Supplementary Materials

Associations between Lifestyle Factors, Physiological Conditions, and Epigenetic Age Acceleration in an Asian Population

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Running title: Lifestyle factors, physiological conditions, and epigenetic age acceleration

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Table S1 The 17 diet-related questions in the TWB questionnaire.

	Dietary Habits/ Food Category	Always	Most of the time	Half of the time	Seldom	Never
1.	When you eat meat (such as pork, beef,	1	2	3	4	5
	mutton, chicken, duck, goose, etc.), do					
	you eat it with fat, suet, or skin?					
2.	When you eat fish or meat, do you prefer	1	2	3	4	5
	cooking it with oil (such as frying, deep-					
	frying, frying, braising, steamed fish					
	topped with oil, etc.)?					
3.	When you eat vegetables, do you prefer	1	2	3	4	5
	cooking them in a stir-fry way?					
4.	When you eat rice or noodles (staple	1	2	3	4	5
	food), do you eat them with marinade,					
	gravy, or lard?					
5.	When you eat soy foods, do you prefer	1	2	3	4	5
	cooking them in a deep-fry way (such as					
	fried tofu, stinky tofu, fried tofu skin,					
	etc.)?					
6.	When you eat bread, do you spread	1	2	3	4	5
	butter, plant-based butter (margarine), or					
	mayonnaise?					
7.	When you have a meal, do you add	1	2	3	4	5
	additional salt, soy sauce, chili sauce, or					
	any other seasoning?					
8.	Are you used to having pickles,	1	2	3	4	5
	fermented tofu, and fermented soybeans					
	as side dishes in a meal?					
9.	Do you eat fruits or vegetables instead of	1	2	3	4	5
	high-fat snacks (such as chips, cakes,					
	doughnuts, etc.) when you have snacks?					
10	. When you prepare meat (such as pork,	1	2	3	4	5
	beef, mutton, chicken, duck, goose, etc.)					
	for a meal, do you cook it roasted or					

Dietary Habits/ Food Category	Always	Most of the time	Half of the time	Seldom	Never
braised instead of deep-frying?					
11. If a food product has a low-fat option	1	2	3	4	5
(such as low-fat ice cream, low-fat milk,					
skim milk, low-fat salad sauce, etc.),					
would you choose it instead of a regular					
product?					
12. Do you eat food with low-sodium	1	2	3	4	5
ingredients (such as low-sodium salt,					
lower-sodium soy sauce, etc.)?					
13. Would you like to eat lower-fat meat	1	2	3	4	5
(such as fish or chicken) instead of					
higher-fat meat (such as beef or pork)?					
14. Would you choose to eat lean meat	1	2	3	4	5
instead of fatty meat?					
15. Would you choose to perform a	1	2	3	4	5
vegetarian and light diet in certain meals					
to reduce the intake of higher-fat food					
such as meat or fat?					
16. Do you eat at least two kinds of	1	2	3	4	5
vegetables a day?					
17. When you have meat, do you	1	2	3	4	5
intentionally eat less?					

This table was Supplementary Table S4 in Lo and Lin (Lo & Lin, 2022).

Table S2 Fifty-two factors not correlated with any epigenetic marker

Males	Females
1243	1231
5.75 (0.86)	5.40 (0.95)
90/1243 (7.24%)	140/1231 (11.37%)
34.76 (4.83)	35.07 (4.15)
122.42 (31.49)	120.42 (33.36)
192.20 (34.17)	197.30 (35.94)
26.03 (11.02)	22.67 (7.64)
0.750 (0.31)	0.619 (0.24)
4.61 (0.25)	4.52 (0.25)
14.01 (3.68)	12.59 (4.29)
32.25 (141.45)	22.70 (89.40)
1.0 second)	
72.83 (18.12)	72.25 (18.07)
70.75 (18.53)	70.99 (20.26)
70.92 (21.21)	69.29 (23.39)
2.78 (1.34)	1.92 (0.97)
4.75 (2.59)	3.09 (1.71)
4.41 (2.45)	2.92 (1.67)
3.19 (1.63)	2.21 (1.19)
1.54 (0.71)	1.08 (0.54)
0.902 (0.41)	0.682 (0.34)
4.60 (2.13)	4.89 (2.09)
1.00 (0.55)	0.753 (0.44)
1.20 (0.56)	0.890 (0.67)
147/1243 (11.83%)	24/1231 (1.95%)
169/1243 (13.60%)	124/1231 (10.07%)
15/1243 (1.21%)	1/1230 (0.08%)
595/1242 (47.91%)	497/1230 (40.41%)
12/690 (1.74%)	11/751 (1.46%)
139/690 (20.14%)	180/752 (23.94%)
	1243 5.75 (0.86) 90/1243 (7.24%) 34.76 (4.83) 122.42 (31.49) 192.20 (34.17) 26.03 (11.02) 0.750 (0.31) 4.61 (0.25) 14.01 (3.68) 32.25 (141.45) 1.0 second) 72.83 (18.12) 70.75 (18.53) 70.92 (21.21) 2.78 (1.34) 4.75 (2.59) 4.41 (2.45) 3.19 (1.63) 1.54 (0.71) 0.902 (0.41) 4.60 (2.13) 1.00 (0.55) 1.20 (0.56) 147/1243 (11.83%) 169/1243 (13.60%) 15/1243 (1.21%) 595/1242 (47.91%) 12/690 (1.74%)

		Males	Females
	Coffee (yes vs. no) h, n/a (%)	247/690 (35.80%)	274/752 (36.44%)
	Tea (yes vs. no) i, n/a (%)	311/690 (45.07%)	244/752 (32.45%)
	Vegetarian (yes vs. no) j, n/a (%)	14/690 (2.03%)	36/752 (4.79%)
	Supper (yes vs. no) k, n/a (%)	255/690 (36.96%)	214/752 (28.46%)
	Eating out (yes vs. no) 1, n/a (%)	661/690 (95.80%)	705/752 (93.75%)
	Supplement (yes vs. no) m, n/a (%)	331/690 (47.97%)	458/752 (60.90%)
	Daily Meals (times)	2.91 (0.34)	2.83 (0.43)
Diet preference			
	D1 ⁿ (1-5 points)	2.87 (1.56)	3.55 (1.40)
	D2 (1-5 points)	2.84 (1.11)	2.97 (1.19)
	D3 (1-5 points)	2.53 (1.02)	2.66 (1.09)
	D4 (1-5 points)	3.70 (1.27)	4.06 (1.08)
	D5 (1-5 points)	3.86 (0.96)	4.06 (0.89)
	D6 (1-5 points)	4.28 (0.99)	4.32 (0.98)
	D7 (1-5 points)	3.72 (1.25)	3.83 (1.22)
	D8 (1-5 points)	4.29 (0.77)	4.29 (0.79)
	D9 (1-5 points)	3.31 (1.49)	3.09 (1.44)
	D10 (1-5 points)	2.65 (1.31)	2.48 (1.32)
	D11 (1-5 points)	3.19 (1.66)	2.77 (1.62)
	D12 (1-5 points)	3.40 (1.64)	3.48 (1.64)
	D13 (1-5 points)	3.15 (1.43)	2.92 (1.42)
	D14 (1-5 points)	3.03 (1.52)	2.65 (1.54)
	D15 (1-5 points)	3.10 (1.42)	2.66 (1.35)
	D16 (1-5 points)	1.61 (0.92)	1.66 (0.96)
	D17 (1-5 points)	3.40 (1.52)	2.95 (1.58)

Note: Data are presented as mean (s.d.) or n/a (%) (n: the number of individuals belonging to this category; a: the total number of individuals responding to this question).

^a Educational attainment: Educational attainment was recorded as an integer ranging from 1 to 7, representing different levels of education. "1" represents no formal education and illiteracy, "2" represents self-taught individuals who are literate, "3" represents the completion of elementary school, "4" represents the completion of junior high school, "5" represents the completion of senior high school or vocational high school, "6" represents the completion of college or technical school, and "7" represents the completion of graduate school or higher education.

^b Drinking: individuals who consumed more than 150 mL of alcoholic beverages per week for at least six

months when participating in the TWB.

- ^c SHS (Secondhand smoke): being exposed to an environment with secondhand smoke (someone smoking nearby) for at least five minutes in the past six months.
- ^d Nut: betel nut chewing in the past six months.
- ^e Sport: exercising for at least 30 minutes thrice a week. Exercise included leisure-time activities such as swimming, jogging, cycling, mountain climbing, dancing, weight training, etc.
- f Drug: taking cough syrup, sedatives, or pain relievers at least once a week within the six months before joining the TWB.
- g Incense: being exposed to incense burning (e.g., during worship or the use of incense powder or rings), mosquito coils (traditional, liquid electric, or electric mosquito repellent), or fragrances (such as essential oils, aromatherapy, air fresheners, sprays, or scented candles) for at least five minutes within the past year before joining the TWB.
- ^h Coffee: coffee drinking thrice a week.
- ⁱ Tea: consuming tea (containing tea leaves, excluding herbal teas) at least once daily within six months before joining the TWB.
- ^j Vegetarian: vegetarian diet for at least six months before joining the TWB.
- ^k Supper: eating supper within an hour before bedtime (including milk and wine).
- ¹ Eating out: eating out at least once in the past month when joining the TWB.
- ^m Supplement: regularly taking vitamins, minerals, or supplements in the past month before joining the TWB.
- ⁿ D1-D17: These 17 questions are included in the diet-related questionnaires listed in Supplementary Table S1.

Table S3 Best-subset selection for 7 DNA methylation-based markers (22 factors passing the partial correlation filtering serve as candidate predictors)

	DNA methylation-based markers								
	HannumEAA	IEAA	PhenoEAA	GrimEAA	DNAmPACKYRS	DNAmPAI1	DunedinPACE		
	14 factors	8 factors	14 factors	15 factors	12 factors	16 factors	17 factors		
Factors									
Chronological age	V		V		V	V	V		
Sex	V	V	V	V	V	V			
BMI	V		V	V	V	V	V		
WHR	V		V	V	V	V	V		
Smoking	V	V	V	V	V	V	V		
SBP						V	V		
DBP	V	V	V						
Fasting glucose	V					V	V		
HbA1c			V	V	V	V	V		
Triglyceride						V	V		
HDL-C				V	V	V	V		
SGPT									
Creatinine	V			V			V		
Uric acid			V	V		V			
GGT			V	V	V	V	V		
Hemoglobin	V		V	V			V		
Hematocrit	V				V		V		
B lymphocytes	V	V	V	V		V			
Natural killer cells	V	V		V	V	V	V		
CD4 ⁺ T cells	V	V	V	V			V		
CD8 ⁺ T cells	V	V	V	V	V	V	V		
Monocytes		V	V	V	V	V	V		
Explanatory variable	Explanatory variables: Factors checked in the above model								
R square	19.7%	5.9%	20.4%	45.0%	53.7%	42.6%	29.6%		
Adjusted R-square	19.3%	5.6%	19.9%	44.7%	53.5%	42.2%	29.1%		

Abbreviations: BMI, body mass index; WHR, waist-hip ratio; SBP, systolic blood pressure; DBP, diastolic blood pressure; HbA1c, hemoglobin A1c; HDL-C, high-density lipoprotein cholesterol; SGPT, serum glutamic pyruvic transaminase; GGT, gamma-glutamyl transferase.

Table S4 The best model for HannumEAA, IEAA, PhenoEAA, DNAmPACKYRS, and DunedinPACE

Chronological age (years)	_	β	Standard error	95% Confid	ence interval	VIF	FDR ^a
Sex (female vs. male) -0.3588 0.1944 $[-0.7400, 0.022]$ 2.1470 $7.3E-02$ BMI (kg/m²) 0.0519 0.0232 $[0.0064, 0.0974]$ 1.6513 $3.2E-02$ WHR 3.7785 1.3272 $[1.1759, 6.3810]$ 1.8318 $6.9E-03**$ CD8† T cells (%) -8.3290 1.6304 $[-11.5261, -5.1318]$ 1.1267 $1.0E-06***$ CD4† T cells (%) -17.3285 1.3785 $[-2.00316, -14.6255]$ 1.2801 $5.8E-34****$ Natural killer cells (%) 18.1273 1.4798 $[15.2256, -21.0291]$ 1.2920 $2.9E-02*$ B lymphocytes (%) -6.4031 2.8184 $[-11.9299, -0.8764]$ 1.2820 $2.9E-02*$ DBP (mmHg) 0.0158 0.0069 $[0.0023, 0.0294]$ 1.3617 $2.8E-02*$ Fasting glucose (mg/dL) 0.0061 0.0035 $[-0.0008, 0.0130]$ 1.1223 $8.6E-02*$ Hemadorit (%) 0.0877 0.0313 $[0.0263, 0.1492]$ 4.3775 $7.7E-03***$ Creatinine (mg/dL) 0.0877	HannumEAA (in years)						
BMI (kg/m²) 0.0519 0.0232 [0.0064, 0.0974] 1.6513 3.2E-02 WHR 3.7785 1.3272 [1.1759, 6.3810] 1.8318 6.9E-03** CD8 T cells (%) -8.3290 1.6304 [-11.5261, -5.1318] 1.1267 1.0E-06*** CD4 T cells (%) -17.3285 1.3785 [-20.0316, -14.6255] 1.2801 5.8E-34**** Natural killer cells (%) 18.1273 1.4798 [15.2256, 21.0291] 1.2192 2.2E-32**** B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, -0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, -0.030] 1.1223 8.6E-02* Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05**** Hematocrit (%) 0.0877 0.0313 [0.023, -0.148] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1985 6.7E-08**	Chronological age (years)	-0.0371	0.0071	[-0.0511,	-0.0231]	1.4180	6.8E-07***
WHR 3.7785 1.3272 [1.1759, 6.3810] 1.8318 6.9E-03** CD8*T cells (%) -8.3290 1.6304 [-11.5261, -5.1318] 1.1267 1.0E-06*** CD4*T cells (%) -17.3285 1.3785 [-20.0316, -14.6255] 1.2801 5.8E-34*** Natural killer cells (%) 18.1273 1.4798 [15.2256, 21.0291] 1.2192 2.2E-32**** B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, -0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, -0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.9338 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, -0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, -0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, -0.8248] 1.1985 <	Sex (female vs. male)	-0.3588	0.1944	[-0.7400,	0.0225]	2.1470	7.3E-02
WHR 3.7785 1.3272 [1.1759, 6.3810] 1.8318 6.9E-03** CD8*T cells (%) -8.3290 1.6304 [-11.5261, -5.1318] 1.1267 1.0E-06*** CD4*T cells (%) -17.3285 1.3785 [-20.0316, -14.6255] 1.2801 5.8E-34*** Natural killer cells (%) 18.1273 1.4798 [15.2256, 21.0291] 1.2192 2.2E-32**** B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, -0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, -0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.9338 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, -0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, -0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, -0.8248] 1.1985 <	BMI (kg/m^2)	0.0519	0.0232	[0.0064,	0.0974]	1.6513	3.2E-02
CD4" T cells (%) -17.3285 1.3785 [-20.0316, -14.6255] 1.2801 5.8E-34*** Natural killer cells (%) 18.1273 1.4798 [15.2256, 21.0291] 1.2192 2.2E-32*** B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, 0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.035 [-0.0008, 0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, 0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.128 6.7E-08*** EVA (fin years) 5 0.2215 [0.8178, 1.6865] 1.3035 4.1E-13*** CD8* T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 <td< td=""><td>WHR</td><td>3.7785</td><td>1.3272</td><td>[1.1759,</td><td>6.3810]</td><td>1.8318</td><td>6.9E-03**</td></td<>	WHR	3.7785	1.3272	[1.1759,	6.3810]	1.8318	6.9E-03**
Natural killer cells (%) 18.1273 1.4798 [15.2256, 21.0291] 1.2192 2.2E-32*** B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, 0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, 0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, 0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CDB* T cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.091 4.6E-02* B lymphocytes (%) -7.4997 3.711	CD8 ⁺ T cells (%)	-8.3290	1.6304	[-11.5261,	-5.1318]	1.1267	1.0E-06***
B lymphocytes (%) -6.4031 2.8184 [-11.9299, -0.8764] 1.2820 2.9E-02* DBP (mmHg) 0.0158 0.0069 [0.0023, 0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, 0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, 0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8* T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4* T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8* T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** B lymphocytes (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	CD4 ⁺ T cells (%)	-17.3285	1.3785	[-20.0316,	-14.6255]	1.2801	5.8E-34***
DBP (mmHg) 0.0158 0.0069 [0.0023, 0.0294] 1.3617 2.8E-02* Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, 0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, 0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) *** Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8* T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4* T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297	Natural killer cells (%)	18.1273	1.4798	[15.2256,	21.0291]	1.2192	2.2E-32***
Fasting glucose (mg/dL) 0.0061 0.0035 [-0.0008, 0.0130] 1.1223 8.6E-02 Hemoglobin (g/dL) -0.3938 0.0959 [-0.5818, -0.2058] 4.9890 9.6E-05*** Hematocrit (%) 0.0877 0.0313 [0.0263, 0.1492] 4.3775 7.7E-03** Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8* T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4* T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3	B lymphocytes (%)	-6.4031	2.8184	[-11.9299,	-0.8764]	1.2820	2.9E-02*
Hemoglobin (g/dL)	DBP (mmHg)	0.0158	0.0069	[0.0023,	0.0294]	1.3617	2.8E-02*
Hematocrit (%)	Fasting glucose (mg/dL)	0.0061	0.0035	[-0.0008,	0.0130]	1.1223	8.6E-02
Creatinine (mg/dL) 0.4035 0.2148 [-0.0178, 0.8248] 1.1985 6.8E-02 Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8+ T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4+ T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) C	Hemoglobin (g/dL)	-0.3938	0.0959	[-0.5818,	-0.2058]	4.9890	9.6E-05***
Smoking (yes vs. no) 1.2521 0.2215 [0.8178, 1.6865] 1.1288 6.7E-08*** IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8* T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4* T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male)<	Hematocrit (%)	0.0877	0.0313	[0.0263,	0.1492]	4.3775	7.7E-03**
IEAA (in years) Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8+ T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4+ T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08**** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04****	Creatinine (mg/dL)	0.4035	0.2148	[-0.0178,	0.8248]	1.1985	6.8E-02
Sex (female vs. male) -1.2757 0.1700 [-1.6090, -0.9423] 1.3035 4.1E-13*** CD8+ T cells (%) 5.1438 1.7686 [1.6757, 8.6120] 1.0535 5.9E-03** CD4+ T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0	Smoking (yes vs. no)	1.2521	0.2215	[0.8178,	1.6865]	1.1288	6.7E-08***
$\begin{array}{c} \text{CD8$^{+}$T$ cells (\%)} & 5.1438 & 1.7686 & [1.6757, 8.6120] & 1.0535 & 5.9E-03**\\ \text{CD4$^{+}$T$ cells (\%)} & 4.3580 & 1.5883 & [1.2435, 7.4724] & 1.3515 & 8.9E-03**\\ \text{Natural killer cells (\%)} & -3.2626 & 1.5755 & [-6.3520, -0.1731] & 1.0991 & 4.6E-02*\\ \text{B lymphocytes (\%)} & -18.5297 & 3.1455 & [-24.6977, -12.3616] & 1.2686 & 1.8E-08***\\ \text{Monocytes (\%)} & -7.4997 & 3.7118 & [-14.7784, -0.2211] & 1.1804 & 5.1E-02\\ \text{DBP (mmHg)} & 0.0287 & 0.0071 & [0.0148, 0.0427] & 1.1514 & 1.2E-04***\\ \text{Smoking (yes vs. no)} & 0.3519 & 0.2463 & [-0.1310, 0.8349] & 1.1097 & 1.5E-01\\ \hline \textbf{PhenoEAA (in years)} \\ \text{Chronological age (years)} & -0.0227 & 0.0089 & [-0.0401, -0.0053] & 1.2739 & 1.5E-02*\\ \text{Sex (female vs. male)} & 0.9992 & 0.2600 & [0.4894, 1.5091] & 2.2250 & 2.7E-04***\\ \textbf{BMI (kg}/m^2) & 0.0932 & 0.0313 & [0.0319, 0.1546] & 1.7413 & 4.7E-03**\\ \textbf{WHR} & 3.7580 & 1.7560 & [0.3146, 7.2013] & 1.8565 & 3.9E-02*\\ \textbf{CD8$^{+}$T$ cells (\%)} & -20.2567 & 2.1276 & [-24.4288, -16.0846] & 1.1115 & 3.2E-20***\\ \textbf{CD4$^{+}$T$ cells (\%)} & -23.5376 & 1.8722 & [-27.2088, -19.8663] & 1.3685 & 5.8E-34***\\ \textbf{B lymphocytes (\%)} & -21.0387 & 3.7095 & [-28.3127, -13.7647] & 1.2872 & 6.3E-08***\\ \textbf{Monocytes (\%)} & 13.2991 & 4.3060 & [4.8553, 21.7429] & 1.1591 & 3.6E-03**\\ \end{array}$	IEAA (in years)						
CD4 ⁺ T cells (%) 4.3580 1.5883 [1.2435, 7.4724] 1.3515 8.9E-03** Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8 ⁺ T cells (%) -20.2567 2.1276	Sex (female vs. male)	-1.2757	0.1700	[-1.6090,	-0.9423]	1.3035	4.1E-13***
Natural killer cells (%) -3.2626 1.5755 [-6.3520, -0.1731] 1.0991 4.6E-02* B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8* T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4* T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	CD8 ⁺ T cells (%)	5.1438	1.7686	[1.6757,	8.6120]	1.0535	5.9E-03**
B lymphocytes (%) -18.5297 3.1455 [-24.6977, -12.3616] 1.2686 1.8E-08*** Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	CD4 ⁺ T cells (%)	4.3580	1.5883	[1.2435,	7.4724]	1.3515	8.9E-03**
Monocytes (%) -7.4997 3.7118 [-14.7784, -0.2211] 1.1804 5.1E-02 DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 <t< td=""><td>Natural killer cells (%)</td><td>-3.2626</td><td>1.5755</td><td>[-6.3520,</td><td>-0.1731]</td><td>1.0991</td><td>4.6E-02*</td></t<>	Natural killer cells (%)	-3.2626	1.5755	[-6.3520,	-0.1731]	1.0991	4.6E-02*
DBP (mmHg) 0.0287 0.0071 [0.0148, 0.0427] 1.1514 1.2E-04*** Smoking (yes vs. no) 0.3519 0.2463 [-0.1310, 0.8349] 1.1097 1.5E-01 PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	B lymphocytes (%)	-18.5297	3.1455	[-24.6977,	-12.3616]	1.2686	1.8E-08***
Smoking (yes vs. no) 0.3519 0.2463 $[-0.1310, 0.8349]$ 1.1097 $1.5E-01$ PhenoEAA (in years)Chronological age (years) -0.0227 0.0089 $[-0.0401, -0.0053]$ 1.2739 $1.5E-02*$ Sex (female vs. male) 0.9992 0.2600 $[0.4894, 1.5091]$ 2.2250 $2.7E-04***$ BMI (kg/ m^2) 0.0932 0.0313 $[0.0319, 0.1546]$ 1.7413 $4.7E-03**$ WHR 3.7580 1.7560 $[0.3146, 7.2013]$ 1.8565 $3.9E-02*$ CD8+ T cells (%) -20.2567 2.1276 $[-24.4288, -16.0846]$ 1.1115 $3.2E-20***$ CD4+ T cells (%) -23.5376 1.8722 $[-27.2088, -19.8663]$ 1.3685 $5.8E-34***$ B lymphocytes (%) -21.0387 3.7095 $[-28.3127, -13.7647]$ 1.2872 $6.3E-08***$ Monocytes (%) 13.2991 4.3060 $[4.8553, 21.7429]$ 1.1591 $3.6E-03**$	Monocytes (%)	-7.4997	3.7118	[-14.7784,	-0.2211]	1.1804	5.1E-02
PhenoEAA (in years) Chronological age (years) -0.0227 0.0089 [-0.0401, -0.0053] 1.2739 1.5E-02* Sex (female vs. male) 0.9992 0.2600 [0.4894, 1.5091] 2.2250 2.7E-04*** BMI (kg/m²) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	DBP (mmHg)	0.0287	0.0071	[0.0148,	0.0427]	1.1514	1.2E-04***
Chronological age (years) -0.0227 0.0089 $[-0.0401, -0.0053]$ 1.2739 $1.5E-02*$ Sex (female vs. male) 0.9992 0.2600 $[0.4894, 1.5091]$ 2.2250 $2.7E-04***$ BMI (kg/ m^2) 0.0932 0.0313 $[0.0319, 0.1546]$ 1.7413 $4.7E-03**$ WHR 3.7580 1.7560 $[0.3146, 7.2013]$ 1.8565 $3.9E-02*$ CD8+ T cells (%) -20.2567 2.1276 $[-24.4288, -16.0846]$ 1.1115 $3.2E-20***$ CD4+ T cells (%) -23.5376 1.8722 $[-27.2088, -19.8663]$ 1.3685 $5.8E-34***$ B lymphocytes (%) -21.0387 3.7095 $[-28.3127, -13.7647]$ 1.2872 $6.3E-08***$ Monocytes (%) 13.2991 4.3060 $[4.8553, 21.7429]$ 1.1591 $3.6E-03**$	Smoking (yes vs. no)	0.3519	0.2463	[-0.1310,	0.8349]	1.1097	1.5E-01
Sex (female vs. male) 0.9992 0.2600 $[0.4894, 1.5091]$ 2.2250 $2.7E-04***$ BMI (kg/ m^2) 0.0932 0.0313 $[0.0319, 0.1546]$ 1.7413 $4.7E-03**$ WHR 3.7580 1.7560 $[0.3146, 7.2013]$ 1.8565 $3.9E-02*$ CD8+ T cells (%) -20.2567 2.1276 $[-24.4288, -16.0846]$ 1.1115 $3.2E-20***$ CD4+ T cells (%) -23.5376 1.8722 $[-27.2088, -19.8663]$ 1.3685 $5.8E-34***$ B lymphocytes (%) -21.0387 3.7095 $[-28.3127, -13.7647]$ 1.2872 $6.3E-08***$ Monocytes (%) 13.2991 4.3060 $[4.8553, 21.7429]$ 1.1591 $3.6E-03**$	PhenoEAA (in years)						
BMI (kg/ m^2) 0.0932 0.0313 [0.0319, 0.1546] 1.7413 4.7E-03** WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8 ⁺ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4 ⁺ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	Chronological age (years)	-0.0227	0.0089	[-0.0401,	-0.0053]	1.2739	1.5E-02*
WHR 3.7580 1.7560 [0.3146, 7.2013] 1.8565 3.9E-02* CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	Sex (female vs. male)	0.9992	0.2600	[0.4894,	1.5091]	2.2250	2.7E-04***
CD8+ T cells (%) -20.2567 2.1276 [-24.4288, -16.0846] 1.1115 3.2E-20*** CD4+ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	BMI (kg/m^2)	0.0932	0.0313	[0.0319,	0.1546]	1.7413	4.7E-03**
CD4 ⁺ T cells (%) -23.5376 1.8722 [-27.2088, -19.8663] 1.3685 5.8E-34*** B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	WHR	3.7580	1.7560	[0.3146,	7.2013]	1.8565	3.9E-02*
B lymphocytes (%) -21.0387 3.7095 [-28.3127, -13.7647] 1.2872 6.3E-08*** Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	CD8 ⁺ T cells (%)	-20.2567	2.1276	[-24.4288,	-16.0846]	1.1115	3.2E-20***
Monocytes (%) 13.2991 4.3060 [4.8553, 21.7429] 1.1591 3.6E-03**	CD4 ⁺ T cells (%)	-23.5376	1.8722	[-27.2088,	-19.8663]	1.3685	5.8E-34***
• • • •	B lymphocytes (%)	-21.0387	3.7095	[-28.3127,	-13.7647]	1.2872	6.3E-08***
DBP (mmHg) 0.0252 0.0091 [0.0074, 0.0430] 1.3613 8.2E-03**	Monocytes (%)	13.2991	4.3060	[4.8553,	21.7429]	1.1591	3.6E-03**
	DBP (mmHg)	0.0252	0.0091	[0.0074,	0.0430]	1.3613	8.2E-03**

	β	Standard error	95% Confid	ence interval	VIF	FDR ^a
HbA1c (%)	0.3938	0.1282	[0.1423,	0.6453]	1.1584	3.7E-03**
Hemoglobin (g/dL)	-0.3782	0.0759	[-0.5271,	-0.2293]	1.8114	1.9E-06***
GGT (U/L)	0.0082	0.0027	[0.0030,	0.0135]	1.1247	3.7E-03**
Uric acid (mg/dL)	0.1855	0.0787	[0.0312,	0.3398]	1.7082	2.4E-02*
Smoking (yes vs. no)	1.8586	0.2908	[1.2885,	2.4288]	1.1252	8.5E-10***
DNAmPACKYRS (DNAm	-based smo	king pack-years	s)			
Chronological age (years)	0.2636	0.0116	[0.2409,	0.2864]	1.4153	2.2E-102***
Sex (female vs. male)	-3.0770	0.2935	[-3.6525,	-2.5015]	1.8486	3.4E-24***
BMI (kg/ m^2)	-0.0598	0.0368	[-0.1321,	0.0124]	1.5964	1.1E-01
WHR (%)	3.8990	2.1914	[-0.3983,	8.1962]	1.8805	8.2E-02
CD8 ⁺ T cells (%)	-7.4933	2.6084	[-12.6082,	-2.3784]	1.0923	6.5E-03**
Natural killer cells (%)	-7.3608	2.4147	[-12.0959,	-2.6257]	1.2345	3.9E-03**
Monocytes (%)	16.7706	5.1642	[6.6439,	26.8973]	1.0718	2.2E-03**
HbA1c (%)	0.5477	0.1647	[0.2248,	0.8706]	1.1645	1.8E-03**
HDL-C (mg/dL)	-0.0165	0.0093	[-0.0349,	0.0018]	1.4102	8.3E-02
Hematocrit (%)	0.0592	0.0305	[-0.0007,	0.1191]	1.5830	6.0E-02
GGT (U/L)	0.0099	0.0035	[0.0030,	0.0168]	1.1038	7.7E-03**
Smoking (yes vs. no)	13.6793	0.3833	[12.9276,	14.4310]	1.0942	3.1E-222***
DunedinPACE						
Chronological age (years)	0.0024	0.0002	[0.0020,	0.0028]	1.5815	7.0E-29***
BMI (kg/m^2)	0.0048	0.0006	[0.0036,	0.0061]	1.6878	2.7E-13***
WHR (%)	0.0518	0.0365	[-0.0197,	0.1234]	1.8730	1.6E-01
CD8 ⁺ T cells (%)	-0.1797	0.0438	[-0.2655,	-0.0938]	1.1023	9.6E-05***
CD4 ⁺ T cells (%)	-0.3499	0.0363	[-0.4211,	-0.2788]	1.2034	1.1E-20***
Natural killer cells (%)	-0.1274	0.0404	[-0.2066,	-0.0482]	1.2335	3.0E-03**
Monocytes (%)	0.4482	0.0898	[0.2721,	0.6243]	1.1785	1.8E-06***
SBP (mmHg)	0.0003	0.0001	[0.0000,	0.0005]	1.4103	3.2E-02*
HbA1c (%)	0.0176	0.0044	[0.0090,	0.0262]	3.1475	1.3E-04***
Fasting glucose (mg/dL)	-0.0003	0.0002	[-0.0006,	[0.0000]	3.1050	8.6E-02
Triglyceride (mg/dL)	0.0000	0.0000	[0.0000,	0.0001]	1.3567	8.2E-02
HDL-C (mg/dL)	-0.0009	0.0002	[-0.0012,	-0.0006]	1.5138	1.8E-07***
Hemoglobin (g/dL)	-0.0135	0.0025	[-0.0185,	-0.0086]	4.6319	2.2E-07***
Hematocrit (%)	0.0014	0.0009	[-0.0003,	0.0031]	4.4342	1.1E-01
GGT (U/L)	0.0001	0.0001	[0.0000,	0.0003]	1.1353	1.1E-02*
Creatinine (mg/dL)	0.0110	0.0056	[0.0000,	0.0220]	1.1078	5.9E-02
Smoking (yes vs. no)	0.0774	0.0060	[0.0655,	0.0892]	1.1410	5.7E-35***

Abbreviations: VIF, variance inflation factor; FDR, false discovery rate; BMI, body mass index; WHR, waist-hip ratio; HbA1c, hemoglobin A1c; HDL-C, high-density lipoprotein cholesterol; GGT, gamma-glutamyl transferase.

FDR $^{\rm a}$: Statistical significance is marked with *, **, and ***, representing an FDR less than 0.05, 0.01, and 0.001, respectively.

Figures



Figure S1 Partial correlation analysis between DNAm-based markers and lifestyle factors

This figure illustrates the partial correlation coefficients (r with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between lifestyle factors and epigenetic age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: Edu, education attainment; LA, living alone; BMI, body mass index; WHR, waist-hip ratio.

Sex: 1 for males; 2 for females.

Edu (Educational attainment): Educational attainment was recorded as an integer ranging from 1 to 7, representing different levels of education. "1" represents no formal education and illiteracy, "2" represents self-taught individuals who are literate, "3" represents the completion of elementary school, "4" represents the completion of junior high school, "5" represents the completion of senior high school or vocational high school, "6" represents the completion of college or technical school, and "7" represents the completion of graduate school or higher education.

LA (Living alone): 1 for not living alone; 2 for living alone.

Drink: consuming more than 150 mL of alcoholic beverages per week for at least six months when participating in the TWB, coded as $1 = N_0$, $2 = Y_0$ es.

Smoke: smoking cigarettes for at least six months when participating in the TWB, coded as 1 = No, 2 = Yes. Sport: exercising for at least 30 minutes thrice a week. Exercise included leisure-time activities such as swimming, jogging, cycling, mountain climbing, dancing, weight training, etc., coded as follows: 1 = No, 2 = Yes.

SHS (Secondhand smoke): being exposed to an environment with secondhand smoke (someone smoking nearby) for at least five minutes in the past six months, coded as 1 = No, 2 = Yes.

Nut: betel nut chewing in the past six months, coded as 1 = No, 2 = Yes.

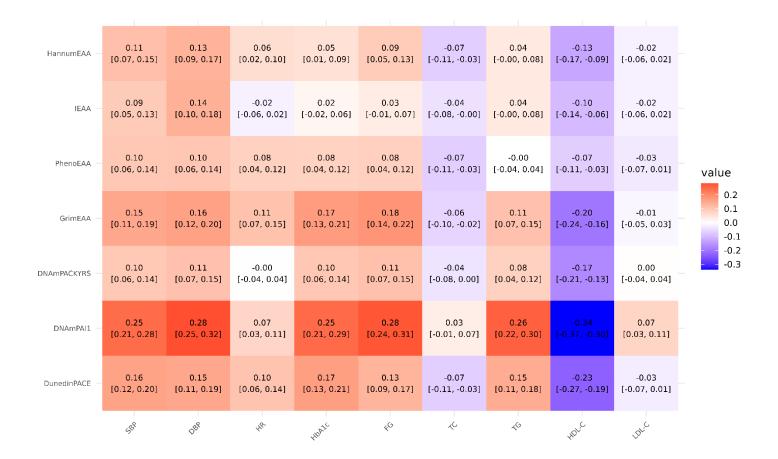


Figure S2 Partial correlation analysis between DNAm-based markers, cardiovascular health metrics, and blood biochemical indicators

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between cardiovascular health metrics, blood biochemical variables, and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: SBP, systolic blood pressure; DBP, diastolic blood pressure; HR, heart rate; HbA1c, hemoglobin A1c; FG, fasting glucose; TC, total cholesterol; TG, triglyceride; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol.

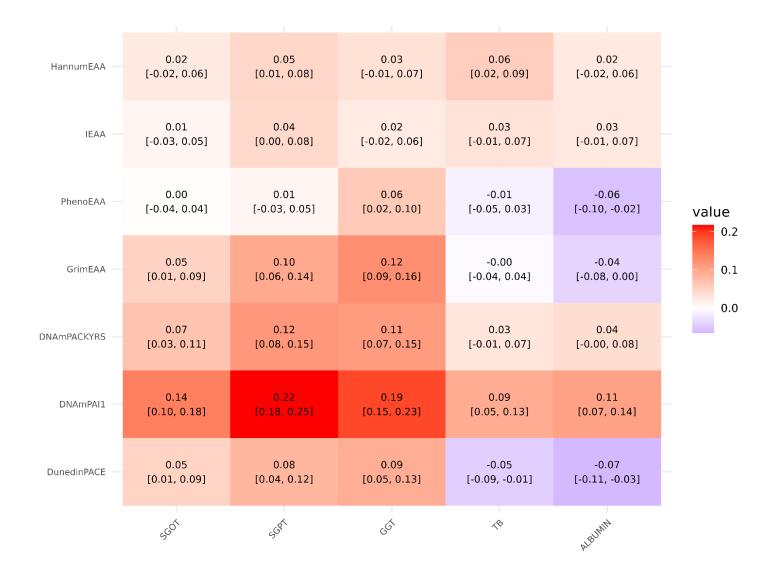


Figure S3 Partial correlation analysis between DNAm-based markers and blood biochemical indicators

Partial correlation analysis was conducted between DNAm-based markers and blood biochemical variables. This figure illustrates the partial correlation coefficients (r with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between blood biochemical variables and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: SGOT, serum glutamic oxaloacetic transaminase; SGPT, serum glutamic pyruvic transaminase; GGT, gamma-glutamyl transferase; TB, total bilirubin.

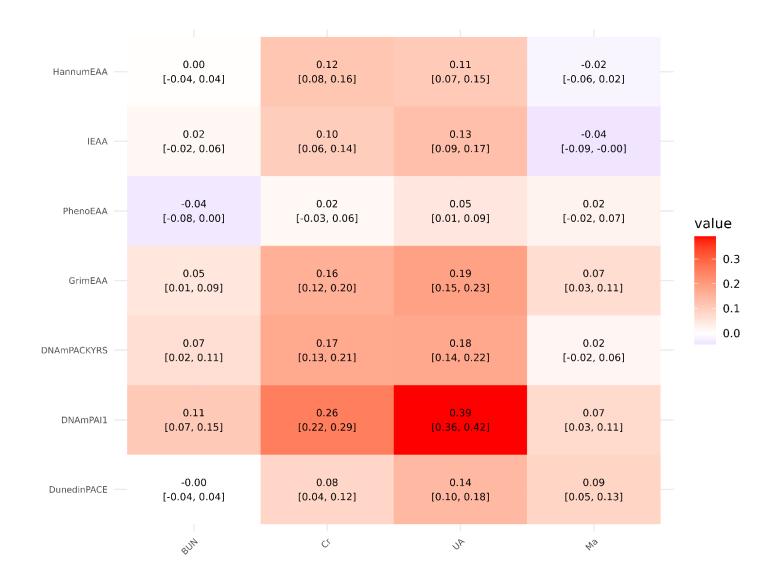


Figure S4 Partial correlation analysis between DNAm-based markers and blood biochemical indicators

Partial correlation analysis was conducted between DNAm-based markers and blood biochemical variables. This figure illustrates the partial correlation coefficients (r with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between blood biochemical indicators and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: BUN, blood urea nitrogen; Cr, creatinine; UA, uric acid; Ma, Microalbumin.

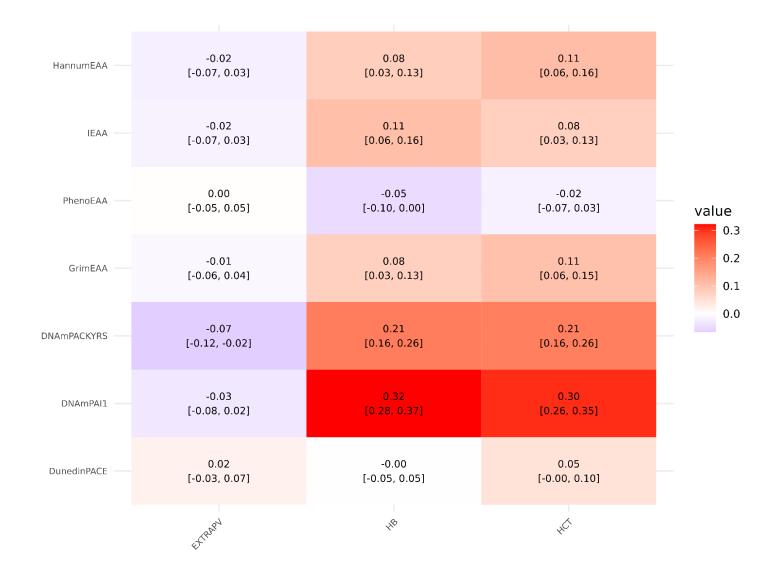


Figure S5 Partial correlation analysis between DNAm-based markers, lung function measures, and blood biochemical indicators

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between lung function measures, blood biochemical indicators, and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: EXTRAPV, (Extrapolated Volume/ forced vital capacity) *100; HB, hemoglobin; HCT, hematocrit.



Figure S6 Partial correlation analysis between DNAm-based markers and lung function measures

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between lung function measures and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: VC, vital capacity; VC/HT, vital capacity/height; FVC, forced vital capacity; FEV1, forced expiratory volume in 1.0 s; FEV1/SVC, (forced expiratory volume in 1.0 s / vital capacity)*100; FEV1/VCPR, forced expiratory volume in 1.0 s /predicted vital capacity; MMEF, maximum mid-expiratory flow; PEF, peak expiratory flow; FEF25, forced expiratory flow at 25%; FEF50, forced expiratory flow at 50%; FEF75, forced expiratory flow at 75%/height.

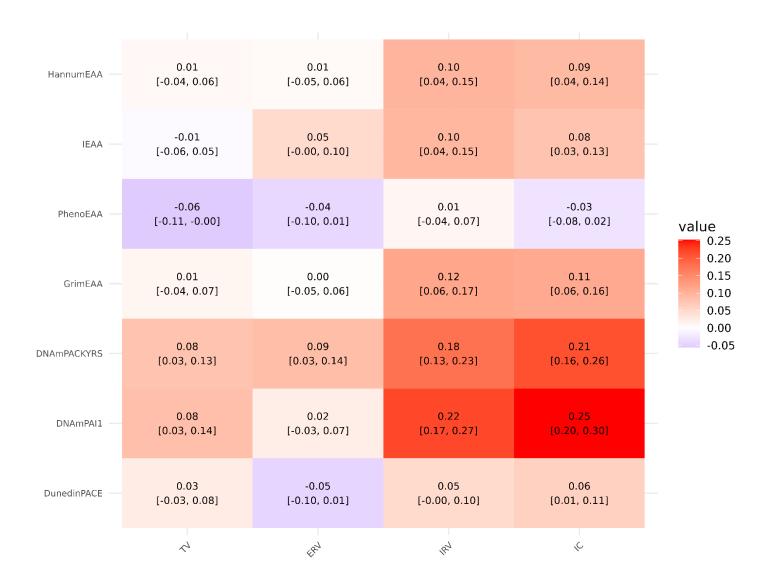


Figure S7 Partial correlation analysis between DNAm-based markers and lung function measures

This figure illustrates the partial correlation coefficients (r with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between lung function measures and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: TV, tidal volume; ERV, expiratory reserve volume; IRV, inspiratory reserve volume; IC, inspiratory capacity.



Figure S8 Partial correlation analysis between DNAm-based markers and diet preference

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between diet preference and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations. D1-D17 are 17 questions in the TWB questionnaires, listed in the supplementary Table S1.

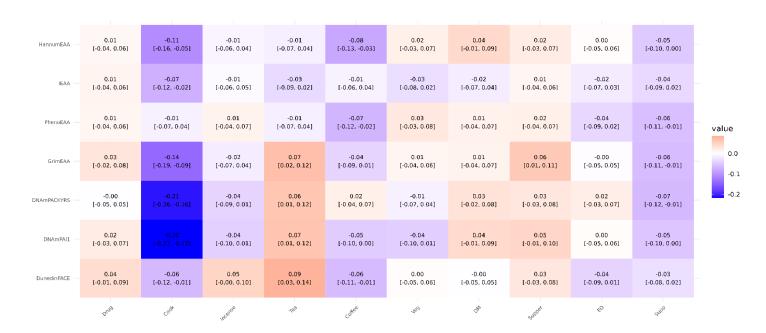


Figure S9 Partial correlation analysis between DNAm-based markers and lifestyle-related factors

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between lifestyle-related factors and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Drug: taking cough syrup, sedatives, or pain relievers at least once a week within the six months before joining the TWB, coded as $1 = N_0$, $2 = Y_{es}$.

Cook: cooking meals by yourselves within six months before participating in the TWB, categorized as $1 = N_0$, $2 = Y_{es}$.

Incense: being exposed to incense burning (e.g., during worship or the use of incense powder or rings), mosquito coils (traditional, liquid electric, or electric mosquito repellent), or fragrances (such as essential oils, aromatherapy, air fresheners, sprays, or scented candles) for at least five minutes within the past year before joining the TWB, categorized as 1 = No, 2 = Yes.

Tea: consuming tea (containing tea leaves, excluding herbal teas) at least once daily within six months before joining the TWB, categorized as $1 = N_0$, $2 = Y_0$ es.

Coffee: coffee drinking thrice a week, categorized as $1 = N_0$, $2 = Y_{es}$.

Veg: vegetarian diet for at least six months before joining the TWB, categorized as 1 = No, 2 = Yes.

DM: Daily meals, number of main meals per day, an integer ranging from 1 to 6.

Supper: eating supper within an hour before bedtime (including milk and wine), categorized as $1 = N_0$, $2 = Y_0$ es.

EO (Eating out): eating out at least once in the past month when joining the TWB, categorized as 1 = No, 2 = Yes.

Supp: regularly taking vitamins, minerals, or supplements in the past month before joining the TWB, categorized as $1 = N_0$, $2 = Y_0$ es.

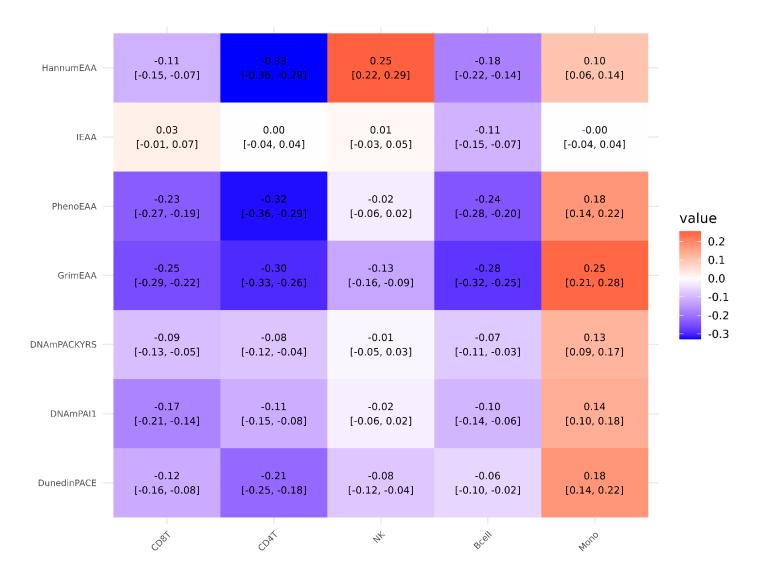


Figure S10 Partial correlation analysis between DNAm-based markers and cell-type proportions

This figure illustrates the partial correlation coefficients (*r* with its 95% confidence interval) adjusted for chronological age, smoking, and drinking status, indicating the relationship between cell-type proportions and DNA methylation age acceleration. Red cells denote positive correlations, while blue cells represent negative correlations.

Abbreviations: CD8T, CD8⁺ T cells; CD4T, CD4⁺ T cells; NK, natural killer cells; Bcell: B lymphocytes; Mono, monocytes.

References:

Lo, Y. H., & Lin, W. Y. (2022). Cardiovascular health and four epigenetic clocks. *Clin Epigenetics*, 14(1), 73. doi:10.1186/s13148-022-01295-7