ONLINE SUPPLEMENT

AORTIC PULSE WAVE VELOCITY PREDICTS FOCAL WHITE MATTER HYPERINTENSITIES IN A BIRACIAL COHORT OF OLDER ADULTS.

Caterina Rosano, MD, MPH^a; Nora Watson, PhD^a*; YueFang Cheng,PhD^b*; Anne B. Newman, MD, MPH^a; Howard J. Aizenstein, MD, PhD^c; Yan Du, PhD^d; Vijay Venkatraman, PhD^e; Tamara B. Harris, MD, MS^f; Emma Barinas-Mitchell, PhD^a; Kim Sutton-Tyrrell <u>PhD^a</u>

*Contributed equally to preparation of manuscript.

- ^a University of Pittsburgh, Department of Epidemiology; Pittsburgh, PA
- ^b University of Pittsburgh, Department of Neurological Surgery; Pittsburgh, PA

^c University of Pittsburgh, Department of Psychiatry; Pittsburgh, PA

^f National Institute on Aging, Laboratory of Epidemiology, Demography, and Biometry; Bethesda, MD

Corresponding Author: Caterina Rosano, MD, MPH 130 N. Bellefield Ave Room 507 Pittsburgh, PA 15213 Phone: 412-383-1294 Fax: 412-383-1308 Email: rosanoc@edc.pitt.edu

Methods

Study Population

The cohort was followed prospectively to evaluate the relationship of changes in body composition, weight, and related health conditions to incident mobility disability. The cohort consisted of 3,075 well-functioning older black and white men and women living in Pittsburgh, PA and Memphis, TN¹. Eligible participants were 70-79 years of age and reported no difficulty walking a quarter of a mile (400 m), climbing 10 steps, or performing activities of daily living¹.

Pulse Wave Velocity

A minimum of ten beats were recorded for each simultaneous recording run. Three separate runs were recorded for each participant, and all usable runs were averaged to calculate the final PWV measure. The distance between the carotid and femoral recording sites was measured above the surface of the body with a tape measure. The distance was divided by the time delay between the foot of the pressure waves at each site to calculate PWV in cm/s.²

Magnetic Resonance Image Acquisition

The magnetization-prepared rapid gradient echo T1-weighted images and fluid-attenuated inversion recovery images were acquired in the axial plane. Diffusion Weighted Images were acquired using single-short spin-echo sequence.³⁻⁵

White matter hyperintensities volume was obtained from T2-weighted FLAIR image using an automated method and normalized for brain volume^{4, 6}. FA was obtained from Diffusion weighted images⁷, using previously published processing steps³⁻⁵. Using the segmentation of white matter, gray matter, and WMH that were obtained from the T1-weighted and T2-weighted FLAIR images, the FA maps were restricted to normal appearing white matter.

The spatial distribution of white matter tracts was obtained using the JHU atlas⁸. Volume of the gray matter was calculated by segmenting the skull-stripped T1-weighted image in native anatomical space using the FAST - FMRIB's Automated Segmentation Tool⁹. Intracranial volume was measured using a T1-weighted image, BET with advanced option (–A) to extract additional skull and scalp surfaces¹⁰.

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| Population Characteristics | Study entry | Time of MRI | |
|-------------------------------------|-----------------|----------------|--|
| PWV, cm/sec | 840.31 (412.68) | | |
| BMI, m/kg ² | 27.21 (4.54) | 27.27 (4.48) | |
| SBP, mmHg | 137.08 (20.12) | 135.14 (18.32) | |
| DBP, mmHg | 73.77 (9.66) | 69.64 (9.65) | |
| Pulse pressure, mmHg | 63.31 (16.53) | 64.74 (16.28) | |
| Current Smoker, N (%) | 15 (5) | 6 (2) | |
| DSST, points | 42.54 (12.59) | 38.50 (12.66) | |
| MMSE, points | 92.45 (6.24) | 92.94 (8.29) | |
| Diabetes, N (%) | 36 (12) | 83 (29) | |
| Cardiovascular diseases, N (%) | 49 (14) | 83 (27) | |
| Stroke, N (%) | 12 (4) | 24 (8) | |
| Hypertension, N (%) | 146 (48) | 211 (70) | |
| WMH volume, cm ³ | | 6.00 (7.00) | |
| Atrophy , cm³ | | 0.91 (1.41) | |
| Fractional anisotropy | | 0.357 (0.014) | |

Table S1. Population characteristics of this study cohort (n=303) measured at study entry in 1997-98 and concurrent with the MRI in 2006-08. Means and standard deviations (SD) are reported, unless otherwise noted.

| Population Characteristics | Study entry 1997-98 | | Follow-up visit in 2006-07 | |
|--------------------------------|------------------------|----------|-------------------------------|--------------------|
| | Mean | SD | Mean | SD |
| PWV, cm/sec | 902.76 | (393.77) | | |
| BMI, m/kg^2 | 27.7 | (4.8) | 27.6 | (4.7) |
| SBP, mmHg | 138.7 | (21.1) | 134.6 | (20.5) |
| DBP, mmHg | 74.3 | (10.5) | 69.2 | (10.4) |
| Pulse pressure, mmHg | 64.5 | (17.3) | 65.4 | (17.5) |
| Current Smoker, N (%) | 147 | (10.1) | 28 | (3) |
| DSST, points | 37.8 | (13.8) | 35.9 | (13.4) |
| MMSE, points | 90.6 | (7.5) | 91.2 | (8.6) |
| Diabetes, N (%) | 215 | (15) | 412 | (28)^ |
| Cardiovascular diseases, N (%) | 410 | (28) | 534 | (37) |
| Stroke, N (%) | 129 | (10) | 171 | (12) |
| Hypertension, N (%) | 955 | (67) | 91 | (93) ^{\$} |

Table S2. Population characteristics of the parent cohort (N=1455) measured at study entry in 1997-98 and at follow-up in 2006-07. Means and standard deviations (SD) are reported, unless otherwise noted.

N=325 missing; n=178 missing.

| | interest. | Median, cm ³ | 25 th - 75 th percentile cm ³ |
|-------------------------------------|-----------|-------------------------|--|
| Superior Longitudinal Fasciculus | Left | 0.001 | 0014 |
| | Right | 0.001 | 0024 |
| Anterior thalamic radiation | Left | 0.033 | .014071 |
| | Right | 0.060 | .027120 |
| Cortico- spinal tracts | Left | 0.001 | 0006 |
| | Right | 0.001 | 0012 |
| Inferior Longitudinal Fasciculus | Left | 0.005 | 0026 |
| | Right | 0.002 | 0009 |
| Uncinate Fasciculus | Left | 0.008 | .002017 |
| | Right | 0.012 | .003031 |
| Cingulum^ | Left | 0.003 | 0130 |
| | Right | 0 | 0008 |
| Corpus Callosum | Frontal | .032 | .014,.069 |
| | Occipital | .017 | .001, .082 |

Table S3. Mean, 25th and 75th percentile of white matter hyperintensities volume for each tract of interest.

^ Upper and lower portions of the Cingulum are reported combined.





Figure Legend

Figure S1. Axial views of white matter tracts of interest are illustrated in color overlays on a T1weighted MPRAGE in standard space (Montreal Neuroimaging Initiative).