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Supplementary Table 1. Spearman correlation coefficient matrix of sRAGE with selected baseline variables.

	N	Spearman's rho
Age (years)	1201	-0.0078
Fasting Glucose (mg/dL)	1168	-0.218
Hemoglobin A1c (%)	1185	-0.2187
Total Cholesterol (mg/dL)	1201	-0.0491
HDL-Cholesterol (mg/dL)	1201	0.0636
Average Systolic BP (mmHg)	1201	-0.1831
Body Mass Index (kg/m^2)	1201	-0.3246
C-reactive Protein (mg/L)	1201	-0.2548
White Blood Cell Count (x1000/mm3)	1200	-0.1172

Supplementary Table 2. Unadjusted and adjusted Prevalence Ratios (PRs) for the lowest quartile of sRAGE.

	Unadjusted PR	95% CI for unadjusted PR	Adjusted PR	95% CI for adjusted PR
Age - per 10 years	1.02	(0.84, 1.24)	1.05	(0.85, 1.29)
Male (vs female)	1.32	(1.05, 1.66)*	1.53	(1.19, 1.96)*
Black (vs white)	3.07	(2.44, 3.85)*	2.78	(2.17, 3.55)*
Cigarette Smoking				
Current Smoker (vs never)	0.95	(0.67, 1.34)	0.97	(0.67, 1.41)
Former Smoker (vs never)	1.41	(1.10, 1.81)*	1.40	(1.08, 1.83)*
Total Cholesterol ≥ 200 mg/dL (vs < 200 mg/dL)	1.03	(0.82, 1.30)	1.10	(0.87, 1.39)
Low HDL Cholesterol†	0.98	(0.78, 1.24)	0.91	(0.72, 1.16)
Hypertension‡	1.71	(1.37, 2.15)*	1.09	(0.85, 1.39)
Body Mass Index (BMI)				
25-30 kg/m^2 (vs < 25 kg/m^2)	1.69	(1.21, 2.35)*	1.33	(0.94, 1.87)
≥ 30 kg/m^2 (vs < 25 kg/m^2)	3.00	(2.18, 4.12)*	1.90	(1.33, 2.73)*
C-reactive protein				
1-3 mg/L (vs < 1 mg/L)	1.69	(1.19, 2.39)*	1.48	(1.04, 2.10)*
≥3 mg/L (vs < 1 mg/L)	2.51	(1.82, 3.47)*	1.93	(1.35, 2.75)*
Diabetes§	1.68	(1.26, 2.24)*	0.95	(0.70, 1.30)

* p-value < 0.05
† Low HDL-cholesterol was defined as < 40 mg/dL in males and < 50 mg/dL in females.
‡ Diastolic blood pressure ≥ 90 mmHg, systolic blood pressure ≥ 140 mmHg, or blood pressure-lowering medication use

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§ Non-fasting glucose ≥ 200 mg/dL, fasting glucose ≥ 126 mg/dL, self-reported diagnosis, medication use, or HbA1c ≥ 6.5%

Supplementary Table 3. Hazard ratios by incident diabetes, coronary heart disease, ischemic stroke, and mortality by sRAGE quartile excluding cases with diagnosed and undiagnosed diabetes at baseline.

	# of Events	HR Model 1†	HR Model 2‡
Incident Diabetes (N=1045)			
Q4 [1263.4, 4650.4]	45	1 (ref)	1 (ref)
Q3 [965.6-1263.4]	61	1.51 (1.02, 2.22)*	1.31 (0.89, 1.94)
Q2 [711.6-966.3]	63	1.68 (1.14, 2.48)*	1.44 (0.98, 2.13)
Q1 [119.4-711.6]	73	2.24 (1.51, 3.33)*	1.61 (1.08, 2.42)*
p-value for trend		<0.001*	0.018*
CHD (N=1051)			
Q4 [1263.4, 4650.4]	32	1 (ref)	1 (ref)
Q3 [965.6-1263.4]	35	1.05 (0.65, 1.70)	1.11 (0.68, 1.80)
Q2 [711.6-966.3]	35	1.08 (0.66, 1.75)	1.13 (0.70, 1.84)
Q1 [119.4-711.6]	48	1.65 (1.03, 2.65)*	2.02 (1.24, 3.29)*
p-value for trend		0.057	0.012*
Ischemic Stroke (N=1051)			
Q4 [1263.4, 4650.4]	9	1 (ref)	1 (ref)
Q3 [965.6-1263.4]	10	0.99 (0.40, 2.46)	1.02 (0.41, 2.53)
Q2 [711.6-966.3]	4	0.32 (0.10, 1.09)	0.33 (0.10, 1.10)
Q1 [119.4-711.6]	9	0.61 (0.22, 1.70)	0.65 (0.23, 1.84)
p-value for trend		0.168	0.196
All-Cause Mortality (N=1051)			
Q4 [1263.4, 4650.4]	30	1 (ref)	1 (ref)
Q3 [965.6-1263.4]	31	0.98 (0.59, 1.63)	0.98 (0.59, 1.63)
Q2 [711.6-966.3]	39	1.19 (0.73, 1.94)	1.18 (0.73, 1.92)
Q1 [119.4-711.6]	56	1.78 (1.11, 2.86)*	1.77 (1.08, 2.89)*
p-value for trend		0.014*	0.022*

* p-value < 0.05
† Model 1: Adjusted for age, race, sex
‡ Model 2: Adjusted for Framingham Risk Score (which includes age (years), sex, current smoker, systolic blood pressure (mmHg), total cholesterol (mg/dL), HDL cholesterol (mg/dL), diabetes status), plus c-reactive protein (mg/L), BMI (kg/m^2), and race group

Supplementary Table 4. Hazard ratios by incident diabetes, coronary heart disease, ischemic stroke, and mortality by sRAGE quartile excluding cases with diagnosed and undiagnosed diabetes at baseline, Black Race Only.

	# of Events	HR Model 1†	HR Model 2‡
Incident Diabetes (N=207)			
Q4 [1263.4, 4650.4]	5	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	7	0.80 (0.25, 2.53)	0.82 (0.26, 2.59)
Q2 [711.6-966.3)	17	1.29 (0.47, 3.50)	1.15 (0.42, 3.15)
Q1 [119.4-711.6)	33	1.34 (0.52, 3.45)	1.22 (0.47, 3.18)
p-value for trend		0.303	0.455
Coronary heart disease (N=261)			
Q4 [1263.4, 4650.4]	6	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	5	0.39 (0.12, 1.28)	0.44 (0.13, 1.44)
Q2 [711.6-966.3)	6	0.23 (0.07, 0.73)*	0.24 (0.07, 0.79)*
Q1 [119.4-711.6)	19	0.39 (0.15, 0.98)*	0.49 (0.19, 1.28)
p-value for trend		0.135	0.314
Ischemic Stroke (N=261)			
Q4 [1263.4, 4650.4]	1	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	2	0.95 (0.09, 10.52)	1.07 (0.10, 11.88)
Q2 [711.6-966.3)	6	1.49 (0.18, 12.60)	0.95 (0.11, 8.40)
Q1 [119.4-711.6)	10	1.27 (0.16, 10.06)	1.21 (0.15, 9.67)
p-value for trend		0.759	0.765
All-Cause Mortality (N=261)			
Q4 [1263.4, 4650.4]	3	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	5	0.73 (0.17, 3.07)	0.83 (0.20, 3.49)
Q2 [711.6-966.3)	12	0.74 (0.21, 2.67)	0.74 (0.20, 2.75)
Q1 [119.4-711.6)	37	1.37 (0.42, 4.49)	1.43 (0.43, 4.75)
p-value for trend		0.139	0.166

* p-value < 0.05
† Model 1: Adjusted for age, race, sex
‡ Model 2: Adjusted for Framingham Risk Score (which includes age (years), sex, current smoker, systolic blood pressure (mmHg), total cholesterol (mg/dL), HDL cholesterol (mg/dL), diabetes status), plus c-reactive protein (mg/L), BMI (kg/m^2), and race group

Supplementary Table 5. Hazard ratios by incident diabetes, coronary heart disease, ischemic stroke, and mortality by sRAGE quartile excluding cases with diagnosed and undiagnosed diabetes at baseline, White Race Only.

	# of Events	HR Model 1†	HR Model 2‡
Incident Diabetes (N=850)			
Q4 [1263.4, 4650.4]	40	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	57	1.71 (1.14, 2.57)*	1.41 (0.93, 2.12)
Q2 [711.6-966.3)	49	1.75 (1.15, 2.66)*	1.42 (0.93, 2.17)
Q1 [119.4-711.6)	45	2.57 (1.67, 3.98)*	1.64 (1.05, 2.55)*
p-value for trend		<0.001*	0.031*
Coronary heart disease (N=940)			
Q4 [1263.4, 4650.4]	31	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	39	1.20 (0.75, 1.93)	1.16 (0.72, 1.88)
Q2 [711.6-966.3)	43	1.50 (0.94, 2.39)	1.48 (0.93, 2.36)
Q1 [119.4-711.6)	43	2.01 (1.26, 3.21)*	2.16 (1.33, 3.50)*
p-value for trend		0.003*	0.002*
Ischemic Stroke (N=940)			
Q4 [1263.4, 4650.4]	11	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	10	0.93 (0.39, 2.20)	0.76 (0.32, 1.82)
Q2 [711.6-966.3)	7	0.76 (0.29, 1.98)	0.57 (0.22, 1.49)
Q1 [119.4-711.6)	6	0.91 (0.33, 2.50)	0.63 (0.22, 1.78)
p-value for trend		0.702	0.254
All-Cause Mortality (N=940)			
Q4 [1263.4, 4650.4]	34	1 (ref)	1 (ref)
Q3 [965.6-1263.4)	38	1.09 (0.68, 1.73)	1.02 (0.64, 1.63)
Q2 [711.6-966.3)	43	1.40 (0.89, 2.20)	1.18 (0.75, 1.87)
Q1 [119.4-711.6)	41	1.85 (1.17, 2.94)*	1.61 (1.00, 2.61)
p-value for trend		0.007*	0.057

* p-value < 0.05
† Model 1: Adjusted for age, race, sex
‡ Model 2: Adjusted for Framingham Risk Score (which includes age (years), sex, current smoker, systolic blood pressure (mmHg), total cholesterol (mg/dL), HDL cholesterol (mg/dL), diabetes status), plus c-reactive protein (mg/L), BMI (kg/m^2), and race group

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Supplementary Figure 1. Kernal density plots of the sRAGE distribution at baseline. A, Overall. B, By hypertension status. C, By diabetes status. D, By Race.