

Online supplement

Hypertensive adults exhibit lower myelin content: A multicomponent relaxometry and diffusion MRI study

John P. Laporte¹, Mary E. Faulkner¹, Zhaoyuan Gong¹, Mohammad A.B.S. Akhonda¹, Luigi Ferrucci², Josephine M. Egan¹, and Mustapha Bouhrara^{1*}

¹Laboratory of Clinical Investigation, National Institute on Aging, National Institutes of Health, Baltimore, 21224 MD, USA.

²Translational Gerontology Branch, National Institute on Aging, National Institutes of Health, Baltimore, 21224 MD, USA.

***Corresponding author:** Mustapha Bouhrara, Magnetic Resonance Physics of Aging and Dementia Unit, Laboratory of Clinical Investigation, National Institute on Aging (NIA), NIH, BRC 05C-222, 251 Bayview Blvd., Baltimore 21224 MD, USA. Tel: 410-558-8541, E-mail: bouhraram@mail.nih.gov

Table S1. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Myelin Water Fraction vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $MWF \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Myelin Water Fraction (MWF)						
	Hypertension β (SE) *10 ⁻²	Age β (SE) *10 ⁻⁴	SBP β (SE) *10 ⁻⁴	Smoking β (SE) *10 ⁻³	Sex β (SE) *10 ⁻³	Diabetes β (SE) *10 ⁻²	Cholesterol β (SE) *10 ⁻⁵
WB	-1.97 (0.68), 0.010	-9.64 (1.76), < 0.001	2.76 (2.17), 0.390	3.10 (5.72), 0.831	-4.44 (5.84), 0.856	0.82 (1.56), 0.898	1.51 (9.14), 0.959
FL	-2.43 (0.67), 0.006	-11.2 (1.73), < 0.001	4.34 (2.14), 0.390	4.52 (5.64), 0.831	-4.89 (5.76), 0.856	0.34 (1.54), 0.898	1.06 (9.00), 0.959
OL	-1.55 (0.87), 0.090	-8.43 (2.23), < 0.001	1.36 (2.76), 0.670	4.26 (7.25), 0.831	-4.04 (7.41), 0.856	1.34 (1.98), 0.898	-0.60 (11.6), 0.959
PL	-2.41 (0.78), 0.010	-10.1 (2.00), < 0.001	3.85 (2.47), 0.390	3.35 (6.50), 0.831	-11.4 (6.64), 0.856	0.41 (1.77), 0.898	-3.44 (10.4), 0.959
TL	-1.84 (0.82), 0.038	-9.28 (2.10), < 0.001	2.16 (2.60), 0.570	3.99 (6.84), 0.831	-3.86 (6.98), 0.856	1.57 (1.86), 0.898	2.56 (10.9), 0.959
CRB	-0.50 (0.73), 0.500	-6.70 (1.89), < 0.001	0.02 (2.33), 0.994	-5.70 (6.14), 0.831	1.69 (6.27), 0.856	1.76 (1.67), 0.898	4.81 (9.80), 0.959
CC	-2.06 (0.69), 0.010	-9.01 (1.77), < 0.001	2.91 (2.19), 0.390	-1.65 (5.76), 0.836	-7.33 (5.88), 0.856	1.24 (1.57), 0.898	2.47 (9.20), 0.959
IC	-1.40 (0.71), 0.067	-9.61 (1.83), < 0.001	1.26 (2.26), 0.670	2.71 (5.94), 0.831	1.72 (6.07), 0.856	1.04 (1.62), 0.898	7.69 (9.49), 0.959
CR	-3.03 (0.88), 0.006	-11.8 (2.27), < 0.001	4.39 (2.80), 0.390	0.24 (7.37), 0.974	-2.25 (7.53), 0.830	-0.33 (2.01), 0.898	2.76 (11.8), 0.959
CP	-0.70 (0.63), 0.285	-4.07 (1.61), 0.013	-1.21 (1.99), 0.670	4.19 (5.23), 0.831	1.40 (5.34), 0.856	2.25 (1.43), 0.898	10.6 (8.36), 0.959
TR	-2.20 (0.83), 0.015	-10.1 (2.14), < 0.001	2.68 (2.65), 0.488	2.52 (6.96), 0.836	0.36 (7.11), 0.960	0.24 (1.90), 0.898	1.31 (11.1), 0.959
FOF	-2.37 (0.77), 0.010	-10.7 (1.98), < 0.001	3.33 (2.45), 0.390	4.31 (6.44), 0.831	4.36 (6.58), 0.856	-0.24 (1.76), 0.898	4.85 (10.3), 0.959
LF	-2.28 (0.80), 0.010	-9.92 (2.06), < 0.001	3.37 (2.55), 0.390	3.02 (6.70), 0.831	-3.19 (6.85), 0.856	0.25 (1.83), 0.898	1.44 (10.7), 0.959
Fr	-2.24 (0.76), 0.010	-10.6 (1.95), < 0.001	2.96 (2.41), 0.390	3.26 (6.35), 0.831	-3.68 (6.49), 0.856	1.04 (1.73), 0.898	-1.92 (10.1), 0.959

MWF, myelin water fraction; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S2. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Longitudinal relaxation rate (R_1) vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $R_1 \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Longitudinal relaxation rate (R_1)						
	Hypertension β (SE) *10 ⁻⁵	Age β (SE) *10 ⁻⁶	SBP β (SE) *10 ⁻⁶	Smoking β (SE) *10 ⁻⁵	Sex β (SE) *10 ⁻⁵	Diabetes β (SE) *10 ⁻⁵	Cholesterol β (SE) *10 ⁻⁷
WB	-5.76 (1.74), 0.004	-1.71 (0.45), < 0.001	1.18 (0.55), 0.075	0.13 (1.46), 0.987	-2.53 (1.49), 0.215	-2.19 (3.97), 0.989	0.44 (2.32), 0.999
FL	-6.77 (1.61), 0.001	-2.06 (0.41), < 0.001	1.60 (0.51), 0.034	0.91 (1.34), 0.987	-2.66 (1.37), 0.215	-3.65 (3.66), 0.989	0.67 (2.15), 0.999
OL	-4.18 (1.98), 0.043	-1.41 (0.51), 0.008	0.70 (0.63), 0.315	0.34 (1.66), 0.987	-2.12 (1.70), 0.260	-1.11 (4.53), 0.989	-1.28 (2.65), 0.999
PL	-6.05 (1.75), 0.003	-1.87 (0.45), < 0.001	1.22 (0.56), 0.075	0.25 (1.47), 0.987	-3.41 (1.50), 0.215	-3.60 (4.00), 0.989	-2.05 (2.34), 0.999
TL	-4.91 (1.83), 0.011	-1.77 (0.47), < 0.001	1.05 (0.58), 0.097	-0.25 (1.53), 0.987	-2.36 (1.56), 0.250	-0.06 (4.17), 0.989	-0.87 (2.44), 0.999
CRB	-2.63 (1.78), 0.145	-1.09 (0.46), 0.021	0.45 (0.57), 0.433	-1.10 (1.49), 0.987	-2.64 (1.52), 0.215	-0.06 (4.07), 0.989	0.02 (2.38), 0.999
CC	-7.41 (1.82), 0.001	-1.90 (0.47), < 0.001	1.35 (0.58), 0.075	-1.10 (1.52), 0.987	-3.39 (1.55), 0.215	-1.86 (4.14), 0.989	-0.15 (2.43), 0.999
IC	-4.91 (1.74), 0.008	-1.77 (0.45), < 0.001	1.18 (0.55), 0.075	-0.57 (1.45), 0.987	-1.99 (1.49), 0.259	-0.76 (3.97), 0.989	-0.19 (2.32), 0.999
CR	-7.79 (1.93), 0.001	-2.16 (0.50), < 0.001	1.30 (0.61), 0.075	0.10 (1.61), 0.987	-2.02 (1.65), 0.830	-4.19 (4.40), 0.989	-0.83 (2.58), 0.999
CP	-3.70 (1.77), 0.043	-0.50 (0.46), 0.273	0.57 (0.56), 0.336	0.27 (1.48), 0.987	-1.48 (1.51), 0.356	-0.26 (4.04), 0.989	1.11 (2.37), 0.999
TR	-5.70 (1.88), 0.006	-2.04 (0.48), < 0.001	1.07 (0.60), 0.097	-0.50 (1.57), 0.987	-2.16 (1.61), 0.259	-2.27 (4.29), 0.989	-1.27 (2.51), 0.999
FOF	-5.95 (1.85), 0.004	-2.09 (0.48), < 0.001	1.14 (0.59), 0.086	0.63 (1.55), 0.987	-0.85 (1.58), 0.592	-2.89 (4.22), 0.989	-0.02 (2.47), 0.999
LF	-5.63 (1.88), 0.006	-2.02 (0.49), < 0.001	1.33 (0.60), 0.075	0.14 (1.58), 0.987	-2.38 (1.61), 0.250	-3.06 (4.30), 0.989	-1.58 (2.52), 0.999
Fr	-6.12 (1.87), 0.004	-2.02 (0.48), < 0.001	1.21 (0.59), 0.080	-0.07 (1.56), 0.987	-2.93 (1.59), 0.215	-1.57 (4.26), 0.989	-1.40 (2.49), 0.999

R_1 , longitudinal relaxation rate; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S3. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Transverse relaxation rate (R_2) vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $R_2 \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Transverse relaxation rate (R_2)						
	Hypertension β (SE) * 10^{-3}	Age β (SE) * 10^{-5}	SBP β (SE) * 10^{-5}	Smoking β (SE) * 10^{-4}	Sex β (SE) * 10^{-4}	Diabetes β (SE) * 10^{-3}	Cholesterol β (SE) * 10^{-6}
WB	-1.20 (0.46), 0.021	-6.12 (1.17), < 0.001	0.99 (1.45), 0.786	-0.33 (3.81), 0.931	-3.13 (3.89), 0.870	0.82 (1.04), 0.829	3.55 (6.09), 0.940
FL	-1.45 (0.45), 0.009	-6.95 (1.15), < 0.001	2.11 (1.42), 0.786	1.80 (3.74), 0.931	-2.59 (3.82), 0.870	0.61 (1.02), 0.829	2.49 (5.97), 0.940
OL	-1.19 (0.57), 0.052	-5.61 (1.47), < 0.001	0.32 (1.82), 0.861	-1.23 (4.79), 0.931	-5.52 (4.89), 0.870	0.93 (1.31), 0.829	-0.56 (7.65), 0.942
PL	-1.35 (0.53), 0.021	-7.06 (1.37), < 0.001	1.84 (1.70), 0.786	-2.97 (4.46), 0.931	-9.07 (4.56), 0.700	0.36 (1.22), 0.846	-1.38 (7.13), 0.940
TL	-1.28 (0.54), 0.027	-5.55 (1.38), < 0.001	1.11 (1.71), 0.786	0.59 (4.50), 0.931	-3.73 (4.59), 0.870	1.26 (1.23), 0.829	3.56 (7.18), 0.940
CRB	-0.20 (0.51), 0.693	-4.71 (1.30), < 0.001	0.43 (1.61), 0.861	-4.20 (4.24), 0.931	0.88 (4.33), 0.903	0.52 (1.16), 0.829	4.19 (6.77), 0.940
CC	-1.72 (0.55), 0.009	-6.78 (1.42), < 0.001	1.02 (1.76), 0.786	-3.86 (4.62), 0.931	-3.07 (4.72), 0.870	1.80 (1.26), 0.829	6.15 (7.39), 0.940
IC	-0.84 (0.43), 0.065	-5.48 (1.11), < 0.001	0.87 (1.37), 0.786	2.85 (3.61), 0.931	0.17 (3.68), 0.963	0.99 (0.98), 0.829	5.37 (5.76), 0.940
CR	-1.71 (0.53), 0.009	-5.84 (1.37), < 0.001	1.31 (1.70), 0.786	0.46 (4.47), 0.931	1.79 (4.56), 0.830	0.24 (1.22), 0.846	5.44 (7.13), 0.940
CP	-0.31 (0.48), 0.561	-3.54 (1.23), < 0.001	-1.11 (1.52), 0.786	1.73 (3.99), 0.931	-1.81 (4.08), 0.870	1.87 (1.09), 0.829	7.63 (6.37), 0.940
TR	-1.38 (0.53), 0.021	-5.34 (1.37), 0.005	0.43 (1.69), 0.861	2.33 (4.44), 0.931	2.31 (4.53), 0.870	0.64 (1.21), 0.829	3.24 (7.09), 0.940
FOF	-1.52 (0.51), 0.010	-6.71 (1.30), < 0.001	1.34 (1.61), 0.786	3.17 (4.24), 0.931	3.32 (4.33), 0.870	0.30 (1.16), 0.846	5.27 (6.77), 0.940
LF	-1.29 (0.50), 0.021	-5.35 (1.29), < 0.001	0.53 (1.59), 0.861	1.38 (4.18), 0.931	-2.71 (4.27), 0.870	0.59 (1.14), 0.829	1.77 (6.68), 0.940
Fr	-1.57 (0.51), 0.009	-6.38 (1.30), < 0.001	1.41 (1.61), 0.786	0.62 (4.23), 0.931	-1.41 (4.32), 0.870	0.97 (1.15), 0.829	1.09 (6.76), 0.940

R_2 , transverse relaxation rate; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S4. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Fractional Anisotropy vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $FA \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Fractional Anisotropy (FA)						
	Hypertension β (SE) * 10^{-2}	Age β (SE) * 10^{-4}	SBP β (SE) * 10^{-4}	Smoking β (SE) * 10^{-3}	Sex β (SE) * 10^{-3}	Diabetes β (SE) * 10^{-3}	Cholesterol β (SE) * 10^{-5}
WB	-1.20 (0.53), 0.041	-6.87 (1.36), < 0.001	0.48 (1.66), 0.774	-3.78 (4.42), 0.884	-2.98 (4.50), 0.830	-2.99 (11.8), 0.931	-0.93 (6.95), 0.950
FL	-2.51 (0.51), < 0.001	-8.65 (1.31), < 0.001	3.34 (1.60), 0.089	-0.62 (4.26), 0.884	-6.37 (4.34), 0.830	-7.57 (11.4), 0.931	-2.56 (6.70), 0.950
OL	-1.81 (0.56), 0.006	-10.3 (1.44), < 0.001	3.38 (1.77), 0.094	3.42 (4.70), 0.884	1.56 (4.79), 0.830	-3.57 (12.5), 0.931	11.3 (7.39), 0.676
PL	-1.08 (0.78), 0.168	-5.02 (2.00), 0.015	5.16 (2.45), 0.089	6.23 (6.50), 0.884	2.53 (6.63), 0.830	8.61 (17.4), 0.931	12.1 (10.2), 0.676
TL	-5.52 (1.36), < 0.001	-19.3 (3.49), < 0.001	8.74 (4.28), 0.089	-13.2 (11.4), 0.884	-3.14 (11.6), 0.830	-19.4 (30.3), 0.931	-2.99 (17.9), 0.950
CRB	-2.50 (1.14), 0.044	-7.99 (2.93), 0.009	8.94 (3.59), 0.089	5.33 (9.53), 0.884	2.09 (9.71), 0.830	-13.9 (25.4), 0.931	21.9 (15.0), 0.676
CC	-1.87 (0.70), 0.021	-11.8 (1.79), < 0.001	4.18 (2.19), 0.094	1.22 (5.81), 0.884	5.74 (5.93), 0.830	-10.9 (15.5), 0.931	0.58 (9.15), 0.950
IC	-2.60 (1.14), 0.041	-6.35 (2.93), 0.033	1.56 (3.59), 0.717	-1.52 (9.53), 0.884	3.91 (9.71), 0.830	3.14 (25.4), 0.931	7.75 (15.0), 0.950
CR	-2.04 (0.64), 0.006	-10.4 (1.64), < 0.001	3.72 (2.01), 0.094	2.28 (5.32), 0.884	10.2 (5.43), 0.830	-10.2 (14.2), 0.931	10.9 (8.38), 0.676
CP	-1.64 (0.92), 0.091	-7.91 (2.35), 0.001	6.06 (2.88), 0.089	1.97 (7.64), 0.884	3.55 (7.79), 0.830	-3.50 (20.4), 0.931	8.65 (12.0), 0.950
TR	-1.76 (0.77), 0.041	-6.86 (1.98), 0.001	3.96 (2.43), 0.125	7.70 (6.45), 0.884	7.40 (6.58), 0.830	-5.60 (17.2), 0.931	16.3 (10.2), 0.676
FOF	-3.24 (0.90), 0.002	-15.0 (2.29), < 0.001	4.98 (2.81), 0.103	-5.96 (7.47), 0.884	-2.18 (7.61), 0.830	-4.38 (19.9), 0.931	2.98 (11.8), 0.950
LF	-6.49 (0.34), 0.074	-7.54 (0.87), < 0.001	2.38 (1.06), 0.089	-1.66 (2.82), 0.884	-3.21 (2.87), 0.830	-5.02 (7.52), 0.931	-1.29 (4.43), 0.950
Fr	-6.92 (0.45), 0.136	-8.91 (1.15), < 0.001	3.01 (1.41), 0.089	-1.41 (3.74), 0.884	-2.02 (3.81), 0.830	0.16 (9.98), 0.988	-3.94 (5.88), 0.950

FA, fractional anisotropy; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S5. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Medial Diffusivity vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $MD \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Medial Diffusivity (MD)							
	Hypertension β (SE) *10 ⁻⁵	Age β (SE) *10 ⁻⁶	SBP β (SE) *10 ⁻⁶	Smoking β (SE) *10 ⁻⁵	Sex β (SE) *10 ⁻⁵	Diabetes β (SE) *10 ⁻⁵	Cholesterol β (SE) *10 ⁻⁷	
WB	1.68 (1.47), 0.276	2.46 (0.38), <0.001	-0.58 (0.46), 0.231	1.39 (1.23), 0.525	4.47 (1.25), 0.009	-0.37 (3.28), 0.912	-1.44 (1.93), 0.801	
FL	2.77 (1.67), 0.128	3.92 (0.43), <0.001	-1.32 (0.52), 0.066	2.02 (1.39), 0.431	2.97 (1.42), 0.114	-1.90 (3.71), 0.912	-0.76 (2.19), 0.801	
OL	2.99 (0.91), 0.021	2.03 (0.23), <0.001	-0.75 (0.29), 0.066	0.44 (0.76), 0.653	2.02 (0.77), 0.076	-0.35 (2.03), 0.912	-0.54 (1.19), 0.801	
PL	2.21 (1.52), 0.175	0.67 (0.39), 0.089	-0.81 (0.48), 0.148	0.86 (1.27), 0.638	-0.40 (1.29), 0.759	4.45 (3.39), 0.484	-0.77 (1.99), 0.801	
TL	5.67 (2.12), 0.032	2.23 (0.54), <0.001	-1.68 (0.67), 0.066	2.37 (1.77), 0.431	1.02 (1.81), 0.661	-1.01 (4.73), 0.912	-2.40 (2.79), 0.801	
CRB	4.55 (2.01), 0.042	2.95 (0.52), <0.001	-1.16 (0.63), 0.121	1.64 (1.68), 0.546	2.12 (1.71), 0.339	-1.62 (4.47), 0.912	-1.50 (2.64), 0.801	
CC	5.79 (2.02), 0.030	1.98 (0.52), <0.001	-0.95 (0.63), 0.164	1.58 (1.68), 0.546	0.87 (1.71), 0.661	-7.33 (4.49), 0.484	-1.24 (2.65), 0.801	
IC	4.05 (1.80), 0.042	2.34 (0.46), <0.001	-1.10 (0.56), 0.121	-1.09 (1.50), 0.638	3.15 (1.53), 0.114	0.69 (4.00), 0.912	1.09 (2.36), 0.801	
CR	2.97 (1.21), 0.037	1.32 (0.31), <0.001	-0.16 (0.38), 0.670	0.05 (1.01), 0.962	0.69 (1.03), 0.830	4.07 (2.69), 0.484	0.19 (1.59), 0.801	
CP	5.01 (1.78), 0.030	2.45 (0.46), <0.001	-1.25 (0.56), 0.099	2.21 (1.49), 0.431	1.01 (1.52), 0.645	9.11 (3.98), 0.345	-0.77 (2.34), 0.801	
TR	2.82 (1.11), 0.036	1.11 (0.28), <0.001	-0.66 (0.35), 0.121	0.33 (0.93), 0.782	1.60 (0.94), 0.186	3.29 (2.47), 0.484	0.63 (1.46), 0.801	
FOF	3.52 (1.56), 0.042	1.36 (0.40), <0.001	-0.73 (0.49), 0.164	1.74 (1.30), 0.431	2.66 (1.33), 0.114	1.09 (3.48), 0.912	-1.25 (2.05), 0.801	
LF	3.75 (2.14), 0.119	6.10 (0.55), <0.001	-1.35 (0.67), 0.121	2.49 (1.79), 0.431	4.00 (1.82), 0.114	-3.30 (4.78), 0.912	-2.53 (2.81), 0.801	
Fr	2.16 (2.98), 0.470	7.84 (0.76), <0.001	-1.43 (0.94), 0.164	3.38 (2.49), 0.431	3.44 (2.54), 0.313	-8.44 (6.64), 0.484	-1.41 (3.91), 0.801	

MD, medial diffusivity; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S6. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Radial Diffusivity vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $RD \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Radial Diffusivity (RD)							
	Hypertension β (SE) *10 ⁻⁵	Age β (SE) *10 ⁻⁶	SBP β (SE) *10 ⁻⁶	Smoking β (SE) *10 ⁻⁵	Sex β (SE) *10 ⁻⁵	Diabetes β (SE) *10 ⁻⁵	Cholesterol β (SE) *10 ⁻⁷	
WB	2.62 (1.60), 0.122	2.87 (0.41), <0.001	-0.59 (0.50), 0.261	-1.63 (1.33), 0.522	4.42 (1.36), 0.023	-0.57 (3.55), 0.940	-1.47 (2.09), 0.842	
FL	4.29 (1.76), 0.026	4.34 (0.45), <0.001	-1.41 (0.55), 0.059	1.99 (1.47), 0.522	3.29 (1.49), 0.145	-1.71 (3.91), 0.940	-0.42 (2.31), 0.878	
OL	3.99 (1.13), 0.002	2.75 (0.29), <0.001	-0.93 (0.35), 0.059	0.34 (0.94), 0.972	1.77 (0.96), 0.241	-0.42 (2.51), 0.940	-1.51 (1.48), 0.842	
PL	1.84 (1.46), 0.229	1.03 (0.37), 0.004	-0.84 (0.46), 0.092	-0.10 (1.22), 0.972	-0.77 (1.24), 0.628	3.02 (3.25), 0.875	-1.51 (1.92), 0.842	
TL	10.1 (2.60), 0.002	3.75 (0.67), <0.001	-1.88 (0.82), 0.064	2.71 (2.17), 0.522	1.39 (2.21), 0.628	1.55 (5.78), 0.940	-2.01 (3.41), 0.842	
CRB	5.21 (2.17), 0.026	2.40 (0.56), <0.001	-1.53 (0.68), 0.064	1.04 (1.81), 0.884	1.74 (1.85), 0.548	0.11 (4.83), 0.981	-2.84 (2.85), 0.842	
CC	6.10 (1.85), 0.004	2.42 (0.47), <0.001	-1.07 (0.58), 0.092	1.17 (1.55), 0.790	0.13 (1.58), 0.989	6.45 (4.13), 0.688	-1.20 (2.43), 0.842	
IC	6.02 (1.90), 0.005	2.33 (0.49), <0.001	-1.13 (0.60), 0.092	-0.22 (1.59), 0.972	2.59 (1.62), 0.264	-2.63 (4.24), 0.940	-0.38 (2.50), 0.878	
CR	4.27 (1.21), 0.002	1.84 (0.31), <0.001	-0.39 (0.38), 0.310	0.09 (1.01), 0.972	0.01 (1.03), 0.830	3.94 (2.69), 0.688	-0.71 (1.59), 0.842	
CP	5.30 (1.78), 0.007	2.49 (0.46), <0.001	-1.34 (0.56), 0.064	1.62 (1.48), 0.555	0.94 (1.51), 0.628	6.23 (3.96), 0.688	-1.65 (2.33), 0.842	
TR	3.76 (1.06), 0.002	1.41 (0.27), <0.001	-0.88 (0.33), 0.059	-0.03 (0.89), 0.972	0.85 (0.91), 0.548	2.12 (2.37), 0.875	-0.93 (1.40), 0.842	
FOF	6.11 (1.94), 0.005	2.60 (0.50), <0.001	-1.16 (0.61), 0.092	2.42 (1.62), 0.522	2.85 (1.65), 0.248	1.46 (4.33), 0.940	-1.12 (2.55), 0.842	
LF	3.86 (2.13), 0.094	6.35 (0.55), <0.001	-1.42 (0.67), 0.073	2.39 (1.78), 0.522	3.98 (1.81), 0.145	-2.90 (4.75), 0.940	-2.27 (2.80), 0.842	
Fr	2.24 (3.02), 0.460	8.21 (0.77), <0.001	-1.51 (0.95), 0.136	3.25 (2.52), 0.522	3.32 (2.57), 0.398	-8.00 (6.72), 0.832	0.94 (3.96), 0.878	

RD, radial diffusivity; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.

Table S7. Regression coefficient (β), including standard error (SE) in italics, and significance (p -value) of Axial Diffusivity vs. Hypertension, age, smoking, sex, diabetes, and cholesterol across 14 WM ROIs. The multiple regression model is given by: $AxD \sim \beta_0 + \beta_{age} \times age + \beta_{Hypertension} \times Hypertension + \beta_{SBP} \times SBP + \beta_{Smoking} \times Smoking + \beta_{sex} \times sex + \beta_{Diabetes} \times Diabetes + \beta_{Cholesterol} \times Cholesterol$. The regression model accounted for sex, smoking status, diabetes status and hypertension as categorical variables. Bolded p -values indicate statistical significance ($p < 0.05$) or close-to-significance ($p < 0.1$).

ROIs	Axial Diffusivity (AxD)						
	Hypertension β (SE) *10 ⁻⁵	Age β (SE) *10 ⁻⁶	SBP β (SE) *10 ⁻⁶	Smoking β (SE) *10 ⁻⁵	Sex β (SE) *10 ⁻⁵	Diabetes β (SE) *10 ⁻⁵	Cholesterol β (SE) *10 ⁻⁷
WB	3.07 (1.70), 0.086	3.06 (0.43), < 0.001	-0.58 (0.53), 0.277	1.77 (1.41), 0.609	4.38 (1.44), 0.045	-0.64 (3.78), 0.867	-1.46 (2.23), 0.905
FL	5.01 (1.83), 0.012	4.54 (0.47), < 0.001	-1.44 (0.58), 0.069	1.93 (1.53), 0.609	3.38 (1.56), 0.158	-1.49 (4.08), 0.867	-0.29 (2.40), 0.905
OL	4.47 (1.29), 0.003	3.11 (0.33), < 0.001	-1.00 (0.40), 0.069	0.33 (1.07), 0.883	1.64 (1.09), 0.385	-0.53 (2.87), 0.867	-1.95 (1.69), 0.905
PL	1.61 (1.59), 0.336	1.21 (0.41), 0.004	-0.92 (0.50), 0.098	-0.30 (1.32), 0.883	-1.16 (1.35), 0.634	1.83 (3.54), 0.867	-1.67 (2.08), 0.905
TL	12.3 (3.16), 0.002	4.43 (0.81), < 0.001	-2.01 (0.99), 0.081	2.80 (2.63), 0.678	1.54 (2.68), 0.763	3.32 (7.03), 0.867	-1.75 (4.14), 0.905
CRB	5.54 (2.36), 0.030	2.10 (0.60), < 0.001	-1.71 (0.74), 0.069	0.78 (1.97), 0.883	1.67 (2.01), 0.634	-0.97 (5.25), 0.867	-3.46 (3.09), 0.905
CC	6.27 (1.83), 0.003	2.62 (0.47), < 0.001	-1.14 (0.57), 0.081	0.92 (1.53), 0.883	-0.24 (1.56), 0.880	6.20 (4.07), 0.867	-1.12 (2.40), 0.905
IC	7.07 (2.22), 0.004	2.27 (0.57), < 0.001	-1.16 (0.70), 0.124	0.31 (1.85), 0.883	2.57 (1.89), 0.415	-4.42 (4.95), 0.867	-0.73 (2.92), 0.905
CR	4.90 (1.30), 0.002	2.09 (0.33), < 0.001	-0.49 (0.41), 0.256	0.16 (1.09), 0.883	-0.39 (1.11), 0.830	3.89 (2.90), 0.867	-1.09 (1.71), 0.905
CP	5.36 (1.91), 0.011	2.52 (0.49), < 0.001	-1.37 (0.60), 0.069	1.28 (1.59), 0.847	0.86 (1.62), 0.763	4.52 (4.25), 0.867	-2.13 (2.50), 0.905
TR	4.22 (1.22), 0.003	1.56 (0.31), < 0.001	-0.98 (0.38), 0.069	-0.19 (1.02), 0.883	0.45 (1.04), 0.776	1.38 (2.71), 0.867	-1.75 (1.60), 0.905
FOF	7.40 (2.21), 0.003	3.21 (0.57), < 0.001	-1.38 (0.69), 0.081	2.73 (1.84), 0.609	2.93 (1.88), 0.385	1.72 (4.92), 0.867	-0.96 (2.90), 0.905
LF	3.87 (2.14), 0.086	6.47 (0.55), < 0.001	-1.47 (0.67), 0.074	2.36 (1.78), 0.609	3.92 (1.82), 0.158	-2.80 (4.76), 0.867	-2.13 (2.80), 0.905
Fr	2.23 (3.05), 0.465	8.41 (0.78), < 0.001	-1.56 (0.96), 0.124	3.16 (2.54), 0.609	3.21 (2.59), 0.439	-8.03 (6.79), 0.867	-0.70 (4.00), 0.905

AxD, axial diffusivity; WM, white matter; ROI, region-of-interest; WB, whole brain; FL, frontal lobe; OL, occipital lobe; PL, parietal lobe; TL, temporal lobe; CRB, cerebellum; IC, internal capsule; CR, corona radiata; CC, corpus callosum; TR, thalamic radiation; FOF, fronto-occipital fasciculus; LF, longitudinal fasciculus; Fr, forceps.