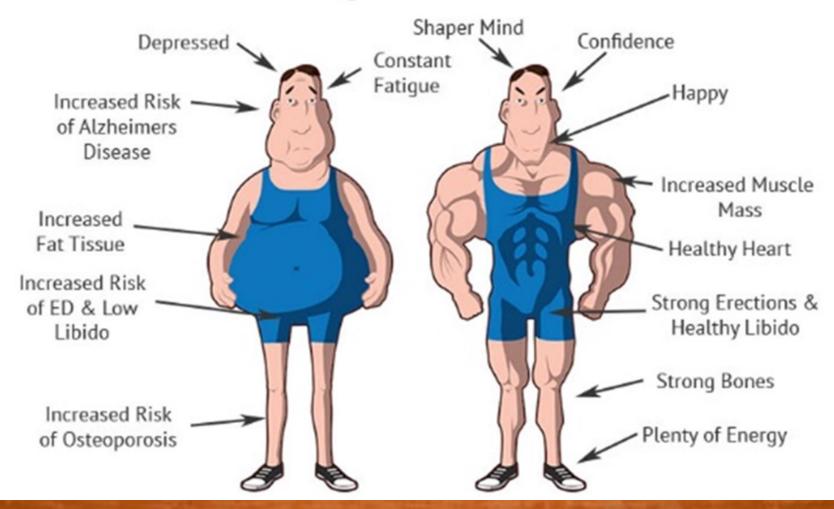
# Double Trouble: Role of testosterone deficiency and obesity in prostate cancer

David S. Lopez, Dr.PH, MPH, MS
Assistant Professor of Epidemiology / Clinical Assistant Professor
Department of Epidemiology
Division of Urology
UTHealth- School of Public Health
UTHealth McGovern Medical School

## Benefits of Optimal Testosterone







Aging, insecurity and the \$2 billion testosterone industry







## Hypothesized Causal Pathway

Race and Ethnicity



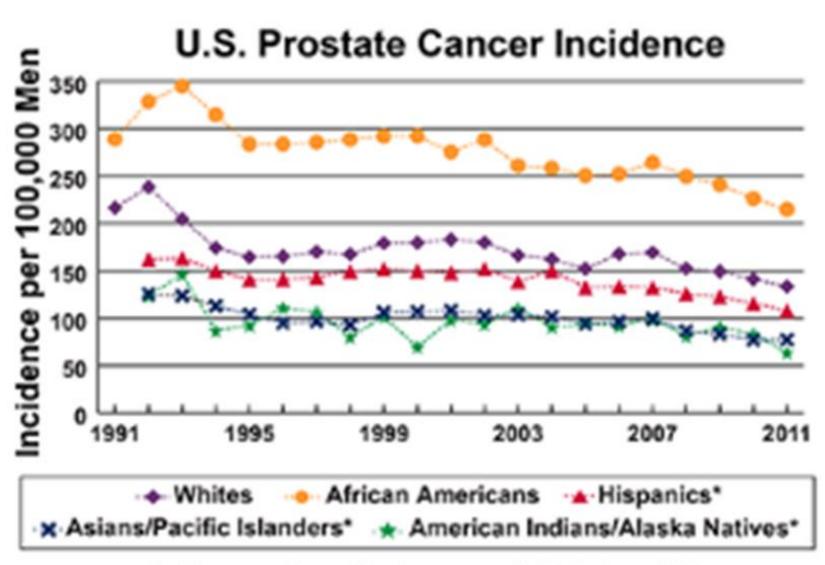
- a) Body Mass Index
  - b) Waist circumference
  - c) % Body Fat
- d) Adipose Tissue

Other Mechanisms

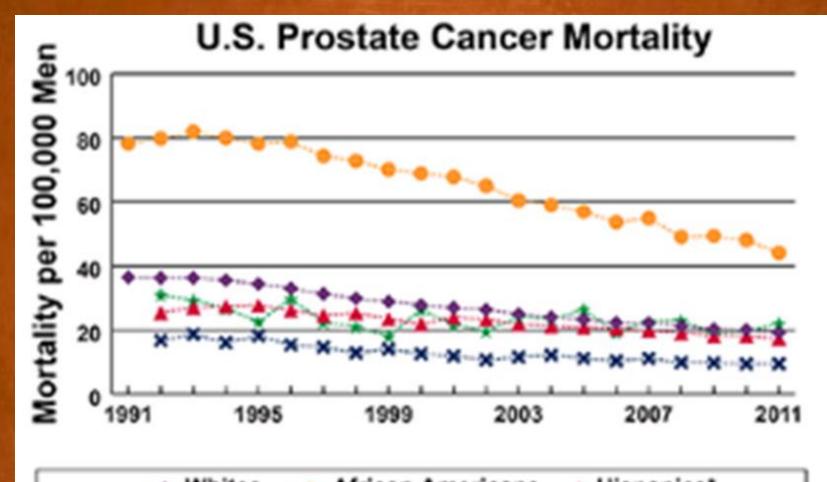
- 1) Insulin-like growth factor-axis
- 2) Inflammation

Prostate Cancer

- A) Grade
- B) Stage
- C) Fatal



\*Incidence and mortality data not available before 1992.



→ Whites → African Americans → Hispanics\*
× Asians/Pacific Islanders\* → American Indians/Alaska Natives\*

\*Incidence and mortality data not available before 1992.

### Prevalence of Obesity in Men

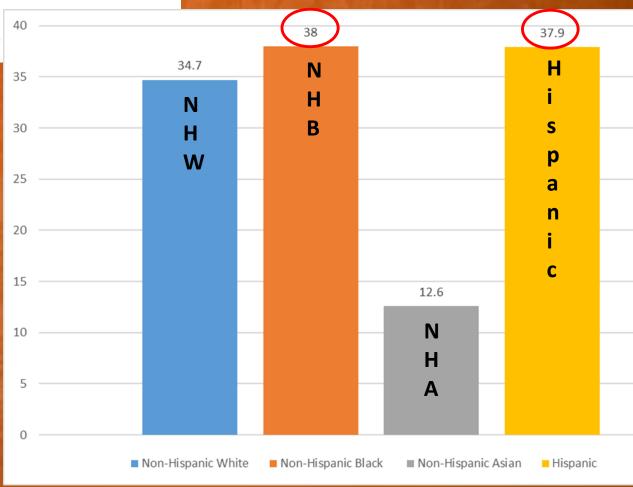
Research

#### **Original Investigation**

Trends in Obesity Among Adults in the United States,

2005 to 2014

Katherine M. Flegal, PhD; Deanna Kruszon-Moran, MS; Margaret D. Carroll, MSPH; Cheryl D. Fryar, MSPH; Cynthia L. Ogden, PhD



### Obesity and Prostate Cancer

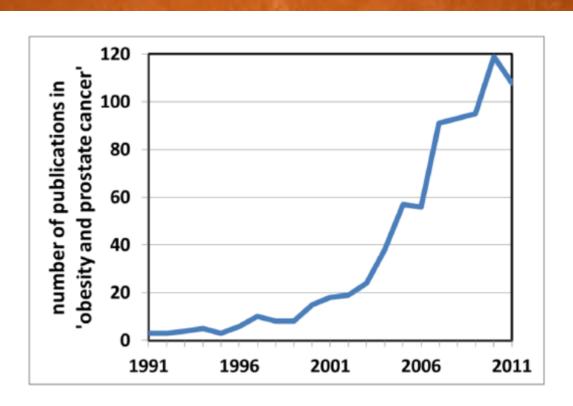
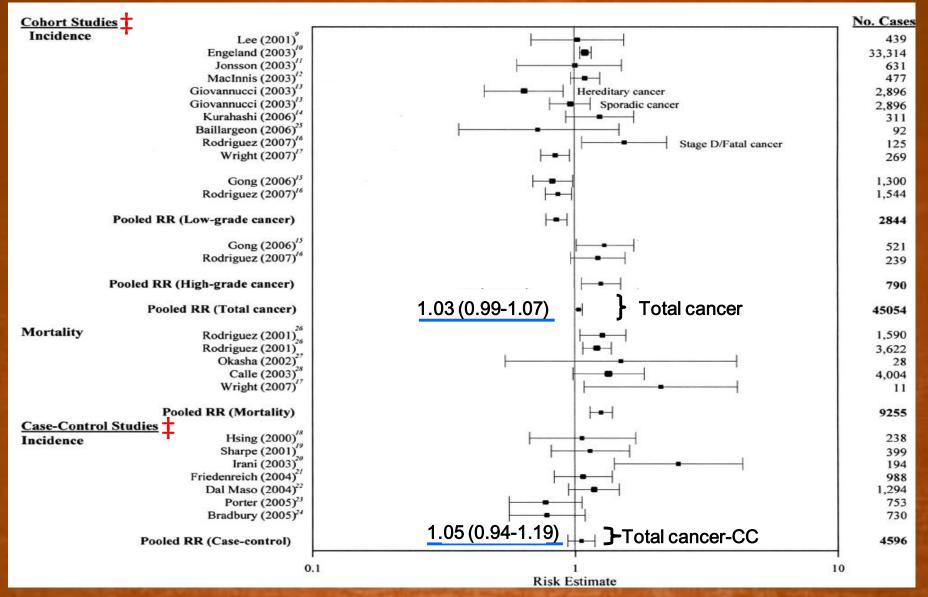


Fig. 1.
Using the search terms *obesity* and *prostate cancer*, the number of PubMed publications has increased in the past 20 yr.



\*Recent meta-analyses of prospective studies show similar observations w/ localized and advanced PCa

## Prostate cancer in Mexican-Americans

The Prostate 68:563 – 570 (2008)

## Prostate Cancer in Mexican-Americans: Identification of Risk Factors

Sara S. Strom,<sup>1</sup>\* Yuko Yamamura,<sup>1</sup> F. Nery Flores-Sandoval,<sup>2</sup> Curtis A. Pettaway,<sup>3</sup> and David S. Lopez<sup>1</sup>

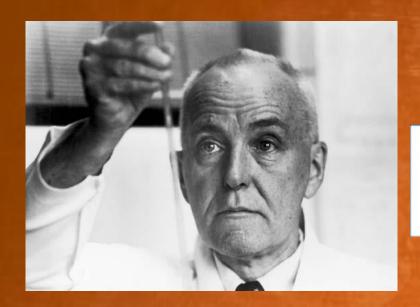
<sup>1</sup>The University of Texas M.D. Anderson Cancer Center, Department of Epidemiology, Houston, Texas <sup>2</sup>St. Luke's Episcopal Hospital, Department of Urology, Houston, Texas <sup>3</sup>The University of Texas M.D. Anderson Cancer Center, Department of Urology, Houston, Texas

## Prostate cancer in Mexican-Americans

TABLE IV. Prostate Cancer Risk Factors by Clinical Stage at Diagnosis				
Variable	Organ-confined ( $N = 127$ )		Regionally advanced/metastatic ( $N = 49$ )	
	Univariable OR (95% CI)	Multivariable OR <sup>a</sup> (95% CI)	Univariable OR (95% CI)	Multivariable OR <sup>a</sup> (95% CI)
Obese at diagnosis No	1.00	_	1.00	1.00
Yes	1.07 (0.65–1.75)	_	1.90 (1.00-3.65)	2.40 (1.11-5.15)

What is the biological mechanism involved in the association between obesity and prostate cancer?





### Charles Huggins

Circa 1941

#### Studies on Prostatic Cancer

The Effect of Castration, of Estrogen and of Androgen
 Injection on Serum Phosphatases in Metastatic
 Carcinoma of the Prostate\*

Charles Huggins, M.D., and Clarence V. Hodges, M.D.

Huggins C. et al. Cancer Res. 1941:1; 293

- Castration decreased acid phosphatase in men with metastatic PCa
- Testosterone administration raised acid phosphatase
- "Cancer of the prostate is activated by testosterone injections."
- Huggins awarded Nobel Prize 1966

#### The Year 2007—Post Doctoral Fellowship



#### Hormone JOHNS HOPKINS Demonstration Program

What is the relationship between race/ethnicity and sex steroid hormones in adults (n=1,413)- National Health and Nutrition and Examination Survey (NHANES III)?

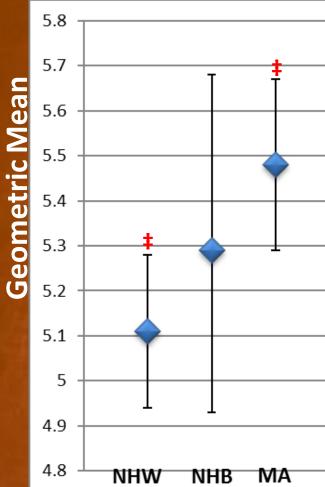
0021-972X/07/\$15.00/0 Printed in U.S.A.

The Journal of Clinical Endocrinology & Metabolism 92(7):2519-2525 Copyright © 2007 by The Endocrine Society doi: 10.1210/jc.2007-0028

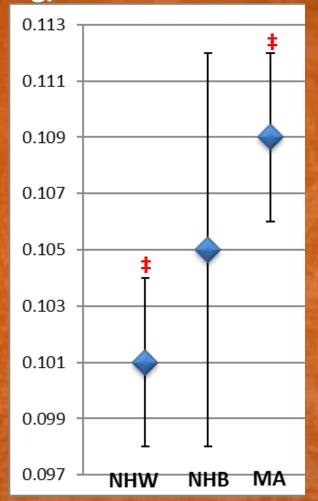
Serum Estrogen, But Not Testosterone, Levels Differ between Black and White Men in a Nationally Representative Sample of Americans

Sabine Rohrmann, William G. Nelson, Nader Rifai, Terry R. Brown, Adrian Dobs, Norma Kanarek, James D. Yager, and Elizabeth A. Platz

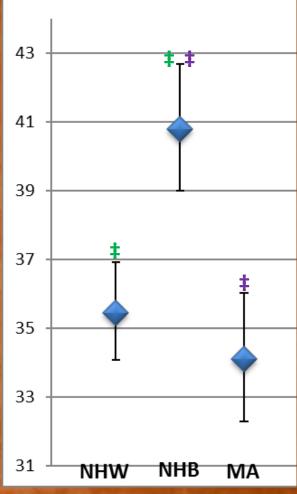
## Total Testosterone, ng/ml



## Free Testosterone, ng/ml



## Estradiol, pg/ml



Model 1 is adjusted for age, BMI, smoking, alcohol consumption, and physical activity; model 2 is the same as model 1 but adjusted for percent body fat instead of BMI; percent body fat is missing for 118 men. MA, Mexican-American; NHB, non-Hispanic black; NHW, non-Hispanic white.

a.b. NHW vs. MA:  $^aP < 0.05$ ;  $^bP < 0.01$ .

ed NHW vs. NHB: eP < 0.05; P < 0.01.

<sup>\*</sup>f NHB vs. MA: \*P < 0.05; fP < 0.01.

## What is the relationship between race/ethnicity and sex steroid hormones in adolescents (n=134)- NHANES III?



#### NIH Public Access

#### **Author Manuscript**

Cancer Causes Control. Author manuscript; available in PMC 2013 December 04.

Published in final edited form as:

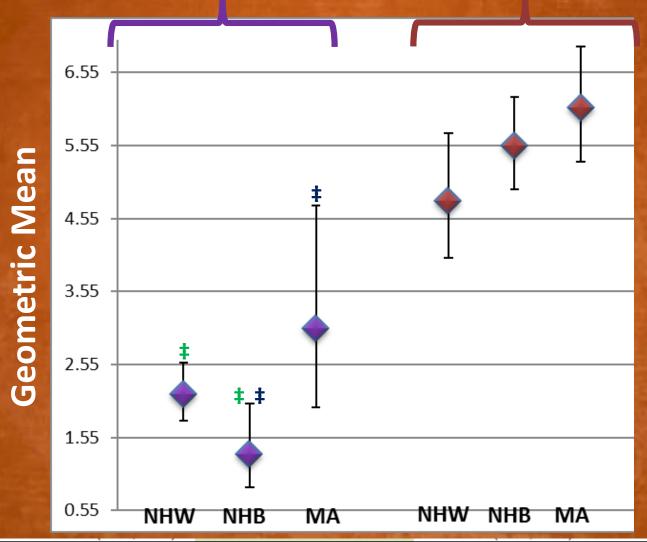
Cancer Causes Control. 2013 April; 24(4): . doi:10.1007/s10552-013-0154-8.

## Racial/ethnic differences in serum sex steroid hormone concentrations in US adolescent males

David S. Lopez<sup>1,2</sup>, Sarah B. Peskoe<sup>1</sup>, Corinne E. Joshu<sup>1</sup>, Adrian Dobs<sup>3,4</sup>, Manning Feinleib<sup>1</sup>, Norma Kanarek<sup>4,5</sup>, William G. Nelson<sup>4,5,6,7</sup>, Elizabeth Selvin<sup>1</sup>, Sabine Rohrmann<sup>8</sup>, and Elizabeth A. Platz<sup>1,4,6</sup>

#### Testosterone, ng/ml

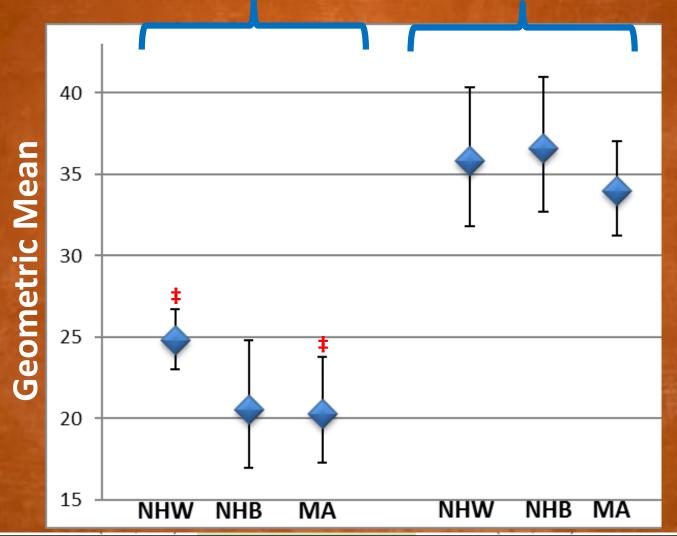
12-15 yrs old (n=57) 16-19 yrs old (n=77)



<sup>&</sup>lt;sup>a</sup> Geometric mean adjusted for age, Tanner stage, percent body fat, waist circumference, frequency of physical activity, cotinine (as an indicator of tobacco smoke exposure), and mutually for the other hormones (total testosterone, total estradiol, and SHBG; free testosterone and free estradiol).

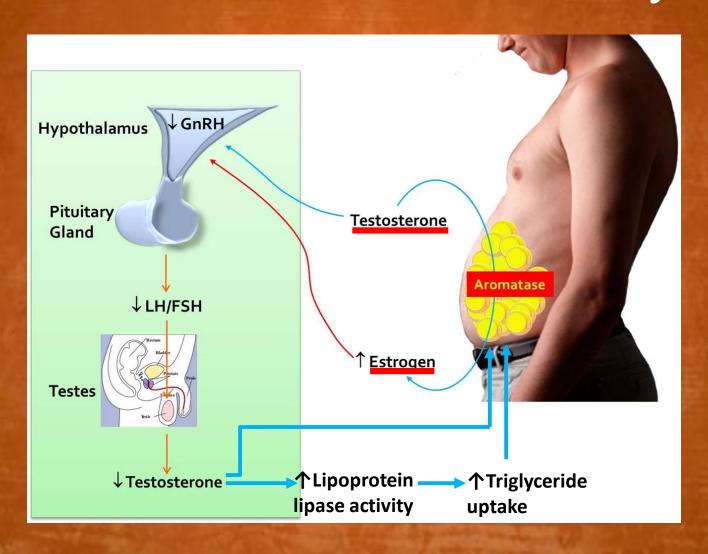
#### Estradiol, pg/ml

12-15 yrs old (n=57) 16-19 yrs old (n=77)



<sup>&</sup>lt;sup>a</sup> Geometric mean adjusted for age, Tanner stage, percent body fat, waist circumference, frequency of physical activity, cotinine (as an indicator of tobacco smoke exposure), and mutually for the other hormones (total testosterone, total estradiol, and SHBG; free testosterone and free estradiol).

### Testosterone and Obesity?



## What is the role of body fatness in sex steroid hormones among adult men (n=1,265) - NHANES III?



#### NIH Public Access

#### **Author Manuscript**

Cancer Causes Control. Author manuscript; available in PMC 2013 November 06.

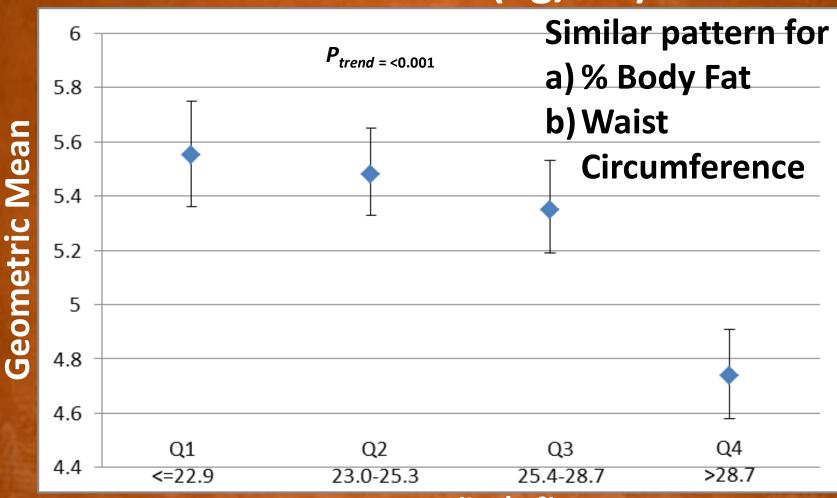
Published in final edited form as:

Cancer Causes Control. 2011 August; 22(8): . doi:10.1007/s10552-011-9790-z.

## Body fatness and sex steroid hormone concentrations in US men – Results from NHANES III

Sabine Rohrmann<sup>1</sup>, Meredith S. Shiels<sup>2</sup>, David S. Lopez<sup>2</sup>, Nader Rifai<sup>3</sup>, William G. Nelson<sup>4,5,6,7</sup>, Norma Kanarek<sup>4,7</sup>, Eliseo Guallar<sup>2,8</sup>, Andy Menke<sup>2,8</sup>, Corinne E Joshu<sup>2</sup>, Manning Feinleib<sup>2</sup>, Siobhan Sutcliffe<sup>9</sup>, and Elizabeth A. Platz<sup>2,4,5</sup>

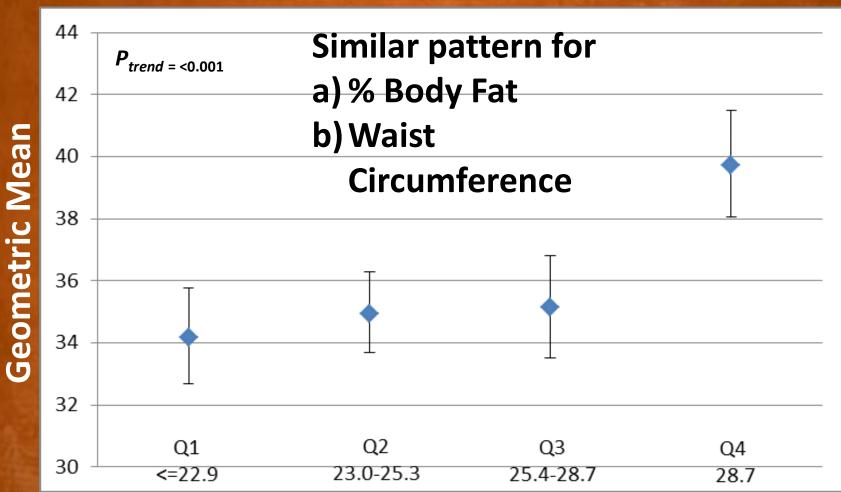
#### Testosterone (ng/mL)



BMI (kg/m<sup>2</sup>)

Adjusted for age, race/ethnicity, smoking, alcohol, physical activity, plus testosterone, estradiol, and SHBH mutually adjusted and free testosterone and free estradiol mutually adjusted.

#### Estradiol (pg/mL)



BMI (kg/m²)

Adjusted for age, race/ethnicity, smoking, alcohol, physical activity, plus testosterone, estradiol, and SHBH mutually adjusted and free testosterone and free estradiol mutually adjusted.

#### Time to Think

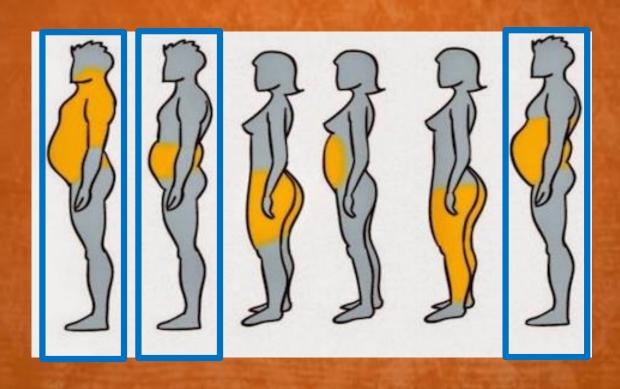
- Incidence and mortality rates of PCa higher in African Americans
- Hugging et al. suggested that higher levels of testosterone increased risk of prostate cancer
- Obesity decreases the risk of low-grade/localized PCa, but increases the risk of high-grade/advanced PCa.
- And obesity decreases testosterone levels

#### HOWEVER,

- We did not see high levels of testosterone in African Americans. Mexican Americans had higher testosterone levels, but they have lower rates of PCa
- If obesity reduces testosterone, how does obesity increase the risk of high-grade/advanced PCa?



## Measurements of Adiposity, Body Fatness, and Race/Ethnicity



# What is the association of overall and central body fatness with sex steroid hormones in adult men (n=1,243) and does it vary by race/ethnicity in NHANES III?

Racial/ethnic differences in the associations of overall and central body fatness

with circulating hormones and metabolic factors in US men

David S. Lopez<sup>1,2</sup>, Sabine Rohrmann<sup>3</sup>, Sarah B. Peskoe<sup>4</sup>, Corinne E. Joshu<sup>4,5</sup>,

Konstantinos K. Tsilidis<sup>6,7</sup>, Elizabeth Selvin<sup>4</sup>, Adrian S. Dobs<sup>5,8</sup>, Norma Kanarek<sup>5,9</sup>,

Steven Canfield<sup>2</sup>, William G. Nelson<sup>5,8,10</sup>, Elizabeth A. Platz<sup>4,5,10</sup>

Submitted to International Journal of Endocrinology and Metabolism

#### Obese by both measures:

High % Body Fat (≥ 25%) and High Waist Circumference (≥ 102 cm)

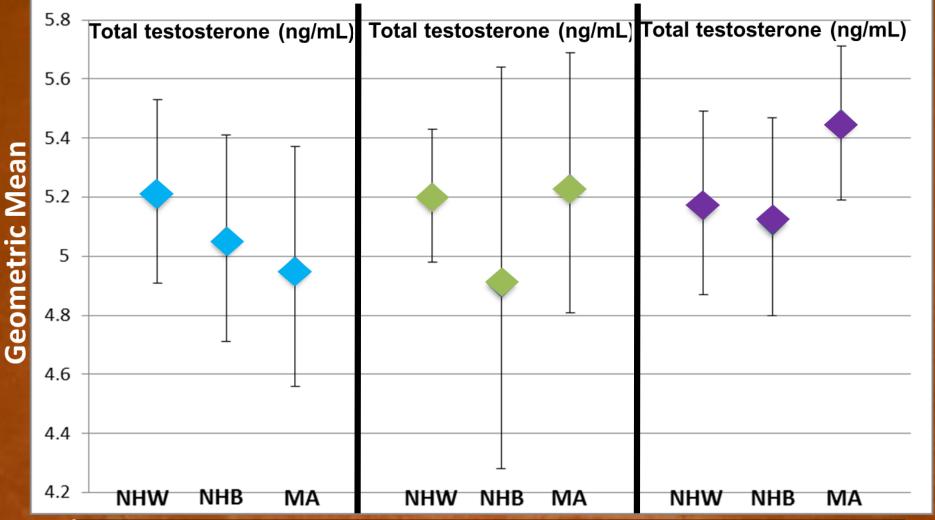
#### Obese by one measure:

High % Body Fat (≥ 25%) and Normal Waist Circumference (< 102 cm)

Normal % Body Fat (< 25%) and High Waist Circumference (≥ 102 cm)

#### Non-obese both measure:

Normal % Body Fat (< 25%) Normal Circumference (< 102 cm)



#### Obese by both measures: Obese by one measure:

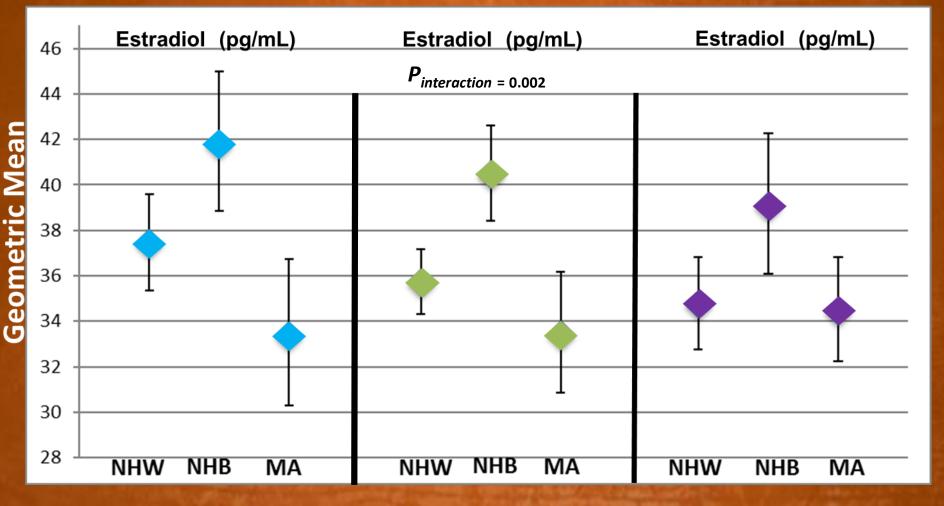
High % Body Fat (≥ 25%) and High Waist Circumference (≥ 102 cm)

High % Body Fat (≥ 25%) and Normal Waist Circumference (< 102 cm)

Normal % Body Fat (< 25%) and High OR Waist Circumference (≥ 102 cm)

#### Non-obese both measure:

Normal % Body Fat (< 25%) Normal Circumference (< 102 cm)



## What about prostate-specific antigen (PSA) test?

Strongly correlated to testosterone and it is still used for disease progression / risk stratification

JOURNAL OF MEN'S HEALTH Volume 11, Number 4, 2014 © Mary Ann Liebert, Inc. DOI: 10.1089/jomh.2014.0058

## Racial and Ethnic Differences in the Association of Metabolic Syndrome with Prostate-Specific Antigen Levels in U.S. Men: NHANES 2001–2006

David S. Lopez, DrPH, MPH, MS,<sup>1,2</sup> Shailesh Advani, MPH,<sup>1,3</sup> Konstantinos K. Tsilidis, PhD,<sup>4</sup> Mike Hernandez, MS,<sup>5</sup> Elaine Symanski, PhD,<sup>1</sup> Sara S. Strom, PhD,<sup>3</sup> Arup Sinha, MS,<sup>6</sup> and Steven Canfield, MD<sup>2</sup>

#### NHANES 2001-2006 n=3528- Objectives:

- 1. Determine the association of PSA with the 3 definitions of metabolic syndrome (ATP, IDF and WHO)
- 2. Investigate whether the relationship of PSA with metabolic syndrome varies by race and ethnicity

#### Adult Treatment Panel III (ATP III)<sup>a</sup>

Any 3 or more of the following criteria:

- 1. Waist Circumference (WC)
  - Men: ≥ 102 cm
- 2. Blood Pressure (BP)b
  - SBP ≥ 130 mmHg OR
  - DBP ≥ 85 mmHg
- 3. HDL Cholesterol
  - Men: < 40 mg/dL</li>
- 4. Serum Triglycerides
  - ≥ 150 mg/dL
- 5. Fasting Plasma Glucose
  - ≥ 110 mg/dL

#### International Diabetes Federation (IDF)<sup>c</sup>

Central obesity (defined as waist circumference) plus any two of the following four criteria:

- Central obesity- waist circumference (WC)
  - White men: ≥94 cm
  - Black men: ≥94 cm
  - Mexican-American men: ≥90 cm
- 2. Blood Pressure (BP)<sup>d</sup>
  - SBP ≥ 130 mmHg OR
  - DBP ≥ 85 mmHg
- 3. HDL Cholesterol
  - Men: < 40 mg/dL</p>
- 4. Serum Triglycerides
  - ≥ 150 mg/dL
- 5. Fasting Plasma Glucose<sup>e</sup>
  - ≥ 100 mg/dL

#### World Health Organization (WHO)

Insulin resistance<sup>†</sup> or Diabetes<sup>‡</sup> and at least two of the following criteria:

- 1. Body Mass Index (BMI)
  - ≥ 30 kg/m<sup>2</sup> AND/OR

Waist-to-Hip Ratio (WHR)

- Men: ≥ 0.90
- 2. Blood Pressure (BP)<sup>g</sup>
  - ≥ 140/90 mmHg

OR

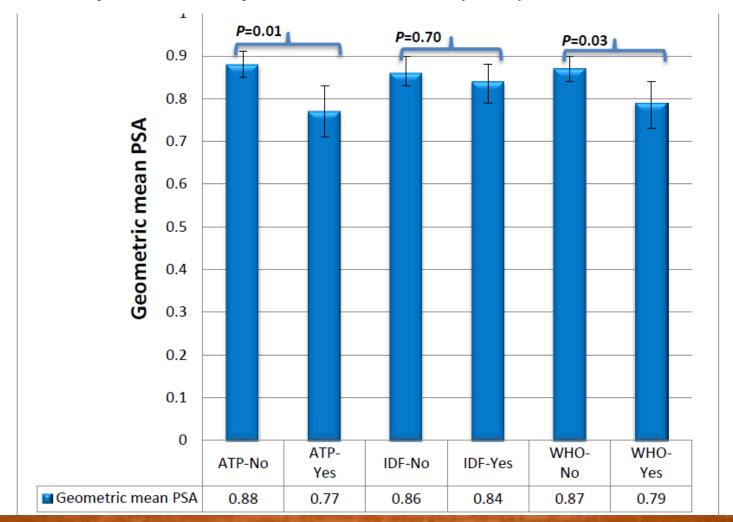
- DBP ≥ 90 mmHg
- 3. HDL Cholesterol
  - Men: < 35 mg/dL

AND/OR

**Serum Triglycerides** 

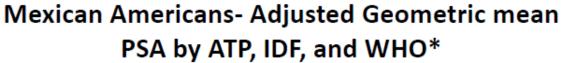
- ≥ 150 mg/dL
- 4. Albumin-to-creatinine ratio
  - ≥ 30 mg/g

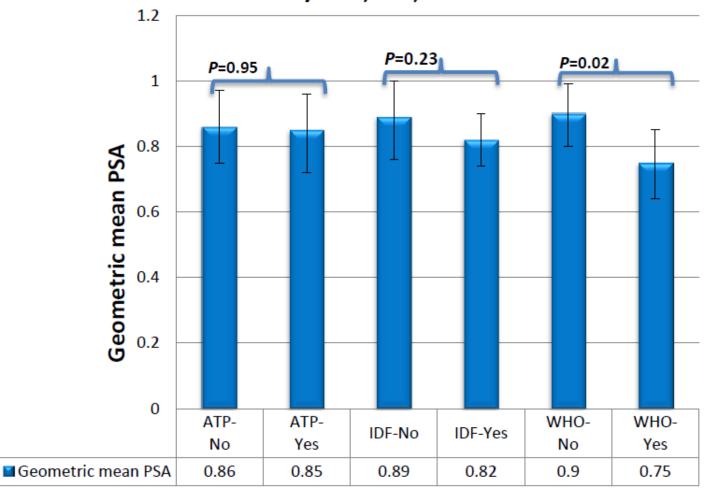
## Multivariable-adjusted geometric mean (95% confidence interval) of PSA by metabolic syndrome definitions, ATP, IDF and WHO.



<sup>&</sup>lt;sup>b</sup>Model 1, geometric mean adjusted for age.

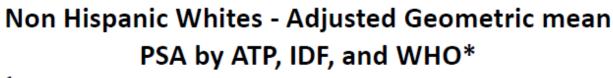
<sup>&</sup>lt;sup>c</sup>Model 2, geometric mean adjusted for age, smoking, moderate and vigorous physical activity, and plasma volume.

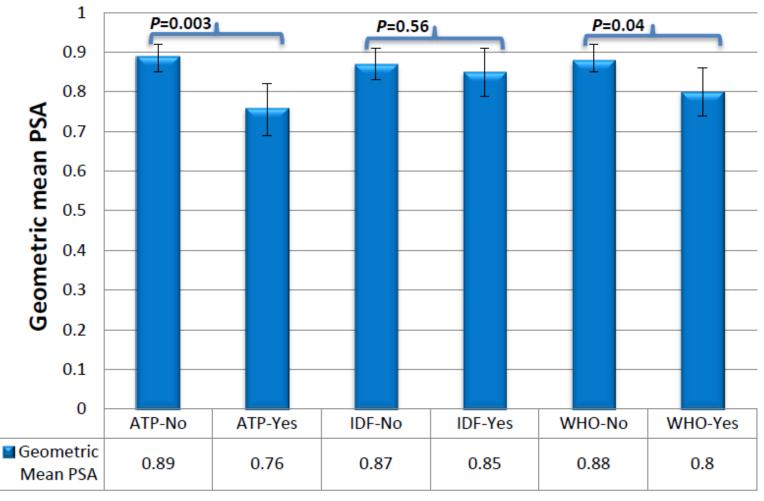




<sup>&</sup>lt;sup>b</sup>Model 1, geometric mean adjusted for age.

<sup>&</sup>lt;sup>c</sup>Model 2, geometric mean adjusted for age, smoking, moderate and vigorous physical activity, and plasma volume.

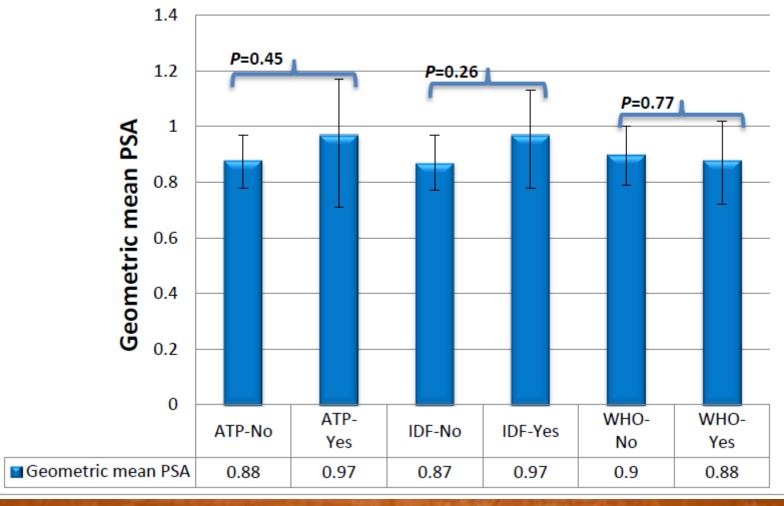




<sup>&</sup>lt;sup>b</sup>Model 1, geometric mean adjusted for age.

<sup>&</sup>lt;sup>c</sup>Model 2, geometric mean adjusted for age, smoking, moderate and vigorous physical activity, and plasma volume.

## Non-Hispanic Black - Adjusted Geometric mean PSA by ATP, IDF and WHO\*



<sup>b</sup>Model 1, geometric mean adjusted for age.

<sup>&</sup>lt;sup>c</sup>Model 2, geometric mean adjusted for age, smoking, moderate and vigorous physical activity, and plasma volume.

# Mortality & Testosterone and Obesity



#### Mortality- Testosterone and Obesity



- Testosterone deficiency (TD) is a common medical condition that is characterized by having total testosterone levels < 300 ng/dL</li>
- Studies show that as many as 38.7% of men over 45y old₁ (24% among men 30-79y old₁) demonstrate TD.
- In the United States, reports show that there are approximately 2.4 million 40-69 yrs men with TD.<sup>17,18</sup>
- Projected that by 2025 approximately 6.5 million American men aged 30 to 80 will suffer from TD.<sup>4</sup>

## A prospective analyses to determine whether the co-occurrence of testosterone deficiency and obesity increase the risk of death in US men-NHANES 1999-2004 & 2011 Mortality Data

Co-occurrence of testosterone deficiency and obesity associated with all-cause mortality in U.S. men

Authors: David S. Lonez, Vueting Oiu, Shailesh Advani, Konstantings K. Tsilidis, Mohit Khara, Abraham A total of 0.49 participants with a modian follow up time of 0.5 years Morgentaler, Rur

Introduction and obje testosterone ≤ 300 na/c relationship between T unclear. Therefore, we circumference [WC] ≥ 1

Methods: The data wer a follow-up of mortality in men who participated measured weight in kild values of T deficiency a

Results: A median follo were fit to estimate haz was a 3.45 fold increase 8.02). No significant as: men had a 4.52 fold inc abdominal obesity had

Conclusion: Men with T deficiency and obesit

A total of 948 participants with a median follow-up time of 9.5 years					
Characteristics	Death / Alive (142 / 806)				
	(1427 000)				
T deficiency (TD, ≤ 300 ng/dL) and Overall					
obesity (BMI ≥ 30 kg/m²)					
No TD and No Overall obesity	63/449				
TD / No Overall obesity	42/130				
Overall obesity / No TD	10/111				
TD and Overall obesity	27/116				
T deficiency (TD, ≤ 300 ng/dL) and abdominal					
obesity (WC ≥ 102 cm)					
No TD / No Abdominal obesity	46/406				
TD / No Abdominal obesity	26/90				
Abdominal obesity / No TD	27/154				
TD and Abdominal obesity	43/156				

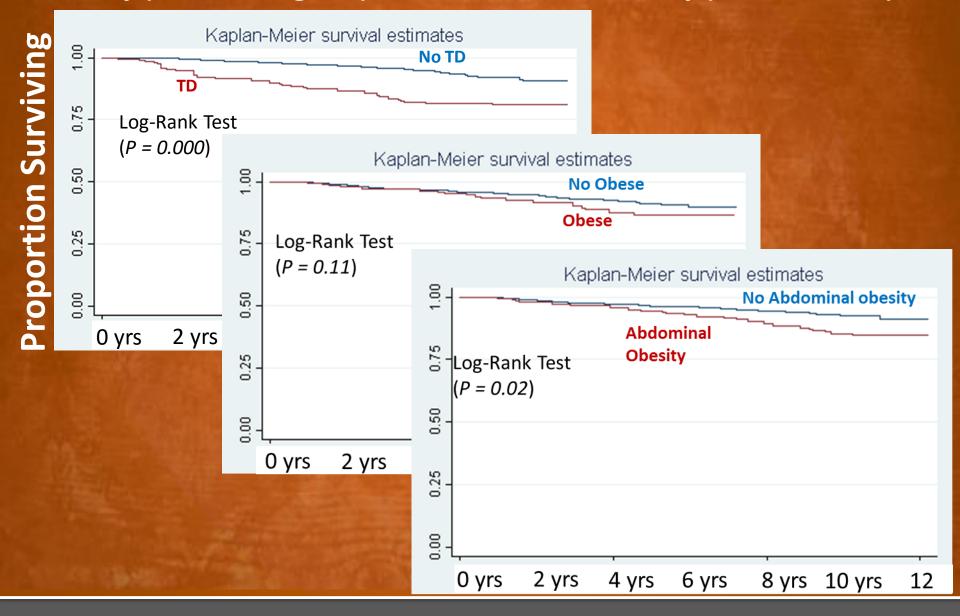
erone (T deficiency: total tes (U.S.). A bidirectional ealth outcomes remains or abdominal obesity (waist

) years old. The latter file was sex hormones were measured 3MI was calculated from mbined the dichotomized

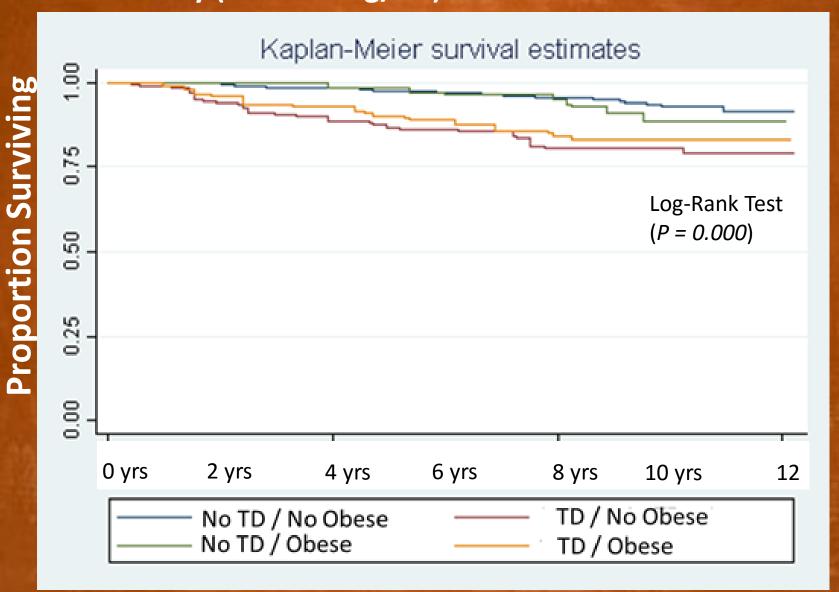
portional hazard models and physical activity. There ence interval [CI] = 1.48 ency and overall obesity, rly, men with T deficiency and 1 = 1.51 - 12.23

/. However, the effect of both

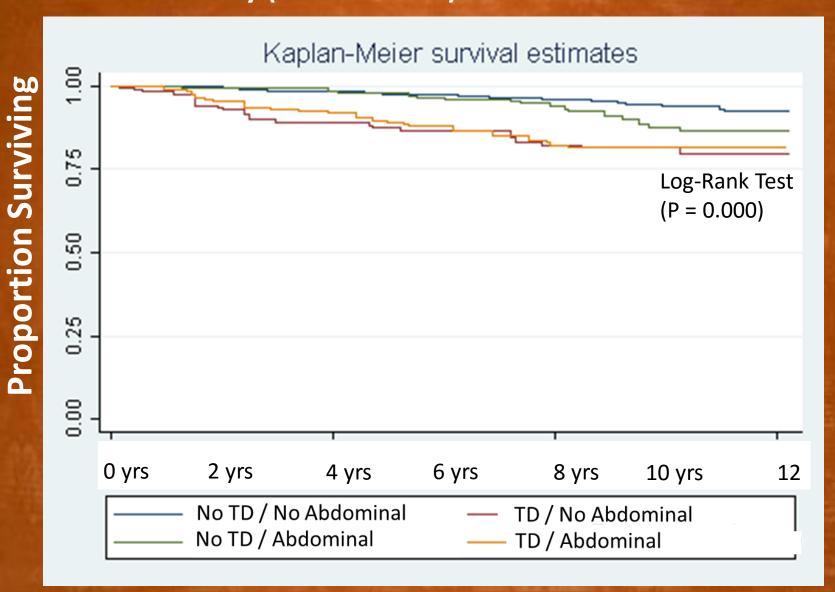
# Testosterone Deficiency (TD, total testosterone ≤ 300 ng/dL), Overall Obesity (BMI ≥ 30 kg/m<sup>2</sup>), and Abdominal Obesity (WC ≥ 102 cm)



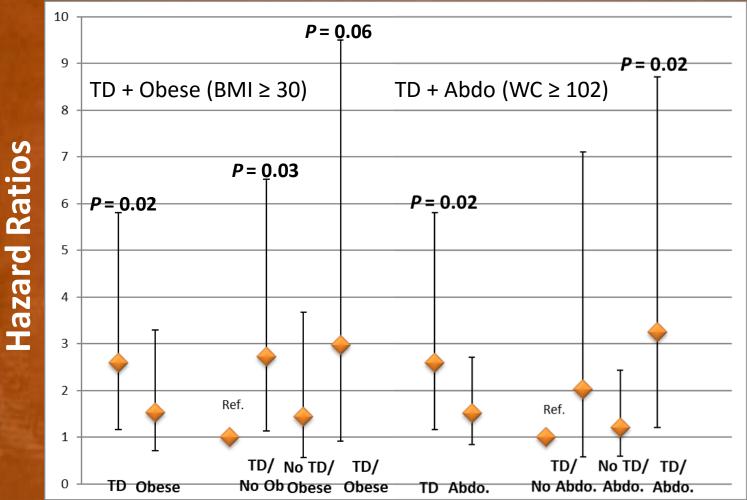
# Testosterone Deficiency (TD, total testosterone ≤ 300 ng/dL) and Overall Obesity (BMI ≥ 30 kg/m²)



# Testosterone Deficiency (TD, total testosterone ≤ 300 ng/dL) and Abdominal Obesity (WC ≥ 102 cm)



# Association of the co-occurrence of testosterone deficiency and obesity with increase risk of death in US men



Adjusted for SHBG, estradiol, age, race/ethnicity, smoking status, alcohol consumption and physical activity

#### Testosterone & Prostate Cancer



#### Charles Huggins

#### Studies on Prostatic Cancer

 The Effect of Castration, of Estrogen and of Androgen Injection on Serum Phosphatases in Metastatic
 Carcinoma of the Prostate\*

Charles Huggins, M.D., and Charence V. Hodges, M.D.

Histories C at al Contac Dec 104

- Castration decreased acid phosphatase in men with metastatic PCa
- Testosterone administration raised acid phosphatase
- "Cancer of the prostate is activated by testosterone injections."
- · Huggins awarded Nobel Prize 1966



**Low Testosterone** 

- Raising testosterone causes rapid PCa growth-
  - -High levels of testosterone or taking testosterone therapy is
  - "Like feeding a hungry tumor" or "pouring gasoline on a fire."

Heal

Conti

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and I 1200

## Testosterone replacement therapy and the heart: friend, foe or bystander?

David S. Lopez<sup>1,2</sup>, Steven Canfield<sup>2</sup>, Run Wang<sup>2</sup>

Department of Epidemiology, Human Genetics and Environmental Sciences, The University of Texas Health Science Center, School of Public

### Endogenous and Exogenous Testosterone and Prostate Cancer: Increased, Decreased or Null Effects

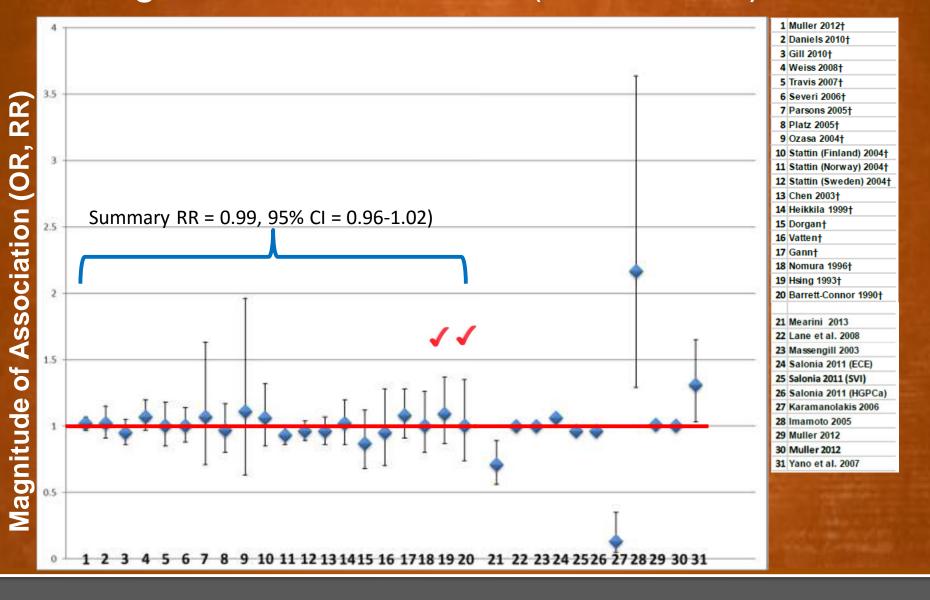
Authors: David S. Lopez, Konstantinos K. Tsilidis, Run Wang, Steven Canfield

<sup>1</sup>Department of Epidemiology, Human Genetics and Environmental Sciences, The University of Texas Health Science Center, School of Public Health, Houston, TX, USA; <sup>2</sup>Division of Urology, The University of Texas Health Science Center, McGovern Medical School, Houston, TX, USA Contributions: (II) Conception and design: All authors; (II) Administrative support: None; (III) Provision of study materials or patients: None; (IV) Collection and assembly of data: DS Lopez; (V) Data analysis and interpretation: All authors; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.

Correspondence to: David S. Lopez, DrPH, MPH, MS. Assistant Professor, Clinical Assistant Professor, Division of Epidemiology, Human Genetics and Environmental Sciences, Division of Urology, University of Texas School of Public Health, University of Texas Medical School at Houston, 1200 Pressler Street, suite E-629, Houston, TX 77030, USA. Email: david.s.lopez@uth.tmc.edu.

- Measures of association (OR, RR, HR's) from multivariable analyses
- Endogenous (continuous and categorical) and exogenous testosterone

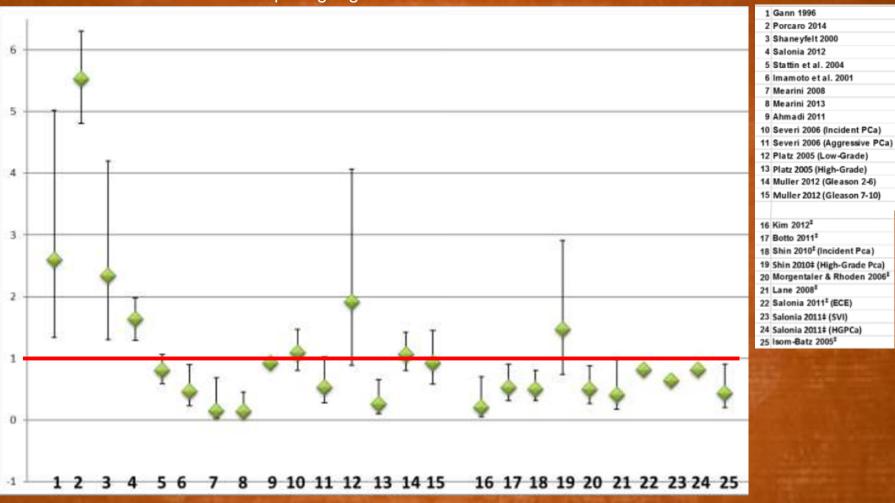
### Review Article Endogenous Testosterone(continuous) & PCa



#### Review Article

### Endogenous Testosterone(categorical) & PCa

Comparing High vs Low levels of Testosterone

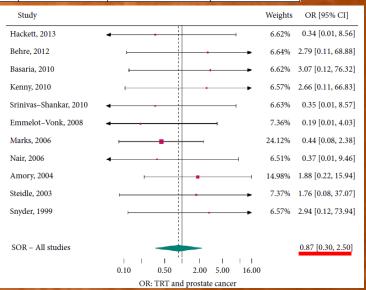


# Testosterone Therapy & PCa (randomized controlled trials)

Table 3. Meta-analyses of randomized controlled trials (RCT's) that investigated the effect of TTh on PCa events (stage and grade). Only included studies that conducted multivariable analyses and reported magnitude of association (Odds Ratio, Relative Risk, etc.)

Study author	Study design	Number of	Number of	TTh (n=) /	Prostate cancer:	Results: only fully-	Statistically
and year		studies /	participants	Placebo (n=)	total, grade	adjusted	significant (SS-
		trials	(n=)		and/or stage	multivariable	P < 0.05) or not
					(N=)	models	significant (NS)
Cui et al. 2014	Randomized	5	1,168	778 / 390	Incident prostate	Summary OR =	NS
	controlled trials-				cancer, n = 12	0.74; 95% CI = 0.25-	
	Short term (<12					4.65	
	months)						
	Randomized	3	379	191 / 188	Incident prostate	Summary OR =	NS
	controlled trials-				cancer, n = 6	0.99; 95% CI = 0.24-	
	Long term (12-36					4.02	
	months)						
Boyle et al. 2016	Randomized	11	2,013	1,114 / 899	Incident prostate	Summary OR =	NS
	controlled trials				cancer, n = 20	0.87; 95% CI = 0.30-	
	(most of the trials <					2.5	
	12 months)						

a testosterone		placebo		Odds Ratio			Odds Ratio			
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% C	<b>Year</b>	M-H, Fixed,	95% CI	
1.1.1 Injection										
Marks LS 2006 I	2	21	4	19	51.5%	0.39 [0.06, 2.45]	2006		-	
Subtotal (95% CI)		21		19	51.5%	0.39 [0.06, 2.45]				
Total events	2		4							
Heterogeneity: Not applicable										
Test for overall effect: Z = 1.00 (P = 0.32)										
1.1.2 Transdermal										
Basaris S 2010 T	1	106	0	103	6.8%	2.94 [0.12, 73.08]			-	-
Steidle C2003 T	2	307	0	99	10.2%	1.63 [0.08, 34.21]		-		
Kawfaman JM 2011 T	2	214	0	37	11.4%	0.88 [0.04, 18.75]		-		
Srinivas-Shank U 2010 T	0	130	1	132	20.1%	0.34 [0.01, 8.32]	2010	•		
Subtotal (95% CI)		757		371	48.5%	1.10 [0.26, 4.65]				
Total events	5		1							
Heterogeneity: Chi <sup>2</sup> = 0.97,	df = 3 (P =	0.81); I	$^{2} = 0\%$							
Test for overall effect: Z = 0.13 (P = 0.90)										
Total (95% CI)		778		390	100.0%	0.74 [0.25, 2.19]				
Total events	7		5							
Heterogeneity: Chi² = 1.67, df = 4 (P = 0.80); l² = 0%										
Test for overall effect: Z = 0.55 (P = 0.58) 0.01 0.1 1 10 100										
Test for subgroup differences: Not applicable testosterone placebo										
						7 7 7 4 4 4 4				

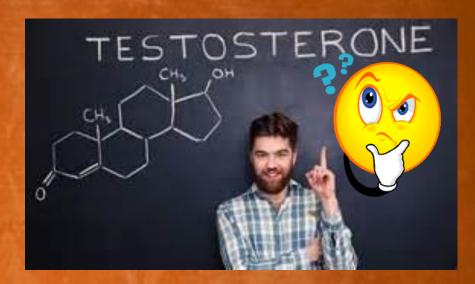


## **Another Moment to Think**

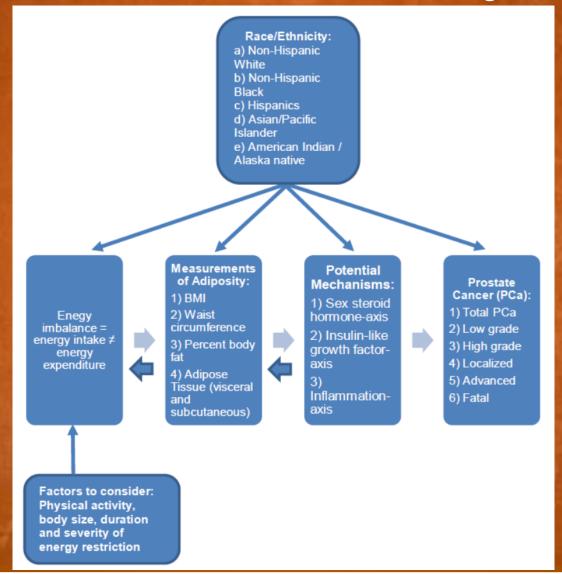
- Not proposing the use of testosterone therapy
- Not proposing to increase the levels of testosterone in men with prostate cancer

#### HOWEVER,

 I want to appreciate the body of literature that negates the the contention that "high levels of testosterone increases prostate cancer."



(Prostate Cancer, Testosterone, Diet, and Weight Management)



(Prostate Cancer and SSB's)



- International consortium of 15 cohort studies from North America, Europe, Australia and Asia.
- 52, 683 primary incident PCa cases, including 4,924 advanced cases among 842, 149 men
- Proposing to investigate SSB's and PCa

 We received pilot funding (\$20,000) to investigate the prospective association between caffeine intake and ED



RESEARCH ARTICLE

Role of Caffeine Intake on Erectile Dysfunction in US Men: Results from NHANES 2001-2004

David S. Lopez<sup>1,2</sup>\*, Run Wang<sup>2</sup>, Konstantinos K. Tsilidis<sup>3</sup>, Huirong Zhu<sup>1</sup>, Carrie R. Daniel<sup>4</sup>, Arup Sinha<sup>5</sup>, Steven Canfield<sup>2</sup>

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Coffee May Reduce Risk of Lethal Prostate

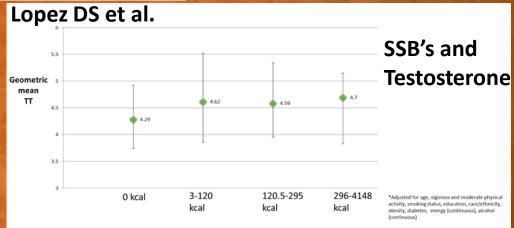
Cancer in Men
Read the press release.

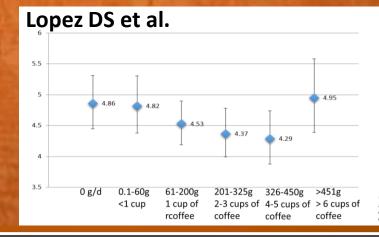
2011 HPFS Newsletter

(Testosterone, diet, physical activity and weight management)

- Currently, investigating the relationship of SSB's and Caffeine intake with testosterone levels.
- We received pilot funding (\$20,000) to conduct a comprehensive analyses between diet, physical activity and weight management and testosterone levels.





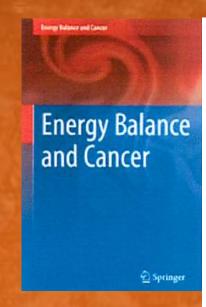


Caffeine intake and Testosterone

\*Adjusted for age, vigorous and moderate physical activity, smoking status, education, race/ethnicity, obesity, diabetes, total water intake (plain and tap), energy (continuous), alcohol (continuous)

## Next Steps and Future Directions

- "Racial/Ethnic
  Differences in the
  Association between
  Energy Balance and
  Prostate Cancer."
- Remain undetermined the interplay between measurement of adiposity, testosterone and advanced/fatal PCa
- We need randomized controlled trials to investigate relationship between diet, physical activity, weight management and PCa



#### 

Springer Science+Business Media Berger/Platz Prostate Cancer Proposal (Volume 13)

#### Co-Edited Volume

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Introduction and Preface - Energy Balance and Prostate Cancer

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