

Sex and Gender Differences

LIPIDS AND LIPOPROTEINS



Lipids are fat molecules and combine with proteins to transport fat and cholesterol throughout the body.

Abbreviations: HDL-C (high-density lipoprotein cholesterol); LDL-C (low-density lipoprotein cholesterol); TG (triglycerides)

FEMALE

ALL

MALE

Hormonal Influence

- Pregnancy: higher estrogen levels lead to increased LDL-C, HDL-C, and TG
- Menopause: small increase in LDL-C
- Polycystic Ovarian Syndrome: higher testosterone levels lead to increased TG and decreased HDL-C

- Estrogen, testosterone, and other sex hormones regulate lipid metabolism in the liver
- Oral estrogen (e.g., some oral contraceptives, feminizing hormone therapy) can affect lipids
- Anabolic steroid use causes very low HDL-C levels
- Gender Affirming Hormone Therapy (HT)
 - Feminizing HT: decreased LDL-C, increased TG, increased HDL-C
 - Masculinizing HT: increased LDL-C, increased TG, decreased HDL-C

- Puberty: HDL-C decreases by 10mg/dL due to testosterone
- Adult men: HDL-C is 10mg/dL lower than in adult women
- Hypogonadism: increased LDL-C and TG
 - May correct with testosterone replacement

Prevalence

- Ideal HDL in females \geq 50 mg/dL
- 8.5% have HDL-C $<$ 40 mg/dL
- Mean TG = 86.8 mg/dL
- 28.1% have LDL-C \geq 130 mg/dL

- Ideal TG $<$ 150 mg/dL
- Ideal LDL $<$ 100 mg/dL

- Ideal HDL in males \geq 40 mg/dL
- 26.6% have HDL-C $<$ 40 mg/dL
- Mean TG = 100.6 mg/dL
- 27.4% have LDL-C \geq 130 mg/dL

Presentation

- Higher incidence of metabolic syndrome
 - Elevated TG
 - Decreased HDL-C

- Dyslipidemia is usually asymptomatic
- LDL-C elevation is associated with atherosclerotic cardiovascular disease (ASCVD)
- CVD is the leading cause of death in U.S. men and women
- Hypercholesterolemia can lead to:
 - Angina
 - Myocardial infarction
 - Ischemic stroke
- Hypertriglyceridemia can lead to pancreatitis

- Onset of atherosclerotic cardiovascular disease occurs 10 years earlier

Treatment

- Statins contraindicated in pregnancy
- Statin treatment:
 - Less likely to be offered despite being eligible
 - Less likely to receive dosing at the guideline-recommended intensity
 - At a higher risk of myopathy as a side effect

- Statin treatment:
 - Similar LDL-C reduction (unless affected by hormone therapy)
 - Similar reduction in risk of ASCVD
- Alternative medications available: bile acid sequestrants, cholesterol absorption inhibitors, PCSK9 inhibitors, fibrates

- Statin treatment:
 - More likely to be offered guideline-recommended therapy
 - More likely to achieve recommended lipid levels

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Definitions & Disclaimers



References

