

NORTHWEST AIDS EDUCATION AND TRAINING CENTER

Hypogonadism and Testosterone Replacement in Men with HIV

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Overview

- 1. Definition and diagnosis of hypogonadism
- 2. Prevalence among HIV infected men and diagnostic confounders
- 3. Risks and benefits of testosterone replacement
- 4. Testosterone delivery and monitoring



Why consider treating male hypogonadism? Androgen (estrogen) Effects

skin

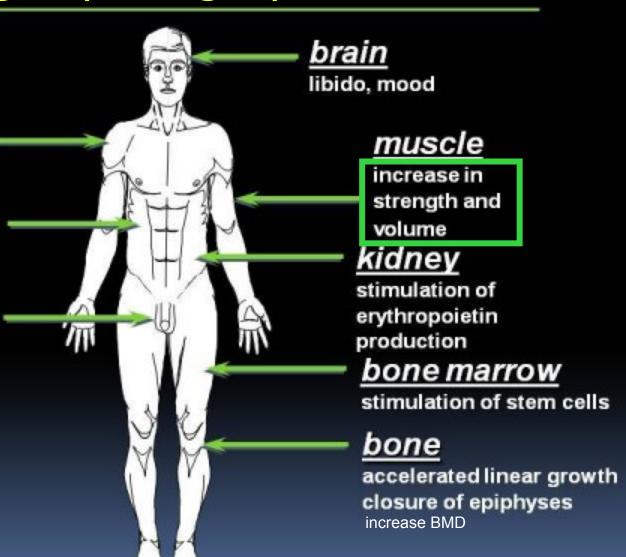
hair growth, balding, sebum production

<u>liver</u>

synthesis of serum proteins

<u>male sexual</u> <u>organs</u>

penile growth
spermatogenesis
prostate growth and
function



- Prevalence of hypogonadism among HIV-infected men pre-HAART 50%, now 13-20%, more common among men with HIV-associated wt. loss
- Among ALL men, incidence of hypogonadism increases with AGE, obesity, and chronic illness
- Endocrine Society 2010 Clinical Practice Guideline for Androgen Deficiency Syndromes: consider screening men with HIV - associated weight loss

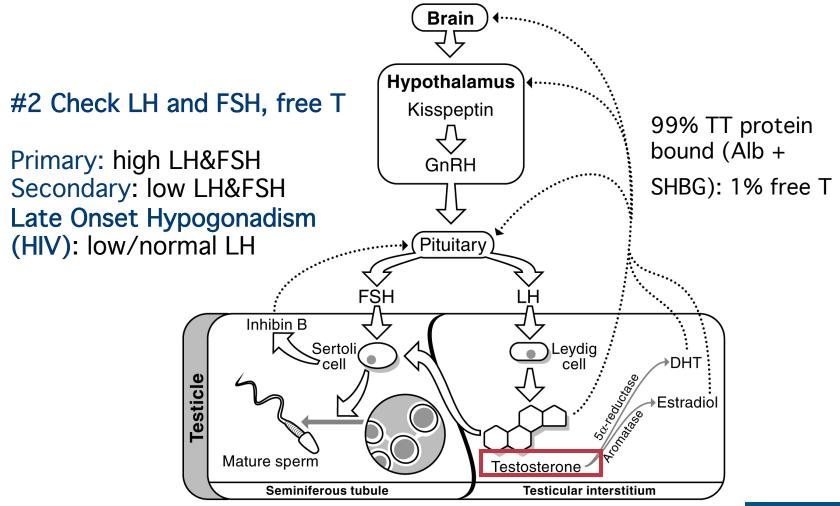


Signs and Symptoms of Hypogonadism in Adult Men

- Sexual complaints (↓ libido, ED).infertility
- Gynecomastia,
 ↓ male pattern hair
- ↓ energy, fatigue, moodiness, ↓ muscle mass and strength, increased fat mass
- NON-SPECIFIC: poor overlap with biochemical criteria, most associated with normal aging



Androgen Production and Metabolism: Diagnosing Hypogonadism



#1 Symptom + Low serum total testosterone AM, fasting, (< 280-300 ng/dl)



Making the Diagnosis of Hypogonadism

- Combination of symptoms + biochemical evidence. Get calculated free T with HIV +
- Primary (high LH): testicular defect
- Secondary (low LH): screen for other Pit hormones, consider MRI, hemochromatosis, anabolic steroids
- Low/normal LH: pit MRI, OPIATES (including methadone), chronic illness, malnutrition, aging



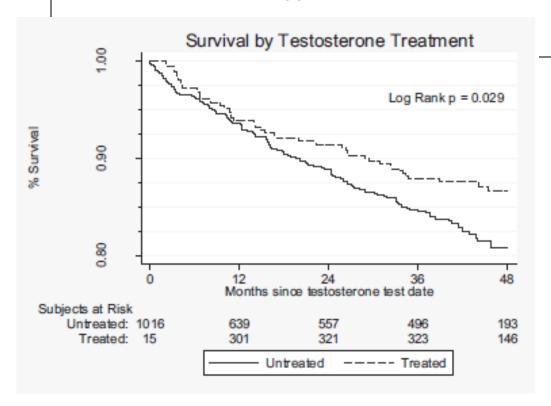
Summary: Risks and benefits of T therapy in men with hypogonadism

- BENEFITS: T therapy will increase lean mass, BMD, libido and HCT and will decrease fat mass. Mild increase in leg strength
 - --- Physical performance, ED, and QOL more mixed results, mild improvement overall
 - --- improvements in IS, metabolic parameters mixed
 - --- benefits generally seen in men with lowest T levels at baseline
- RISKS: unknown. RCT no increase AE.
 - PROSTATE: increases PSA but ? PCa, BPH
 - CVD: lowers HDL, LDL and Total Chol
 - HCT
 - \$\$\$



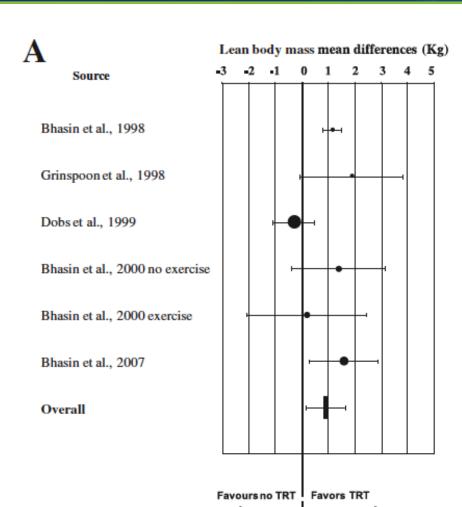
Testosterone and Mortality

- No appropriate RCT to evaluate mortality benefit
- Observational data: multiple large, prospective studies in last 5 years show association between low serum T and earlier mortality (MMAS, VA, Mr. OS)
 - Recent data suggestive of possible treatment benefit (no harm)





RCT of HIV-infected men treated with Testosterone

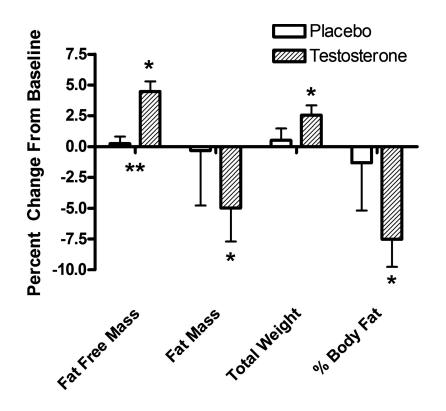


Limitations

- ALL SMALL (n <100) and short (24 weeks max)
- all done with men with HIV-associated wt. loss
- all but one in pre-HAART era
- LBM not functional outcome



Testosterone as an Anabolic agent in HIV



- 3x supraphysiologic T given to HIV+ men with unintentional wt. loss
- NO INCREASE vs.
 PLACEBO in strength, mood, stair climb
- No adverse events



Options for testosterone replacement therapy

- Depot formulations
 - Intramuscular (q2week)
 - Subcutaneous implant
- Daily dosing
 - Transdermal (\$\$)
 - Buccal
 - Oral
- Follow-up (3-6 months)
 - HCT
 - PSA (if have baseline)





Summary

- Incidence of hypogonadism may be slightly higher in HIV infected men then HIV- but main RF is age
- Diagnosis based on low T levels + sx, check free T, LH and FSH (or refer based on low T)
- Consider use of anabolic steroids, opiates, and concomitant illness when making the diagnosis
- T therapy will increase LBM, BMD, libido and HCT. Effect on strength likely positive. QOL effects unclear. Risks not known (\$\$). Monitor for benefit

