

Supplementary Table 1: Constructs & Variables from LIFE-Adult

Construct	Variable	LIFE-Adults Questionnaire name and field	Timepoint for which measure is available
Variables for exclusion			
Conditions to ensure we are looking at a healthy sample	Life-time diagnosis of MS, PD, epilepsy	Medical anamnesis (T00173) MEDANAM_F0171, MEDANAM_F0179, MEDANAM_F0167	Baseline
		Medical anamnesis (T01228) MEDIZ_AN_F26 MEDIZ_AN_F27 MEDIZ_AN_F31 (free field, used to screen for epilepsy)	Followup
	Previous stroke	Medical anamnesis (T00173)	Baseline
		Cardiovascular anamnesis (T01226)	Followup
		Lesion diagnosed by radiologist (ischemic, hemorrhagic, traumatic lesions)	Baseline & Followup
	Incidental findings/non-usability of MRI	Radiologist's rating ("non usable") Radiologist's rating (incidental finding)	Baseline
		Radiologist's rating ("non usable") Radiologist's rating (incidental finding)	Followup
	Dementia or cognitive impairment	T00043 SIDAM MMSE < 24	Baseline
		T00043 SIDAM MMSE < 24 or dementia diagnosis in medical anamnesis (T01228) MEDIZ_AN_F30	Followup
	Intake of centrally active medication	D00038 ATC codes/groups: M03B MUSCLE RELAXANTS, CENTRALLY ACTING AGENTS N02A OPIOIDS N03 ANTIEPILEPTICS N04 ANTI-PARKINSON DRUGS N05 PSYCHOLEPTICS N06A ANTIDEPRESSANTS	Baseline Followup

		N06B PSYCHOSTIMULANTS, AGENTS USED FOR ADHD AND NOOTROPICS N06D ANTI-DEMENTIA DRUGS (except for N06DX02, Ginkgo folium) N07A PARASYMPATHOMIMETICS	
Used for	Covariates		
All analyses	Baseline Age	Age at first MRI or cognitive assessment	Baseline
	Age Change	Time between first and second MRI assessment	Baseline Followup
	Gender	self-reported binary gender	Baseline
	eTIV	Estimated total intracranial volume	Summary value of Baseline and Followup
Model M1	Hypertensive medication	Medical (T00173) or medication anamnese (D00038) Intake of anti-hypertensive medication (ATC code starting with "C02 "C03", "C07", "C08" "C09") or treatment because of hypertension (MEDANAM_F0039)	Baseline
		Medication or cardiovascular Anamnese (T01226) Intake of anti-hypertensive medication (ATC code starting with "C02 "C03", "C07", "C08" "C09") or treatment because of hypertension (KARD_AN_F10_3)	Followup
Model M2	CES-D (log-transformed)	D00041 CES_D_SCORE_SUM_CES_D	Baseline
		T00013 CES_D_SCORE_SUM_CES_D	Followup
	Education	socioeconomic status (D00140) Binary variable based on whether participant has tertiary degree (SES2_sesbldg)	Baseline
	Predictors of interest		
Blood pressure	Systolic/diastolic blood pressure	T00049 Average of three consecutive measurements	Baseline
		T01170	Followup

		Average of three consecutive measurements	
Visceral adiposity	Waist-to-hip ratio	D00074 Anthropometry	Baseline
		T01169 Anthropometry	Followup
Cerebral small vessel disease	WML (asinh-transformed) Voxelwise probability of new WML	From FLAIR imaging	Baseline Followup
Memory function	CERAD learning & delayed recall & recognition	T00044 CERAD-plus test battery	Baseline
		T00044 CERAD-plus test battery	Followup
Executive function	CERAD verbal fluency (S and animals) TMTB/TMTA	T00044 CERAD-plus test battery T00042 CERAD Animals T00041 TMT	Baseline
			Followup
Processing speed	TMTA	T00041 TMT	Baseline
		T00041 TMT	Followup