

## **Supplement 1. Supplement methods**

### *Data source and study cohort*

In Korea, there is a single health insurance system administered by the NHIS, and all data related to insurance eligibility and medical claims for all Koreans are stored in the NHIS database. The NHIS provides health checkups at least once every two years for the purpose of early detection and prevention of diseases. NHIS-HEALS was established based on 514,866 Koreans (aged 40 to 79 in 2002), who randomly selected 10% of the national health screening participants from 2002 to 2003. It includes data on regular blood tests, physical examinations, such as BMI and blood pressure, and questionnaires regarding medical history and lifestyle.

### *Study population*

To define the prediabetes cohort, those with a history of diabetes (ICD-10: E10-E14 and  $\geq 1$  claim for antihyperglycemic medication) prior to the index date were excluded. Prediabetes is defined by the presence of IFG and/or IGT and/or A1C 5.7–6.4%, which is defined as fasting serum glucose 100 to 125mg/dL at the first health screening as IFG since the data of check-up did not contain 2h-plasma glucose value during 75g oral glucose tolerance test and A1c.(1) Regarding the prehypertension cohort, those with a history of hypertension (ICD-10: I10-I15 and  $\geq 1$  claim for an-tihypertensive medication) prior to the index date were excluded. Prehypertension was defined as systolic pressure of 120 to 139mmHg or diastolic pressure of 80 to 89 mmHg at the first health screening.(2, 3) The antihyperglycemic medications that were considered in this study were metformin, sulfonylureas, thiazolidinediones, alpha glucosidase inhibitors, glucagon-like peptide-1 receptor agonists, Dipeptidyl peptidase 4 inhibitors, sodium-glucose cotransporter 2 inhibitors, and insulin. The

antihypertensive medications that were considered in this study were diuretics, angiotensin-converting enzyme inhibitors, beta adrenergic receptor blockers, alpha adrenergic receptor blockers, angiotensin II receptor blockers, calcium channel blockers, vasodilators and centrally acting sympatholytics.

### *Assessment of lifestyle changes*

Change of BMI was assessed change in BMI categories (normal:  $< 23.0 \text{ kg/m}^2$ , overweight:  $23.0 \leq \sim < 25.0 \text{ kg/m}^2$ , and obesity:  $\geq 25.0 \text{ kg/m}^2$ ). Data on current smoking, alcohol intake, and physical activity were collected based on self-reported questionnaires. In the second health screening, compared to the previous period, whether each of the four factors worsened or improved was evaluated. For example, if an individual's BMI category changed from normal at the first screening to overweight at the second screening, he or she has been evaluated as '1 factor worsened', and if the participant continued smoking from the first to the second health screening, he or she has been evaluated as 'no change in lifestyle.' On the other hand, if the participant answered 'never' for exercise at the first health screening and answered 'exercise 2-3 times per month' at the second screening, he or she has been evaluated as '1 factor improved' (Supplementary table 1).

### *Statistical analysis*

We estimated adjusted HR under controlling by age, income level (quantiles), current smoking status, alcohol consumption, physical activity, CCI score, BMI, systolic blood pressure, total cholesterol, fasting serum glucose level at the first health screening, and usage of statin medication prior to the index date. Charlson comorbidity Index score (CCI score) is a

tool used to predict the risk of mortality for patients based on their comorbid conditions and estimate their 10-year survival. We also assessed the impact of lifestyle changes on health risks by adjusting aspirin, antihyperglycemic drugs, or antihypertensive drugs.

Since weight change can be considered to be a result of changes in lifestyle, lifestyle effects were evaluated by considering only their current smoking status, drinking habits, and frequency of physical activity, excluding BMI level, among lifestyle factors. As a sensitivity analysis, to avoid potential reversal causality, we excluded subjects with incident outcomes that occurred within the first 2 years.

## References

1. Committee ADAPP, Committee: ADAPP. 2. Classification and diagnosis of diabetes: Standards of Medical Care in Diabetes—2022. *Diabetes Care* 2022;45(Supplement\_1):S17-S38.
2. O'Neal WT, Soliman EZ, Qureshi W, Alonso A, Heckbert SR, Herrington D. Sustained pre-hypertensive blood pressure and incident atrial fibrillation: the Multi-Ethnic Study of Atherosclerosis. *J Am Soc Hypertens* 2015;9(3):191-196.
3. Mancia G, Fagard R, Narkiewicz K, Redán J, Zanchetti A, Böhm M, et al. 2013 Practice guidelines for the management of arterial hypertension of the European Society of Hypertension (ESH) and the European Society of Cardiology (ESC): ESH/ESC Task Force for the Management of Arterial Hypertension. *J Hypertens* 2013;31(10):1925-1938.

**Supplement 2. Distribution pattern according to the changes in lifestyle pattern**

<b>Prehypertension Group</b>												
	<b>All</b>				<b>Male</b>				<b>Female</b>			
<b>Lifestyle change</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>
4 factors worsened	42	0.05	22,778	26.6	41	0.08	15,245	28.37	1	0	7,533	23.61
3 factors worsened	575	0.67			501	0.93			74	0.23		
2 factors worsened	4,451	5.2			3,354	6.24			1,097	3.44		
1 factor worsened	17,710	20.68			11,349	21.12			6,361	19.94		
No change of lifestyle	33,602	39.24	33,602	39.24	19,440	36.17	19,440	36.17	14,162	44.4	14,162	44.4
1 factor improved	21,738	25.38	29,262	34.17	13,413	24.96	19,059	35.46	8,325	26.1	10,203	31.99
2 factors improved	6,498	7.59			4,761	8.86			1,737	5.45		
3 factors improved	975	1.14			834	1.55			141	0.44		
4 factors improved	51	0.06			51	0.09						
<b>Prediabetes Group</b>												
	<b>All</b>				<b>Male</b>				<b>Female</b>			
<b>Lifestyle change</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>	<b>Patient (n)</b>	<b>Percentage (%)</b>
4 factors worsened	27	0.05	13,667	26.13	26	0.08	9,479	27.8	1	0.01	4,188	23.01
3 factors worsened	341	0.65			301	0.88			40	0.22		
2 factors worsened	2,678	5.12			2,117	6.21			561	3.08		

1 factor worsened	10,621	20.31			7,035	20.63			3,586	19.7		
No change of lifestyle	20,417	39.04	20,417	39.04	12,292	36.05	12,292	36.05	8,125	44.64	8,125	44.64
1 factor improved	13,492	25.8	18,218	34.83	8,714	25.56	12,328	36.16	4,778	26.25	5,890	32.36
2 factors improved	4,105	7.85			3,065	8.99			1,040	5.71		
3 factors improved	582	1.11			511	1.5			71	0.39		
4 factors improved	39	0.07			38	0.11			1	0.01		

**Supplement 3. Univariate analysis of total cardiovascular disease events and mortality associated with lifestyle changes between two biennial health screening periods in the prehypertension group**

Outcome	Parameter	Change	Male		Female		
			Hazard ratio [95% CI]	<i>P</i> -value	Hazard ratio [95% CI]	<i>P</i> -value	
MACE	Unhealthy lifestyle	Worsened	1.06 [0.98–1.15]	0.162	0.987 [0.89–1.10]	0.810	
		Improved	0.986 [0.91–1.07]	0.724	0.913 [0.83–1.01]	0.076	
	Unhealthy lifestyle (exclude BMI)	Worsened	1.064 [0.98–1.16]	0.136	0.886 [0.79–0.99]	0.035	
		Improved	0.994 [0.92–1.08]	0.889	0.836 [0.76–0.93]	0.001	
	BMI	Worsened	1.047 [0.95–1.16]	0.366	1.073 [0.94–1.23]	0.296	
		Improved	1.043 [0.95–1.15]	0.400	1.048 [0.93–1.19]	0.454	
	Currently smoking	Worsened	1.039 [0.91–1.19]	0.571	2.404 [1.65–3.52]	< 0.001	
		Improved	0.982 [0.88–1.09]	0.741	1.926 [1.32–2.82]	0.001	
	Frequency of drinking	Worsened	1.047 [0.97–1.13]	0.263	0.762 [0.65–0.89]	0.001	
		Improved	1.096 [1.01–1.18]	0.020	0.96 [0.84–1.10]	0.561	
	Frequency of physical activity	Worsened	0.994 [0.91–1.08]	0.889	0.956 [0.85–1.07]	0.439	
		Improved	0.884 [0.82–0.96]	0.002	0.802 [0.72–0.89]	< 0.001	
	Non-fatal MI	Unhealthy lifestyle	Worsened	1.004 [0.87–1.17]	0.956	0.908 [0.71–1.16]	0.441
			Improved	0.951 [0.83–1.10]	0.485	0.877 [0.70–1.10]	0.256
Unhealthy lifestyle (exclude BMI)		Worsened	1.053 [0.91–1.22]	0.495	0.88 [0.68–1.13]	0.324	
		Improved	0.965 [0.84–1.11]	0.626	0.834 [0.66–1.05]	0.124	
BMI		Worsened	1.027 [0.86–1.23]	0.776	0.686 [0.48–0.98]	0.036	

	Currently smoking	Improved	1.113 [0.94–1.32]	0.228	0.989 [0.75–1.30]	0.936
		Worsened	1.073 [0.84–1.36]	0.564	2.206 [0.91–5.33]	0.079
	Frequency of drinking	Improved	1.172 [0.97–1.41]	0.092	3.264 [1.69–6.32]	< 0.001
		Worsened	0.995 [0.86–1.15]	0.947	0.915 [0.66–1.27]	0.599
	Frequency of physical activity	Improved	0.979 [0.85–1.13]	0.773	0.867 [0.63–1.20]	0.393
		Worsened	0.992 [0.86–1.15]	0.913	0.908 [0.70–1.18]	0.463
Non-fatal stroke	Unhealthy lifestyle	Worsened	1.113 [1.01–1.23]	0.030	0.99 [0.88–1.12]	0.876
		Improved	1.024 [0.93–1.12]	0.625	0.904 [0.81–1.01]	0.077
	Unhealthy lifestyle (exclude BMI)	Worsened	1.098 [1.00–1.21]	0.058	0.891 [0.79–1.01]	0.067
		Improved	1.024 [0.93–1.12]	0.617	0.813 [0.73–0.91]	0.000
	BMI	Worsened	1.047 [0.93–1.18]	0.442	1.097 [0.95–1.27]	0.210
		Improved	1.017 [0.91–1.14]	0.779	1.051 [0.92–1.20]	0.478
	Currently smoking	Worsened	1.026 [0.88–1.20]	0.754	2.496 [1.65–3.77]	< 0.001
		Improved	0.92 [0.81–1.05]	0.218	1.461 [0.91–2.36]	0.120
	Frequency of drinking	Worsened	1.086 [0.99–1.19]	0.086	0.742 [0.62–0.89]	0.001
		Improved	1.149 [1.05–1.26]	0.003	0.967 [0.83–1.13]	0.673
	Frequency of physical activity	Worsened	0.999 [0.91–1.10]	0.977	0.962 [0.85–1.09]	0.541
		Improved	0.866 [0.79–0.95]	0.002	0.777 [0.69–0.87]	< 0.001
CVD death	Unhealthy lifestyle	Worsened	1.084 [0.79–1.48]	0.611	0.85 [0.54–1.34]	0.484
		Improved	0.891 [0.65–1.22]	0.466	0.606 [0.38–0.96]	0.033
	Unhealthy lifestyle (exclude BMI)	Worsened	1.039 [0.76–1.42]	0.810	0.69 [0.42–1.15]	0.153

	BMI	Improved	0.859 [0.63–1.17]	0.332	0.69 [0.44–1.09]	0.108	
		Worsened	0.856 [0.56–1.31]	0.473	1.444 [0.87–2.39]	0.154	
	Currently smoking	Improved	1.099 [0.76–1.60]	0.619	0.868 [0.49–1.55]	0.632	
		Worsened	1.657 [1.08–2.53]	0.020	5.004 [1.59–15.8]	0.006	
	Frequency of drinking	Improved	0.758 [0.47–1.23]	0.261	8.087 [3.55–18.4]	< 0.001	
		Worsened	1.084 [0.80–1.47]	0.609	0.417 [0.17–1.02]	0.056	
	Frequency of physical activity	Improved	0.991 [0.73–1.35]	0.955	0.77 [0.40–1.48]	0.432	
		Worsened	0.79 [0.57–1.10]	0.158	0.713 [0.43–1.19]	0.194	
	<b>All-cause death</b>	Unhealthy lifestyle	Improved	0.717 [0.52–0.98]	0.038	0.439 [0.26–0.75]	0.003
			Worsened	1.013 [0.92–1.11]	0.793	0.896 [0.76–1.06]	0.211
Unhealthy lifestyle (exclude BMI)		Improved	0.997 [0.91–1.09]	0.947	0.816 [0.70–0.96]	0.013	
		Worsened	1.092 [1.00–1.20]	0.063	0.816 [0.68–0.98]	0.026	
BMI		Improved	1.026 [0.94–1.12]	0.567	0.731 [0.62–0.86]	< 0.001	
		Worsened	0.831 [0.74–0.94]	0.003	1.08 [0.88–1.33]	0.472	
Currently smoking		Improved	0.973 [0.87–1.09]	0.636	1.144 [0.95–1.38]	0.162	
		Worsened	1.109 [0.96–1.28]	0.167	2.874 [1.66–4.97]	< 0.001	
Frequency of drinking		Improved	0.945 [0.83–1.07]	0.377	3.979 [2.61–6.08]	< 0.001	
		Worsened	1.068 [0.98–1.17]	0.155	0.935 [0.74–1.18]	0.574	
Frequency of physical activity		Improved	1.115 [1.02–1.22]	0.015	0.898 [0.71–1.13]	0.354	
		Worsened	0.844 [0.77–0.93]	< 0.001	0.869 [0.73–1.04]	0.132	
		Improved	0.791 [0.72–0.86]	< 0.001	0.699 [0.59–0.83]	< 0.001	

MACE, major adverse cardiovascular events; BMI, body mass index; CVD, cardiovascular disease.



**Supplement 4. Univariate analysis of total cardiovascular disease events and mortality associated with lifestyle change between two biennial health screening periods in the prediabetes group**

Outcome	Parameter	Change	Male		Female		
			hazard ratio [95% CI]	<i>P</i> -value	hazard ratio [95% CI]	<i>P</i> -value	
MACE	Unhealthy lifestyle	Worsened	1.134 [1.04–1.24]	0.005	0.934 [0.83–1.05]	0.268	
		Improved	1.06 [0.98–1.15]	0.173	0.904 [0.81–1.01]	0.071	
	Unhealthy lifestyle (exclude BMI)	Worsened	1.165 [1.07–1.27]	0.001	0.816 [0.72–0.93]	0.002	
		Improved	1.092 [1.01–1.19]	0.037	0.862 [0.77–0.96]	0.009	
	BMI	Worsened	1.095 [0.98–1.22]	0.099	1.133 [0.98–1.31]	0.090	
		Improved	1.006 [0.91–1.12]	0.906	0.998 [0.87–1.14]	0.976	
	Currently smoking	Worsened	1.086 [0.95–1.25]	0.244	1.006 [0.52–1.94]	0.987	
		Improved	0.985 [0.87–1.11]	0.810	2.271 [1.59–3.24]	< 0.001	
	Frequency of drinking	Worsened	1.065 [0.98–1.16]	0.150	0.82 [0.70–0.97]	0.019	
		Improved	1.109 [1.02–1.20]	0.013	0.824 [0.70–0.97]	0.019	
	Frequency of physical activity	Worsened	1.047 [0.96–1.14]	0.303	0.803 [0.70–0.92]	0.002	
		Improved	0.942 [0.87–1.02]	0.157	0.845 [0.75–0.95]	0.004	
	Non-fatal MI	Unhealthy lifestyle	Worsened	1.27 [1.07–1.50]	0.006	0.957 [0.74–1.24]	0.734
			Improved	1.174 [1.00–1.38]	0.051	0.815 [0.64–1.04]	0.094
Unhealthy lifestyle (exclude BMI)		Worsened	1.194 [1.01–1.41]	0.040	0.759 [0.57–1.00]	0.052	
		Improved	1.184 [1.01–1.39]	0.036	0.757 [0.59–0.97]	0.026	
BMI		Worsened	1.088 [0.89–1.34]	0.418	1.223 [0.91–1.65]	0.191	

	Currently smoking	Improved	0.837 [0.68–1.04]	0.106	0.899 [0.66–1.22]	0.493
		Worsened	1.148 [0.88–1.49]	0.302	1.576 [0.51–4.91]	0.433
	Frequency of drinking	Improved	1.169 [0.94–1.45]	0.154	3.065 [1.58–5.94]	0.001
		Worsened	0.948 [0.80–1.12]	0.532	0.852 [0.60–1.21]	0.374
	Frequency of physical activity	Improved	1.036 [0.88–1.21]	0.662	0.736 [0.51–1.07]	0.104
		Worsened	1.079 [0.92–1.27]	0.366	0.73 [0.54–0.98]	0.037
Non-fatal stroke	Unhealthy lifestyle	Improved	0.932 [0.80–1.09]	0.388	0.743 [0.58–0.95]	0.020
		Worsened	1.11 [1.00–1.23]	0.047	0.956 [0.84–1.09]	0.511
	Unhealthy lifestyle (exclude BMI)	Improved	1.026 [0.93–1.13]	0.613	0.921 [0.82–1.04]	0.189
		Worsened	1.169 [1.06–1.29]	0.003	0.828 [0.72–0.96]	0.009
	BMI	Improved	1.06 [0.96–1.17]	0.241	0.862 [0.76–0.98]	0.019
		Worsened	1.076 [0.95–1.22]	0.254	1.124 [0.96–1.32]	0.156
	Currently smoking	Improved	1.034 [0.92–1.17]	0.583	1.023 [0.88–1.19]	0.759
		Worsened	1.123 [0.96–1.32]	0.155	0.969 [0.46–2.04]	0.934
	Frequency of drinking	Improved	0.938 [0.81–1.08]	0.383	2.343 [1.59–3.45]	< 0.001
		Worsened	1.128 [1.02–1.25]	0.017	0.801 [0.67–0.96]	0.019
	Frequency of physical activity	Improved	1.15 [1.05–1.27]	0.004	0.818 [0.68–0.98]	0.029
		Worsened	1.036 [0.94–1.15]	0.494	0.827 [0.71–0.96]	0.012
CVD death	Unhealthy lifestyle	Improved	0.929 [0.84–1.02]	0.135	0.855 [0.75–0.97]	0.015
		Worsened	1.039 [0.77–1.39]	0.798	0.735 [0.50–1.20]	0.191
	Unhealthy lifestyle (exclude BMI)	Worsened	1.027 [0.77–1.37]	0.859	0.843 [0.50–1.30]	0.466

	BMI	Improved	0.921 [0.70–1.22]	0.563	0.775 [0.50–1.20]	0.234	
		Worsened	1.185 [0.83–1.68]	0.343	1.171 [0.70–2.00]	0.556	
	Currently smoking	Improved	1.256 [0.91–1.73]	0.166	1.15 [0.70–1.80]	0.559	
		Worsened	0.701 [0.40–1.22]	0.212	NA	NA	
	Frequency of drinking	Improved	0.812 [0.53–1.24]	0.339	4.583 [1.90–11.20]	0.001	
		Worsened	0.883 [0.66–1.18]	0.397	0.532 [0.30–1.10]	0.084	
	Frequency of physical activity	Improved	0.834 [0.63–1.11]	0.213	0.568 [0.30–1.10]	0.102	
		Worsened	0.999 [0.75–1.34]	0.997	0.911 [0.60–1.50]	0.699	
	<b>All-cause death</b>	Unhealthy lifestyle	Improved	0.876 [0.66–1.16]	0.354	0.749 [0.50–1.20]	0.191
			Worsened	1.067 [0.97–1.18]	0.205	0.969 [0.82–1.15]	0.719
Unhealthy lifestyle (exclude BMI)		Improved	1.041 [0.95–1.14]	0.405	0.796 [0.68–0.94]	0.007	
		Worsened	1.071 [0.97–1.18]	0.173	0.901 [0.75–1.08]	0.255	
BMI		Improved	1.005 [0.92–1.10]	0.925	0.715 [0.60–0.85]	< 0.001	
		Worsened	0.949 [0.84–1.08]	0.423	0.839 [0.66–1.06]	0.148	
Currently smoking		Improved	1.036 [0.92–1.16]	0.555	1.035 [0.86–1.25]	0.723	
		Worsened	1.044 [0.89–1.23]	0.600	1.696 [0.81–3.57]	0.164	
Frequency of drinking		Improved	1.111 [0.98–1.27]	0.108	3.303 [2.14–5.09]	< 0.001	
		Worsened	1.067 [0.97–1.18]	0.183	0.813 [0.64–1.04]	0.097	
Frequency of physical activity		Improved	1.046 [0.95–1.15]	0.344	0.838 [0.66–1.06]	0.142	
		Worsened	0.904 [0.82–1.00]	0.049	0.819 [0.68–0.99]	0.039	
		Improved	0.836 [0.76–0.92]	< 0.001	0.574 [0.48–0.69]	< 0.001	

MACE, major adverse cardiovascular events; BMI, body mass index; CVD, cardiovascular disease.

**Supplement 5. Multivariate analysis of all-cause death, non-fatal MI, and non-fatal stroke associated with lifestyle change between two biennial health screening periods in the prehypertension group**

Outcome	Parameter	Change	Male		Female		
			Adjusted hazard ratio	<i>P</i> -value	Adjusted hazard ratio	<i>P</i> -value	
			[95% CI]		[95% CI]		
All-cause death	Unhealthy lifestyle	Worsened	1.11 [1.00–1.22]	0.043	1.0 [0.84–1.22]	0.889	
		Improved	0.96 [0.88–1.05]	0.372	0.92 [0.78–1.09]	0.348	
	Unhealthy lifestyle (exclude BMI)	Worsened	1.20 [1.09–1.32]	< 0.001	1.00 [0.81–1.23]	0.961	
		Improved	0.93 [0.85–1.02]	0.139	0.83 [0.70–0.99]	0.04	
	BMI	Worsened	0.80 [0.71–0.91]	< 0.001	1.03 [0.83–1.29]	0.775	
		Improved	1.19 [1.05–1.35]	0.007	1.25 [1.01–1.54]	0.039	
	Currently smoking	Worsened	1.53 [1.31–1.78]	< 0.001	1.61 [0.91–2.86]	0.101	
		Improved	0.72 [0.63–0.83]	< 0.001	1.25 [0.72–2.16]	0.421	
	Frequency of drinking	Worsened	1.15 [1.04–1.26]	0.006	1.12 [0.88–1.43]	0.358	
		Improved	1.02 [0.92–1.13]	0.687	1.00 [0.70–1.43]	0.991	
	Frequency of physical activity	Worsened	1.00 [0.92–1.13]	0.996	1.23 [0.93–1.62]	0.148	
		Improved	0.91 [0.83–1.00]	0.043	0.91 [0.77–1.09]	0.301	
	Non-fatal MI	Unhealthy lifestyle	Worsened	1.05 [0.90–1.22]	0.571	0.97 [0.74–1.27]	0.826
			Improved	0.91 [0.78–1.05]	0.183	0.89 [0.71–1.13]	0.339
Unhealthy lifestyle (exclude BMI)		Worsened	1.07 [0.92–1.25]	0.384	0.95 [0.70–1.28]	0.72	
		Improved	0.94 [0.81–1.09]	0.395	0.89 [0.70–1.14]	0.349	
BMI		Worsened	1.16 [0.96–1.41]	0.132	0.71 [0.49–1.05]	0.066	

	Currently smoking	Improved	1.05 [0.85–1.23]	0.809	0.81 [0.60–1.11]	0.187
		Worsened	1.45 [1.12–1.88]	0.005	1.71 [0.70–4.14]	0.236
	Frequency of drinking	Improved	0.85 [0.70–1.03]	0.105	0.82 [0.37–1.81]	0.624
		Worsened	1.02 [0.88–1.19]	0.796	1.11 [0.79–1.56]	0.556
	Frequency of physical activity	Improved	1.16 [0.98–1.37]	0.089	1.04 [0.62–1.66]	0.957
		Worsened	0.95 [0.80–1.14]	0.605	1.10 [0.76–1.60]	0.622
<b>Non-fatal stroke</b>	Unhealthy lifestyle	Worsened	1.20 [1.09–1.33]	< 0.001	1.10 [0.97–1.25]	0.155
		Improved	0.94 [0.85–1.03]	0.185	0.93 [0.83–1.05]	0.246
	Unhealthy lifestyle (exclude BMI)	Worsened	1.15 [1.04–1.27]	0.007	1.04 [0.90–1.20]	0.603
		Improved	0.94 [0.85–1.03]	0.178	0.88 [0.78–0.99]	0.038
	BMI	Worsened	1.14 [1.01–1.29]	0.042	1.10 [0.94–1.28]	0.229
		Improved	0.95 [0.84–1.08]	0.445	1.00 [0.86–1.16]	0.982
	Currently smoking	Worsened	1.31 [1.11–1.54]	0.002	1.72 [1.13–2.62]	0.012
		Improved	0.77 [0.67–0.89]	< 0.001	0.57 [0.32–1.00]	0.052
	Frequency of drinking	Worsened	1.13 [1.02–1.24]	0.018	0.90 [0.75–1.08]	0.266
		Improved	1.08 [0.97–1.20]	0.145	1.13 [0.88–1.46]	0.333
	Frequency of physical activity	Worsened	1.15 [1.02–1.30]	0.022	1.24 [1.03–1.49]	0.021
		Improved	0.91 [0.82–1.00]	0.039	0.89 [0.79–1.00]	0.057
<b>CVD death</b>	Unhealthy lifestyle	Worsened	1.13 [0.82–1.56]	0.453	1.10 [0.67–1.80]	0.712
		Improved	0.89 [0.65–1.23]	0.478	0.76 [0.47–1.24]	0.278
	Unhealthy lifestyle (exclude BMI)	Worsened	1.07 [0.77–1.48]	0.696	0.95 [0.52–1.74]	0.857

		Improved	0.85 [0.62–1.17]	0.32	0.93 [0.58–1.51]	0.773
	BMI	Worsened	0.81 [0.52–1.26]	0.348	1.60 [0.94–2.71]	0.082
		Improved	1.05 [0.69–1.61]	0.817	0.94 [0.49–1.78]	0.845
	Currently smoking	Worsened	2.41 [1.53–3.78]	< 0.001	1.68 {0.41–6.86}	0.469
		Improved	0.56 [0.34–0.92]	0.023	2.70 [0.76–9.63]	0.125
	Frequency of drinking	Worsened	1.10 [0.80–1.52]	0.549	0.54 [0.22–1.35]	0.186
		Improved	1.04 [0.72–1.50]	0.834	0.82 [0.27–2.49]	0.72
	Frequency of physical activity	Worsened	0.83 [0.55–1.27]	0.391	0.99 {0.43–2.30}	0.986
		Improved	0.83 [0.61–1.14]	0.258	0.66 [0.39–1.14]	0.138

BMI, body mass index; CVD, cardiovascular disease.

**Supplement 6. Multivariate analysis of all-cause death, non-fatal MI, and non-fatal stroke associated with lifestyle change between two biennial health screening periods in the prediabetes group**

Outcome	Parameter	Change	Male		Female		
			Adjusted hazard ratio	<i>P</i> -value	Adjusted hazard ratio	<i>P</i> -value	
			[95% CI]		[95% CI]		
All-cause death	Unhealthy lifestyle	Worsened	1.18 [1.06–1.31]	0.002	1.10 [0.91–1.33]	0.325	
		Improved	1.04 [0.95–1.15]	0.397	0.56 [0.72–1.02]	0.076	
	Unhealthy lifestyle (exclude BMI)	Worsened	1.19 [1.08–1.32]	< 0.001	1.10 [0.89–1.35]	0.395	
		Improved	0.97 [0.88–1.06]	0.471	0.80 [0.67–0.96]	0.017	
	BMI	Worsened	0.88 [0.76–1.00]	0.056	0.86 [0.67–1.10]	0.224	
		Improved	1.21 [1.06–1.37]	0.004	1.00 [0.81–1.24]	0.968	
	Currently smoking	Worsened	1.55 [1.31–1.84]	< 0.001	1.18 [0.56–2.49]	0.664	
		Improved	0.89 [0.77–1.02]	0.102	0.98 [0.58–1.65]	0.931	
	Frequency of drinking	Worsened	1.10 [0.99–1.22]	0.072	0.96 [0.74–1.23]	0.724	
		Improved	1.03 [0.92–1.14]	0.648	0.87 [0.60–1.27]	0.475	
	Frequency of physical activity	Worsened	1.11 [0.97–1.26]	0.133	1.13 [0.83–1.52]	0.436	
		Improved	0.96 [0.87–1.06]	0.433	0.75 [0.62–0.91]	0.003	
	Non-fatal MI	Unhealthy lifestyle	Worsened	1.35 [1.13–1.61]	< 0.001	1.16 [0.88–1.53]	0.294
			Improved	1.17 [0.99–1.38]	0.066	0.86 [0.67–1.10]	0.219
Unhealthy lifestyle (exclude BMI)		Worsened	1.24 [1.04–1.47]	0.018	0.94 [0.68–1.30]	0.688	
		Improved	1.20 [1.02–1.41]	0.03	0.84 [0.65–1.09]	0.199	
BMI		Worsened	1.21 [0.98–1.51]	0.08	1.33 [0.97–1.82]	0.082	

	Currently smoking	Improved	0.76 [0.60–0.95]	0.016	0.86 [0.62–1.18]	0.339
		Worsened	1.55 [1.17–2.04]	0.002	1.34 [0.43–4.17]	0.619
	Frequency of drinking	Improved	0.92 [0.73–1.16]	0.487	1.24 [0.53–2.90]	0.621
		Worsened	0.97 [0.81–1.15]	0.695	0.99 [0.69–1.42]	0.963
	Frequency of physical activity	Improved	1.23 [1.03–1.47]	0.024	0.93 [0.54–1.62]	0.804
		Worsened	1.20 [0.97–1.47]	0.088	0.94 [0.60–1.46]	0.779
<b>Non-fatal stroke</b>	Unhealthy lifestyle	Worsened	1.21 [1.09–1.34]	< 0.001	1.10 [0.95–1.27]	0.212
		Improved	1.01 [0.92–1.12]	0.825	0.97 [0.86–1.11]	0.686
	Unhealthy lifestyle (exclude BMI)	Worsened	1.23 [1.11–1.37]	< 0.001	0.96 [0.82–1.14]	0.658
		Improved	1.04 [0.95–1.15]	0.398	0.97 [0.85–1.10]	0.631
	BMI	Worsened	1.15 [1.01–1.32]	0.041	1.23 [1.03–1.46]	0.02
		Improved	0.96 [0.84–1.09]	0.506	0.95 [0.81–1.12]	0.539
	Currently smoking	Worsened	1.40 [1.19–1.66]	< 0.001	0.85 [0.40–1.79]	0.667
		Improved	0.93 [0.80–1.09]	0.388	1.11 [0.68–1.82]	0.682
	Frequency of drinking	Worsened	1.15 [1.03–1.27]	0.011	0.94 [0.77–1.14]	0.498
		Improved	1.18 [1.06–1.32]	0.003	1.03 [0.77–1.38]	0.847
	Frequency of physical activity	Worsened	1.22 [1.07–1.38]	0.003	1.07 [0.86–1.34]	0.536
		Improved	0.97 [0.87–1.07]	0.483	1.04 [0.88–1.15]	0.93
<b>CVD death</b>	Unhealthy lifestyle	Worsened	1.13 [0.83–1.53]	0.429	0.88 [0.53–1.45]	0.609
		Improved	0.96 [0.72–1.27]	0.759	0.98 [0.64–1.49]	0.918
	Unhealthy lifestyle (exclude BMI)	Worsened	1.10 [0.81–1.49]	0.54	1.06 [0.61–1.84]	0.841



		Improved	0.90 [0.67–1.19]	0.448	1.10 [0.70–1.71]	0.689
	BMI	Worsened	1.17 [0.81–1.70]	0.407	1.34 [0.76–2.36]	0.317
		Improved	1.27 [0.89–1.82]	0.191	1.17 [0.71–1.93]	0.55
	Currently smoking	Worsened	1.24 [0.70–2.20]	0.471	N/A	N/A
		Improved	0.61 [0.39–0.96]	0.032	2.30 [0.62–8.57]	0.216
	Frequency of drinking	Worsened	0.87 [0.63–1.18]	0.368	0.66 [0.32–1.37]	0.26
		Improved	0.80 [0.58–1.11]	0.183	0.65 [0.22–1.88]	0.424
	Frequency of physical activity	Worsened	1.16 [0.80–1.70]	0.435	1.88 [0.78–4.55]	0.16
		Improved	1.07 [0.80–1.42]	0.665	1.13 [0.73–1.76]	0.587

MACE, major adverse cardiovascular events; BMI, body mass index; CVD, cardiovascular disease.