMENDATIONS

From the feminist empiricist perspective, we should "Engender" knowledge about disease suffered by both sexes by:

Developing Systematic Reviews

Analysing research limitations

- Explicitly stated by the authors
- Other limitations that can be observed in the studies
"Looking-twice" at the results, stratifying them by sex; and developing Meta-analysis

Requesting how the cut-off points criteria relating to normality/abnormality were established for women in different diagnostic tests :

- Are they established from a sample size of women?
- Are they established from a sample size of men and inferred to be applicable to women?

Evaluating the effectiveness of diagnostic protocols because many have been established for CT's and may not include some women's symptoms/illness, taken as "atypical"

## MAKE VISIBLE THE INVISIBLE. THAT'S ART



