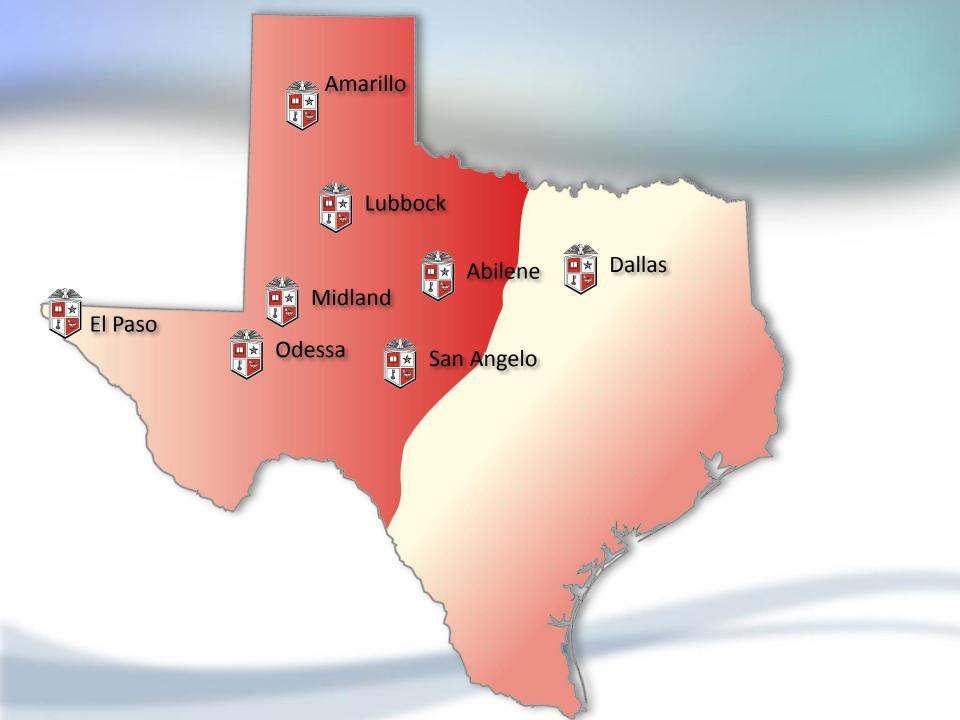
Gender Health Disparities and Clinical Implications

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Health Disparities: Definition

"Health disparities are differences in the incidence, prevalence, mortality, burden of diseases and other adverse health conditions or outcomes that exist among specific population groups in the United States."

National Institutes of Health, 1999

Basis of Health Disparities

- Gender
- Age
- Ethnicity
- Socioeconomic status
- Geography
- Sexual orientation
- Disability or special health care needs

Determinants of Health Disparities

- Natural, biological variation
- Inadequate access to essential health and other basic services
- Transient health advantage of one over another (first adopters of health promoting behavior)
- Health damaging behavior: no restriction on lifestyle choices
- Health damaging behavior: degree of lifestyle choices is severely restricted
- Exposure to unhealthy, stressful living and working conditions
- Natural selection (health related mobility) Sick people tend to move down the social scale.

Gender-Specific medicine the *science* of how normal function and the experience of disease is influenced by the sex of the patient.



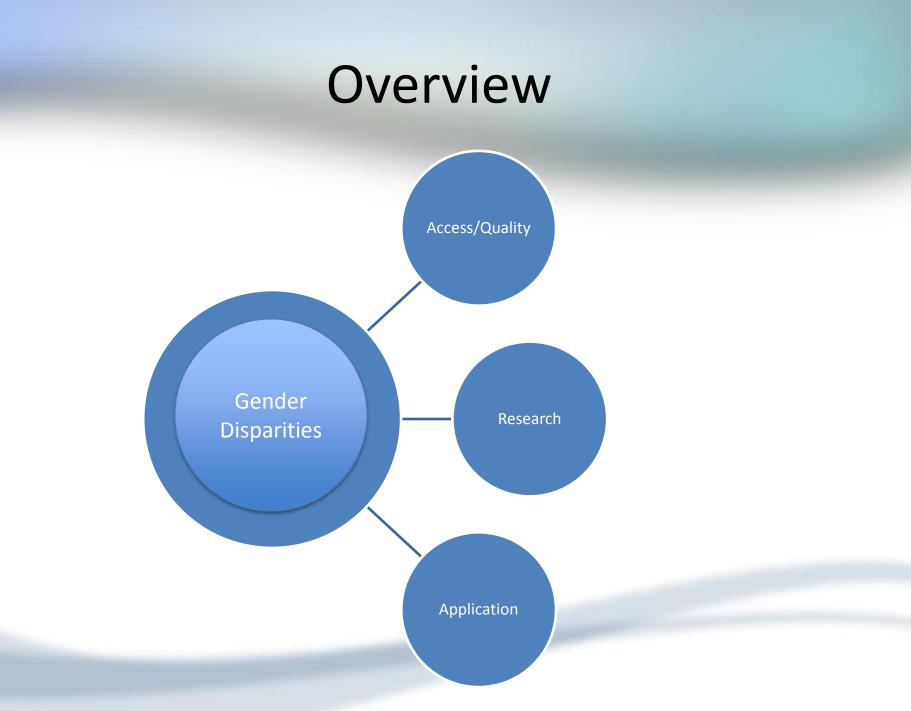
Gender and Sex: Definition

- Gender: introduced in the 1970s as an alternative to 'sex'
- Gender: a social construct
- Sex: a biological term

Sex, Gender & Health

- Explanations for many population and individual level health outcomes are not attributable to biology
- Income, income inequality, social connectedness, and social capital all show some association with health and illness

Sex and gender health association is a complex intertwined relationship



Access & Quality

Socioeconomic Disparity

- Below 100% of the Poverty Level (19-64 yo)
 - Women: 17%
 - Men: 13%
- Median earnings (>15 yo)
 - Women: \$22,224
 - Men: \$32,486
- Women earn 76.5 cents/\$1.00 compared to men

Figure 7. Women Are More Likely Than Men to Have Access Problems in Past Year Because of Cost

Percent of adults ages 19–64 reporting the following problems in past year because of cost

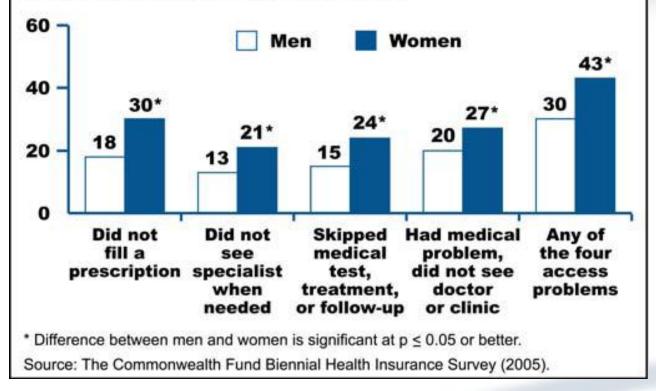
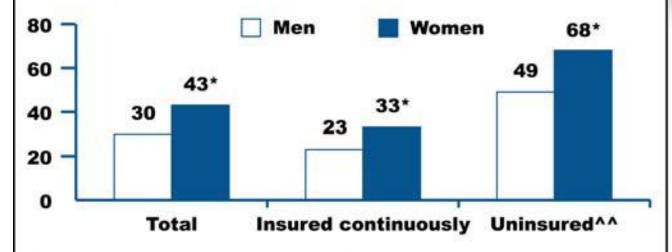


Figure 6. Women Are More Likely Than Men to Have Cost-Related Access Barriers

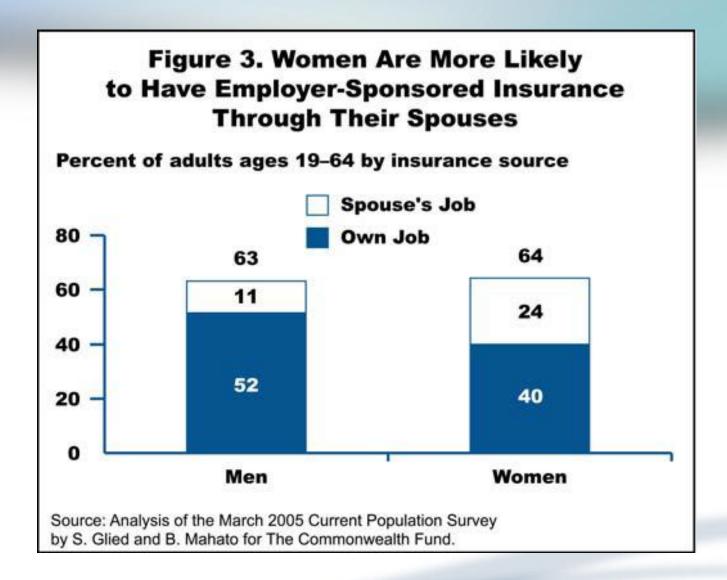
Percent of adults ages 19–64 who have difficulty accessing health care[^]

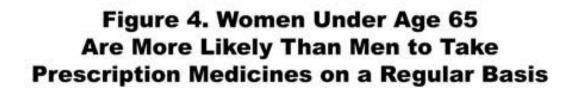


Difference between men and women is significant at p ≤ 0.05 or better.

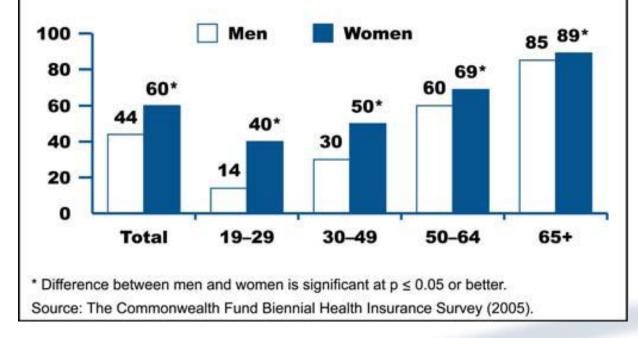
^ Did not fill a prescription; did not see a specialist when needed; skipped recommended medical test, treatment, or follow-up; had a medical problem but did not visit doctor or clinic. ^^ Uninsured combines currently uninsured and currently insured but had a time uninsured in the past 12 months.

Source: The Commonwealth Fund Biennial Health Insurance Survey (2005).





Percent of adults who take prescription medicines on a regular basis



Gender and Health Care Access

Consumer Issue	Gender Majority
Consumers of Health Care	Women
Part-time employment	Women
Preventative Services	Women
Lower Income	Women
Higher out-of-pocket expenses	Women
Coverage through spouse	Women

Connection to the System

• In general, women

- " are more likely to have a usual primary care provider than men" (Agency for Healthcare Research and Quality, 2004)
- " are more likely to enter treatment through primary care [as opposed to emergency services]" (Hauenstein et al., 2006)
- with a connection to the health care system are more likely to have their mental health needs met" (Sherbourne et al., 2001)
- may have fragmented care

• In general, men

- use less medical services
- have less frequent preventative ambulatory visits
- tend to wait until there is an emergent need

Measurable Outcomes

HEDIS: The nation's most widely used tool of health care quality measurement, developed and maintained by NCQA

Gender Difference

Cholesterol

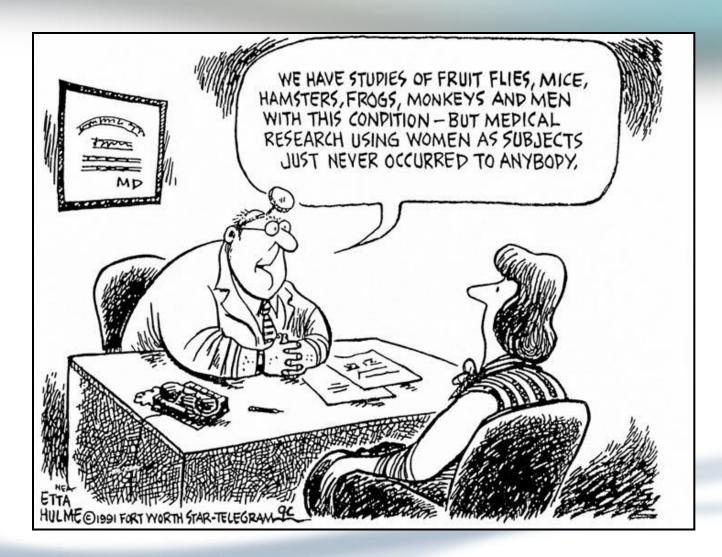
No Gender Difference

- Blood Pressure
- HgA1C

National Committee for Quality Assurance (NCQA), 2005

Research

Clinical Research in Women



Formation of "Women's Health"

• 1980-1990's

- Public Health Services Task Force on Women's Health
- Creation of the Office of Research on Women's Health
- FDA reversed its 1977 edict excluding women from clinical trials
- National Institutes of Health Revitalisation Act 1993
 - 1998 65% of NIH studies included women

Institute of Medicine Report: "Does Sex Matter" (2001)

- Scientists had made the assumption without confirmatory testing and that what is learned from studying men could be extrapolated to women without modification.
- At the time researchers did not report gender analysis, perform gender sub-analysis, or power studies to assess the impact of gender
- 2/3 of all research on diseases that affected both sexes had been performed exclusively in men

Cardiovascular Disease

AHA Guidelines

Evidence-Based Guidelines for Cardiovascular Disease Prevention in Women

Expert Panel/Writing Group*

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Mosca, L et al Circulation 2007

Hypertension

31 studies (103,268 men and 87,349 women)

Major Drug Classes Included

ACE Inhibitors Angiotensin-receptor Blockers B-Blockers Calcium Channel Blockers Diuretics

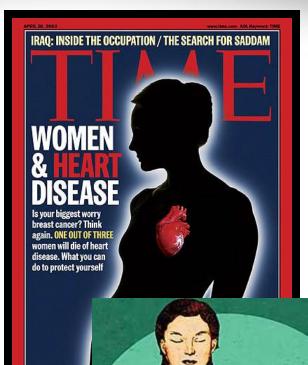
Major Outcomes

Non-fatal stroke or death from cerebrovascular disease Non-fatal myocardial infarction or deaths from CHD Heart failure causing death or requiring hospitalization Total major cardiovascular events Total cardiovascular deaths Total mortality

Achieved blood pressure reductions were comparable for both men and women.

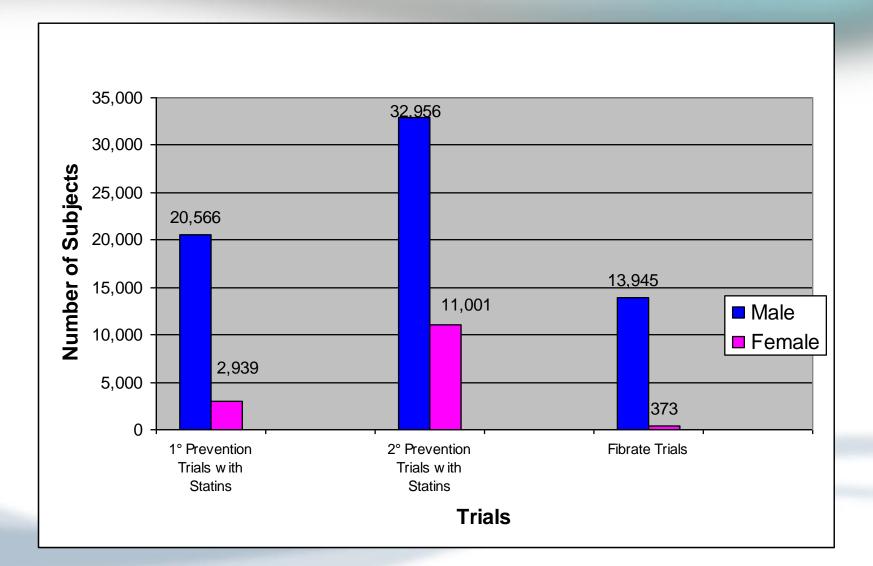
Turnbull, F. Woodward M, Neal B. et al. European Heart Journal. 2008;29(21):2669-2680.

Lipids

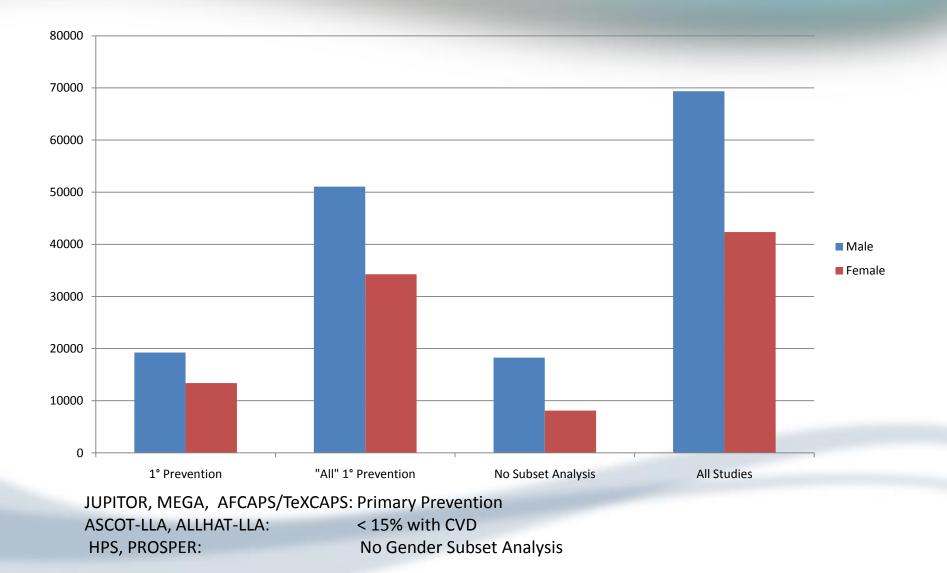


Do Satins Work Equally Well for Men and Women? *Time Magazine March 31, 2010*

Representation of \bigcirc **in Clinical Trials of Cholesterol Lowering Medicines**



Representation of \bigcirc **in Clinical Trials of Cholesterol Lowering Medicines**



Justification of the Use of Statins (JUPITER)

- Study Design
 - Randomized placebo controlled trial
 - 20mg rosuvastatin
 - **11,00**
 - Men>50 yo Women > 60 yo
 - CRP < 2.0 ng/l LDL < 130 mg/dl</p>
- The rates of the primary end point were 0.77 and 1.36 per 100 person-years of follow-up in the rosuvastatin and placebo groups
- The rosuvastatin group did have a higher incidence of physician-reported diabetes.
- NNT 120 for 1.9 years to prevent 1 event

Justification of the Use of Statins (JUPITER)

Outcome Event	Men p-value	Women p-value
Primary Event	<0.0001	0.02
Non-fatal MI	<0.0001	0.18
Any MI	0.0006	0.11
Any Stroke	0.0005	0.40
Arterial revascularization and hospitalization	0.002	<0.0001
MI, Stroke, Death from CVD	<0.0001	0.16
Any Death	0.08	0.12
New Onset DM	0.29	0.008

Carotid Atorvastatin Study in Hyperlipidemic Postmenopausal Women (CASHMERE)

- 12 month trial
- Postmenopausal Women
- Moderate Dyslipidemia
- Randomized Placebo Controlled
- 80mg Atorvastatin vs. Placebo (n=192 vs 206)
- Results: No change in intimal medial thickness (IMT)

Primary Prevention with Statins in Women

The risk benefit ratio of statins for primary prevention of CVD in women has not been <u>definitively</u> established.

The differing results in men compared to women may be due to the diminished effect of elevated LDL in women compared to men or the lack of evaluation of more unique risk factors

The Aspirin Connection



Women's Health Study Aspirin New England Journal March 2005

- 40,000 Women
- Aspirin 100 mg every other day vs placebo
- Followed for 10 years
- Results
 - Non-fatal Stroke
 - Non-fatal Heart Attack
 - Hemorrhagic Stroke
 - Cardiovascular disease death

Aspirin Results in Men vs Women

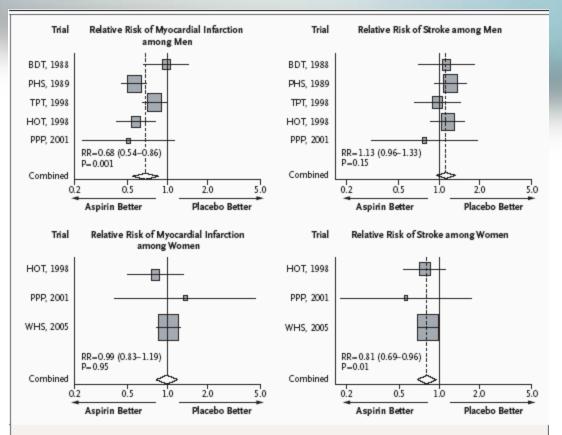


Figure 3. Aspirin in the Primary Prevention of Myocardial Infarction and Stroke among Men and Women.

The results of a sex-specific random-effects meta-analysis of data from six trials are shown: the British Doctors' Trial (BDT), the Physicians' Health Study (PHS), the Thrombosis Prevention Trial (TPT), the Hypertension Optimal Treatment (HOT) study, the Primary Prevention Project (PPP), and the current Women's Health Study (WHS). The relative risk (RR) and 95 percent confidence interval (in parentheses) are shown for each trial (indicated by the box and horizontal line through each box, respectively), and the relative risk is shown for the combined results (indicated by the diamond and the dashed line in each graph). For the relative risk of myocardial infarction among women, the dashed line is coincident with the solid line at 1.00. The size of the box is proportional to the amount of information in the corresponding trial.

stratified I	by percentage	e of ma	le participants				
Male (%)	Non-fatal MI	р*	Fatal MI	p*	Both fatal and non-fatal MI	р*	
0–66	0.87 (0.71–1.06)	0.26	0.87 (0.65–1.17)	0.86	0.86 (0.79–0.95)	0.52	
70–89	0.72 (0.61–0.86)	0.23	0.91 (0.75–1.11)	0.24	0.82 (0.71–0.95)	0.13	
100	0.62 (0.54–0.71)	0.48	0.55 (0.20–1.53)	0.01	0.63 (0.46–0.85)	0.02	
Total	0.72 (0.64–0.81)	0.03	0.88 (0.75–1.03)	0.19	0.79 (0.72–0.87)	< 0.05	

Relative risks and 95% confidence intervals of myocardial infarction stratified by percentage of male participants

MI, myocardial infarction.

*Test for heterogeneity, p value is from χ^2 test. Ransom effects model was used in all combinations.

Yerman *et al. BMC Medicine* 2007 total of 23 trials (n = 113,494 participants)

Gender and Aspirin

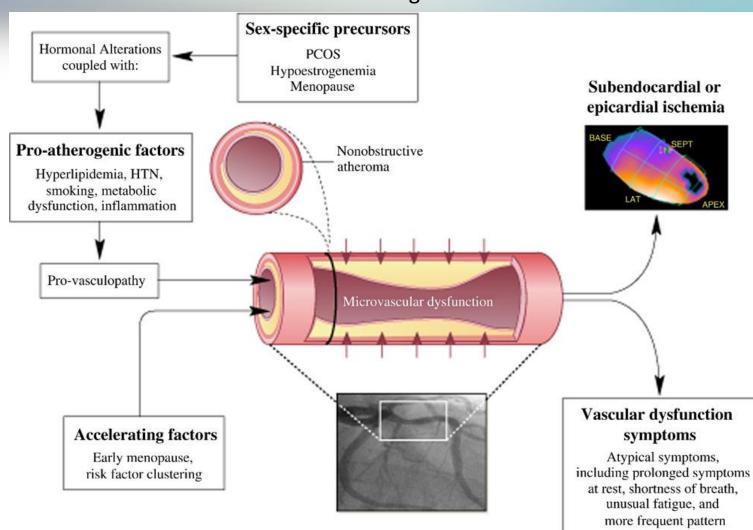
Men

- Decrease Risk of MI
- No/Slight Decrease
 Risk of Stroke
- Recommendations
 - Men with known CAD
 - Men with a <u>></u> 10% 10 yr risk
 - Men > 45 yo when benefit > risk

- Women
 - ? decrease of MI
 - Decrease in Risk of Stroke
 - Recommendations
 - Women with known CAD
 - Women with a <u>>10%</u>
 10 yr risk
 - Women > 55 yo if benefit > risk

Cardiovascular Disease

- Women arrive at the hospital later than men
- Younger women were less likely to be treated with aspirin
- Women were more likely to be given a diagnosis other than coronary artery disease
- Less women discharged on beta-blockers
- Women have a slightly higher 30 day mortality
- Women had longer door-to-needle time (12.5 min)



Model of Microvascular Angina in Women

Shaw, L. et al, JACC Nov 2009

Mental Health

Antidepressants

- Prescribing rate for antidepressants increased by 48% between 1995-2002
- 118 million prescriptions (CDC, 2005)
- Most commonly prescribed medication (CDC, 2005)

Gender and Depression

Parameter	Women as compared to Men
Age at onset	Earlier
Lifetime Prevalence Rate	Higher (20% vs. 10%)
Duration of episodes	Longer
Relapse Rate	Higher
Effect of Seasonal Changes	Greater
Response to SSRI	Women
Co-morbid anxiety	Greater
Co-morbid Alcohol and Substance Abuse	Less
Association of Gonadal Hormones	Greater
Suicide Attempts	Greater
Suicide Completion	Less

Kornstein S, Primary Psychiatry. 2003;10(12):58-61

Nicotine Addiction

Effect of Nicotine on Women

Nicotine

- Women: calming effects on stress-induced mood changes
- Men: enhance aggressive moods
- Nicotine + alcohol
 - Women: appears to enhance the effects of alcohol.
 - Men: appears to dilute some of the sedating and intoxicating effects of alcohol.
- Women may be more responsive than men to nonnicotine stimuli

Smoking Cessation

- Quitting Smoking
 - Women less successful than men
 - Women relapse more often and for different reasons than men
- Women > Men
 - Join smoking cessation groups
 - Experience severe withdrawal symptoms
 - Worry about weight gain
- Men > Women
 - Benefit from nicotine replacement therapy
- Women = men
 - Effectiveness of bupropion

Gender and Diabetes

- Women vs. Men with OSA
 - New-onset DM: OR 11.8
- Severe Diabetic Retinopathy
 - M > F
- Risk of sudden death in patients with adult-onset DM
 - 300% (F) vs 50% (M)
- Skin Disorders in DM
 - Lipodystrophy F>M
 - Acanthosis Nigricans F>M
- Baseline albuminuria

M>F

Celen YT et al, J Clin Sleep Med. 2010 Jun 15;6(3):244-50 Chatziralli IP et al, BMC Res Notes. 2010 Jun 1;3:153 Oseaghdha DM et al Am J Kidney Dis. 2010 Jul 2

Gender-Specific Medicine

- Encompasses all Aspects of Medicine
- Proven Gender Differences
- Future Applications in Clinical Practice
- Improved Clinical Care
- Expand Research and Education Programs
- Benefits Both Men and Women

Gender-Specific Health Disparities

The study of sex- and gender-based differences in healthcare continues to be limited by

- a historical deficit in data on women
- an emerging understanding of unique issues about women's gender specific medicine
- the absence of standardized reporting mechanisms to collect and analyze data

Challenges

- To determine where and when sex- and gender-based differences in health emerge
- To determine the clinical significance of observed differences

Gender Disparities Exist



Thank you for Your Attention