The Continuing Significance of Obesity for Survival to Old Age: A Re-Examination of Age Effect on the Body Weight-Mortality Relationship

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Question: Does the Weight-Mortality Link Vary by Age?

Significance

- 1. Optimal Weight management
- 2. Age-specific weight guidelines
- Two contrasting hypotheses
 - 1. Age-as-leveler
 - Changes in the effect of body fat
 - Co-morbidities and short span
 - Weight change
 - Mortality selection
 - 2. Cumulative disadvantage: Permanent scarring

• Measure of mortality differentials: Mortality ratio

Previous Studies: Findings and Limitations

- Findings: Inconsistent but leaning toward a diminishing effect (e.g., Stevens et al. 1998; Bender et al. 1999; Visscher et al. 2000)
- Limitations:
 - 1. Comparison of baseline age groups: Cohort effect
 - Cohort variations in BMI mortality differentials may distort age variations because at one given time, different cohorts are at different age.

2. Outcome not age-specific: Age overlapping in age groups

Previous Studies: Time since Baseline as Analysis Time Compare Baseline Age groups



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Previous Studies: Time since Baseline as Analysis Time

Compare Points of Time in Follow-up



Analysis Time: Time Since Baseline

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This Study: Age as Analysis Time Compare Age-Specific BMI Differentials within Cohorts



Data: 1971-1975 NHANES I

- National Health and Nutrition Examination Survey: Representative of non-institutionalized US population
- A sub-sample of those aged 24-77 at baseline: medical examination, with height and weight measured
- Mortality through 1992
- Body Mass Index (BMI): kg/m^2
- Baseline BMI groups: WHO guidelines
 - Underweight (BMI < 18.5): excluded in the analysis
 - Normal weight (18.5 \leq BMI < 25): reference category

- Overweight ($25 \leq BMI < 30$)
- Obese (BMI ≥ 30)

Data: NHANES I, Sample Selections

- Total: 14407 cases
- Deletions: 1035 cases
 - Immediately lost after baseline: 546 cases
 - Missing baseline BMI: 5 cases
 - Baseline underweight: 484 cases
- Analytic Sample: 5496 men and 7876 women
- Five birth cohorts:
 - 1895-1904
 - 1905-1914
 - 1915-1924
 - 1925-1934
 - 1935-1950

Methods: Gompertz Model

• Baseline:
$$h(a) = exp(\beta + \gamma a)$$

• BMI effect on β only: No Age variations

$$\beta = \beta_0 + \beta_1 W$$

• BMI effect on β and γ : Age variations

$$\beta = \beta_0 + \beta_1 W$$

$$\gamma = \gamma_0 + \gamma_1 W.$$

• Cohort-BMI interaction: Cohort variations

$$\beta = \beta_0 + \beta_1 W + \beta_2 C + \beta_3 WC$$

$$\gamma = \gamma_0 + \gamma_1 W + \gamma_2 C + \gamma_3 WC.$$

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Methods: Cont'd Frailty, Model Selection, and Robustness

- Gompertz with Frailty: $h(a) = \alpha exp(\beta + \gamma a)$
 - α : Mean 1 and variance θ
 - 1. Gamma distribution
 - 2. Inverse-Gaussian Distribution
- Analytic Strategy: The simplest model that fits the data best

- The AIC criteria
- Semi-parametric Cox model

For the Sake of Replication

Compare Baseline Age Group: Men



For the Sake of Replication

Compare Baseline Age Group: Women



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Results for Men: Mortality by Baseline BMI and Cohort



Age in Years

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Results for Men: Mortality Ratio, Baseline Obese/Normal



Age in Years

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Results for Men: Estimated Ratio and 95% C.I. Baseline Obese/Normal



Age in Years

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Results for Men: Estimated Ratio and 95% C.I. Baseline Obese/Normal



Age in Years

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Results for Women: Mortality by Baseline BMI and Cohort



Age in Years

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Results for Women: Mortality Ratio, Obese/Normal



Age in Years

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Results for Women: Estimated Ratio and 95% C.I. Baseline Obese/Normal



Age in Years

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Results for Women: Estimated Ratio and 95% C.I. Baseline Obese/Normal



Age in Years

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Discussions

- Findings
 - 1. Age variations: Supportive of cumulative disadvantage
 - Men: Increase by age
 - Women: Constant
 - 2. Cohort variations: Increase from earlier to later cohorts
 - Implication for age variations
- Next steps
 - 1. Contribution of disease composition to age patterns
 - 2. Cohort changes:
 - Temporal changes as discussed in Su 2005 and Flegal et al. 2005
 - Body weight as more salient risk factor?
 - Health selection changed over time?
 - Weight change: Weight Measured at different ages?
 - 3. Analytic issues:
 - Smoking behavior
 - Early deaths: How many years of death should we delete?
 - Sample size: NHANES II