ARIC Manuscript Proposal #770

PC Reviewed: 02/15/01 Status: A Priority: 2 SC Reviewed: 03/01/01 Status: A Priority: 2

1.a. Full Title: Identification of psychological attributes associated with the risk of incident diabetes

b. Abbreviated Title (Length 26 characters): Psychological traits and diabetes

2. Writing Group (list individual with lead responsibility first):

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3. Timeline: Analysis to begin immediately

4. Rationale:

Identification of modifiable risk factors for diabetes is important for its prevention. There has been a growing interest in psychological traits as novel risk factors for diabetes. Possible mechanisms include the influence of these traits on behaviors, such as activity and diet, or their influence on the activity of the hypothalamic-pituitary-adrenal (HPA) axis.

The prevalence of depression in diabetic individuals is increased compared to the general population (1;2) and several studies have demonstrated that the presence of depressive symptoms is predictive of incident diabetes (3-6). Disordered carbohydrate metabolism has been demonstrated in non-diabetic individuals with depression/mood disorders (7;8). Individuals with depression are also known to have hypercortisolism (7). Thus, a possible mechanism by which depression may lead to the development of diabetes is via neuroendocrine activation of the hypothalamic-pituitary-adrenal (HPA) axis or the sympathetic nervous system. Both cortisol and catecholamines are counterregulatory hormones that can lead to a state of insulin resistance.

Depressive symptoms have also been shown to be predictive of cardiovascular disease (9-12), as have other psychological traits, including trait anger (13) and vital exhaustion, which is characterized by a state of fatigue, lack of energy, feelings of hopelessness, loss of libido, and increased irritability (14;15).

Cardiovascular disease and diabetes share common metabolic precursors (16), so they may also share antecedent psychological risk factors. Previous epidemiological studies of psychological risk factors in diabetes have been limited by small sample sizes and limited physiological data. Vital exhaustion and trait anger have not been evaluated in relation to incident diabetes. ARIC is an attractive study in which to evaluate this relationship because data are available on the above psychological factors in a large cohort of individuals. While ARIC does not have direct assessment of depressive symptoms, data is available on anti-depressant medication use at baseline, which can serve as a proxy for the presence of depression. Previous associations of trait anger and vital exhaustion with incident CHD suggest a potential common soil.

5. Main Hypothesis/Study Questions:

- 1.) Independent predictors of incident type 2 diabetes will include:
 - a.) Vital exhaustion
 - b.) Trait anger
 - c.) Anti-depressant medication use
- 2.) These relationships will be mediated by obesity, insulin resistance (as indicated by hyperinsulinemia), physical inactivity, and/or high dietary energy intake.

6. Data (variables, time window, source, inclusions/exclusions):

Visit 1 variables: anti-depressant medication use, prevalent diabetes, fasting insulin, smoking status

Visit 2 variables: age, race, gender, body-mass index, waist-to-hip ratio, presence of hypertension, fasting glucose, prevalent diabetes, prevalent CHD, HDL-cholesterol, total triglycerides, systolic blood pressure, diastolic blood pressure, anti-hypertensive medication use, anti-depressant medication use, physical activity score, caloric intake, white blood cell count, heart rate variability (R-R interval), responses to part B of Health and Life Profile Questionnaire, responses to part C of Health and Life Profile Questionnaire

Visits 3 and 4 variables: incident diabetes

Individuals with prevalent diabetes at visits 1 and 2 and individuals missing data on response to parts B and C of the Health and Life Profile Questionnaire will be excluded from analysis.

Analysis Plan: Logistic regression and Cox Proportional Hazards modeling

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