

Table S1: Baseline characteristics of the Golestan Cohort Study participants

Characteristics	Total N=50045	Women N=28811	Men N=21234
Age at baseline			
• <50	24615 (49.19)	14526 (50.42)	10089 (47.51)
• 50-59	15573 (31.12)	9290 (32.24)	6283 (29.59)
• 60-69	7359 (14.70)	3918 (13.60)	3441 (16.21)
• ≥ 70	2498 (4.99)	1077 (3.74)	1421 (6.69)
Ethnicity			
• Turkman	37253 (74.44)	21202 (73.59)	16051 (75.59)
• Others	12792 (25.56)	7609 (26.41)	5183 (24.41)
Marital status			
• Married	43955 (87.83)	23286 (80.82)	20669 (97.34)
• Unmarried*	6090 (12.17)	5525 (19.18)	565 (2.66)
Residence			
• Urban	10034 (20.05)	6100 (21.17)	3934 (18.53)
• Rural	40011 (79.95)	22711 (78.83)	17300 (81.47)
Education			
• No formal schooling	35118 (70.17)	24700 (85.73)	10418 (49.06)
• Primary Education	8463 (16.91)	2914 (10.11)	5549 (26.13)
• Middle school or higher	6464 (12.92)	1197 (4.15)	5267 (24.80)
Wealth score (Quintiles)			
• First	10746 (21.47)	6416 (22.27)	4330 (20.39)
• Second	9288 (18.56)	5450 (18.92)	3838 (18.07)
• Third	10874 (21.73)	6257 (21.72)	4617 (21.74)
• Fourth	9482 (18.95)	5383 (18.68)	4099 (19.30)
• Fifth	9655 (19.29)	5305 (18.41)	4350 (20.49)
Alcohol use			
• Never used	48316 (96.55)	28790 (99.93)	19526 (91.96)
• Ever used	1729 (3.45)	21 (0.07)	1708 (8.04)
Tobacco smoking/Nass chewing			
• Never used	39170 (78.28)	27744 (96.31)	11426 (53.82)
• Ever used	10867 (21.72)	1063 (3.69)	9804 (46.18)
Opium consumption			
• Never used	41545 (83.02)	26458 (91.83)	15087 (71.05)
• Ever used	8500 (16.98)	2353 (8.17)	6147 (28.95)
Physical activity (MET min/week)			
• 1 st tertile	17601 (35.25)	4882 (16.98)	12719 (60.07)
• 2 nd tertile	15815 (31.68)	11351 (39.84)	4464 (21.08)
• 3 rd tertile	16512 (33.07)	12520 (43.54)	3992 (18.85)
Body mass index (kg/m ²)			
• <18.5	2410 (4.82)	1153 (4.00)	1257 (5.92)
• 18.5-24.9	17929 (35.83)	8317 (28.87)	9612 (45.28)
• 25-29.9	16973 (33.92)	9701 (33.67)	7272 (34.26)
• ≥ 30	12725 (25.43)	9637 (33.45)	3088 (14.55)
Vegetable/fruit consumption (g/day)			
• 1 st tertile	16390 (33.33)	10393 (36.73)	5997 (28.73)
• 2 nd tertile	16390 (33.33)	9544 (33.73)	6846 (32.80)
• 3 rd tertile	16390 (33.33)	8358 (29.54)	8032 (38.48)
High blood pressure			
• No	28805 (57.56)	15519 (53.86)	13286 (62.57)
• Yes	21240 (42.44)	13292 (46.14)	7948 (37.43)
Diabetes mellitus			
• No	46601 (93.12)	26464 (91.85)	20137 (94.83)
• Yes	3444 (6.88)	2347 (8.15)	1097 (5.17)

Data are n (%). *Single, divorced, widowed, and others.

Table S2: Causes of death in the Golestan Cohort Study (2004-Aug 2017)*

Disease classification based on ICD-10 (Version:2016)	ICD code	<70	>=70	All
All		4018	2329	6347
I Certain infectious and parasitic diseases	A00-B99	112 (2.79)	98 (4.21)	210 (3.31)
Tuberculosis	A15-A19	36 (32.14)	16 (16.33)	52 (24.76)
Others		76 (67.86)	82 (83.67)	158 (75.24)
II Neoplasms	C00-D48	858 (21.35)	384 (16.49)	1242 (19.57)
Malignant neoplasm of oesophagus	C15	153 (17.83)	83 (21.61)	236 (19.00)
Malignant neoplasm of stomach	C16	155 (18.07)	83 (21.61)	238 (19.16)
Malignant neoplasm of colorectal	C18-20	47 (5.48)	18 (4.69)	65 (5.23)
Malignant neoplasms of hepatobiliary organs	C22-24	52 (6.06)	23 (5.99)	75 (6.04)
Malignant neoplasm of pancreas	C25	52 (6.06)	27 (7.03)	79 (6.36)
Malignant neoplasms of other digestive organs		7 (0.82)	1 (0.26)	8 (0.64)
Malignant neoplasm of bronchus and lung	C34	61 (7.11)	22 (5.73)	83 (6.68)
Malignant neoplasm of breast	C50	27 (3.15)	1 (0.26)	28 (2.25)
Malignant neoplasms of female genital organs	C51-58	56 (6.53)	6 (1.56)	62 (4.99)
Malignant neoplasms of male genital organs	C60-63	9 (1.05)	16 (4.17)	25 (2.01)
Malignant neoplasms of urinary tract	C64-C68	24 (2.80)	11 (2.86)	35 (2.82)
Malignant neoplasms of eye, brain and other parts of central nervous system	C69-72	47 (5.48)	10 (2.60)	57 (4.59)
Malignant neoplasms, stated or presumed to be primary, of lymphoid, haematopoietic and related tissue	C81-96	61 (7.11)	29 (7.55)	90 (7.25)
Other Malignant neoplasms		104 (12.12)	53 (13.80)	157 (12.64)
D00-D48In situ, benign, or unknown	D00-D48	3 (0.35)	1 (0.26)	4 (0.32)
III Diseases of the blood and blood-forming organs and certain disorders involving the immune mechanism	D50-D89	1 (0.02)	1 (0.04)	2 (0.03)
IV Endocrine, nutritional and metabolic diseases	E00-E90	54 (1.34)	14 (0.60)	68 (1.07)
Diabetes mellitus	E10-E14	47 (87.04)	13 (92.86)	60 (88.24)
Others		7 (12.96)	1 (7.14)	8 (11.76)
V Mental and behavioural disorders	F00-F99	30 (0.75)	61 (2.62)	91 (1.43)
VI Diseases of the nervous system	G00-G99	35 (0.87)	36 (1.55)	71 (1.12)
VII Diseases of the eye and adnexa	H00-H59	0	0	0
VIII Diseases of the ear and mastoid process	H60-H95	0	0	0
IX Diseases of the circulatory system	I00-I99	1669 (41.54)	922 (39.59)	2591 (40.82)
Ischaemic heart diseases	I20-I25	864 (51.77)	371 (40.24)	1235 (47.66)
Cardiac arrest		146	180 (10.78)	95 (10.30)
Heart failure		150	106 (6.35)	67 (7.27)
Cerebrovascular diseases	I60-I69	473 (28.34))	354 (38.39)	827 (31.92)
Others		46 (2.76)	35 (3.80)	81 (3.13)
X Diseases of the respiratory system	J00-J99	175 (4.36)	142 (6.10)	317 (4.99)
Acute infection	J00-J22	43 (24.57)	55 (38.73)	98 (30.91)
Chronic lower respiratory diseases	J40-J47	117 (66.86)	75 (52.82)	192 (60.57)
Others		15 (8.57)	12 (8.45)	27 (8.52)
XI Diseases of the digestive system	K00-K93	125 (3.11)	57 (2.45)	182 (2.87)
Diseases of Liver	K70-77	75 (60.00)	22 (38.60)	97 (53.30)
Others		50 (40.00)	35 (61.40)	85 46.70)
XII Diseases of the skin and subcutaneous tissue	L00-L99	3 (0.07)	1 (0.04)	4 (0.06)
XIII Diseases of the musculoskeletal system and connective tissue	M00-M99	9 (0.22)	0	9 (0.14)
XIV Diseases of the genitourinary system	N00-N99	74 (1.84)	50 (2.15)	124 (1.95)
Renal failure	N17-N19	71 (95.95)	48 (96.00)	119 (95.97)
Others		3 (4.05)	2 (4.00)	5 (4.03)
XV Pregnancy, childbirth and the puerperium	O00-O99	0	0	0
XVI Certain conditions originating in the perinatal period	P00-P96	0	0	0
XVII Congenital malformations, deformations and chromosomal abnormalities	Q00-Q99	2 (0.05)	0	2 (0.03)
XVIII Symptoms, signs and abnormal clinical and laboratory findings, not elsewhere classified	R00-R99	185 (4.60)	150 (6.44)	335 (5.28)
XIX Injury, poisoning and certain other consequences of external causes	S00-T98	9 (0.22)	6 (0.26)	15 (0.24)
XX External causes of morbidity and mortality	V01-Y98	235 (5.85)	52 (2.23)	287 (4.52)
Transport accidents	V01-V99	166 (70.64)	33 (63.46)	199 (69.34)
Others		69 (29.36)	19 (36.54)	88 (30.66)
XXI Factors influencing health status and contact with health services	Z00-Z99	0	0	0
XXII Codes for special purposes	U00-U85	0	0	0

* Cause of death not yet determined in 797 mortality cases: 442 deaths <70 and 355 deaths ≥70 years. In columns, n (%) of each main subgroups are calculated (Bold); also, within subgroups n (%) are shown, separately.

Table S3: The top 10 causes of death (Global Health Estimation, WHO Methods) in the Golestan Cohort Study*, after exclusion of deaths within the first year of follow-up

	40-49 years N=357	50-59 years N=1213	60-69 years N=1710	All premature death (<70) N=3280
1	IHD ^a 125 (35.01)	IHD ^a 385 (31.74)	IHD ^a 589 (34.44)	IHD ^a 1099 (33.51)
2	Road injury 45 (12.61)	Stroke 153 (12.61)	Stroke 274 (16.02)	Stroke 461 (14.05)
3	Stroke 34 (9.52)	Road injury 59 (4.86)	Stomach cancer 95 (5.56)	Stomach cancer 159 (4.85)
4	Esophagus cancer 19 (5.32)	Stomach cancer 58 (4.78)	Esophagus cancer 83 (4.85)	Esophagus cancer 153 (4.66)
5	Cirrhosis ^b 10 (2.80)	Esophagus cancer 51 (4.20)	COPD ^c 64 (3.74)	Road injury 151 (4.60)
6	Breast cancer 9 (2.52)	COPD ^c 38 (3.13)	Road injury 47 (2.75)	COPD ^c 106 (3.23)
7	Kidney diseases 9 (2.52)	Kidney diseases 32 (2.64)	Kidney diseases 42 (2.46)	Kidney diseases 83 (2.53)
8	Liver cancer 7 (1.96)	Colorectal cancers ^e 25 (2.06)	Lung cancers ^f 35 (2.05)	Lung cancers ^f 63 (1.92)
9	Brain cancers ^d 7 (1.96)	Cirrhosis ^b 25 (2.06)	Alzheimer ^h 33 (1.93)	Cirrhosis ^b 62 (1.89)
10	Diabetes mellitus 6 (1.68)	Pancreas cancer 23 (1.90)	Ovary cancer 20 (1.65)	Ovary cancer 27 (1.58)
	Stomach cancer 6 (1.68)	Lung cancers ^f 23 (1.90)	Cirrhosis ^b 27 (1.58)	Pancreas cancer 48 (1.46)
	Asthma 4 (1.12)	Ovary cancer 20 (1.65)		

*After redistribution of the garbage codes (295 Cardiovascular disease cases: I10, I26.9, I46, I47.2, I49.0, and I50; 4 Cancer cases: C76, C80, and C97; and 8 External cause cases: Y10-Y34 and Y87.2); and 158 cases of symptoms, signs and ill-defined conditions (R00-R94, R96-R99). Excluded: 441 undetermined mortality causes.

^aIschemic heart disease, ^bCirrhosis of the liver, ^cChronic obstructive respiratory diseases, ^dBrain and nervous system cancers, ^eColon and rectum cancers,

^fTrachea, bronchus, lung cancers, ^gLower respiratory tract infection, ^hAlzheimer and other dementias.

Table S4: Hazard ratios for associations between demographic and lifestyle risk factors and premature mortality after exclusion of deaths within the first year of follow-up, in the Golestan Cohort Study*

Variable	Adjusted HR (95% CI) **
Sex	
• Women	1
• Men	1.39 (1.25-1.54)
Ethnicity	
• Turkman	1
• Non-Turkman	0.99 (0.91-1.08)
Marital Status	
• Married	1
• Other	1.17 (1.05-1.31)
Residence	
• Urban	1
• Rural	0.91 (0.82-1.02)
Education	
• No formal schooling	1
• Primary Education	0.86 (0.77-0.96)
• Middle school or higher	0.84 (0.73-0.97)
Wealth Score	
• 1 st quintile	1
• 2 nd quintile	0.88 (0.79-0.97)
• 3 rd quintile	0.69 (0.62-0.77)
• 4 th quintile	0.70 (0.63-0.79)
• 5 th quintile	0.56 (0.49-0.64)
Alcohol Use	
• Never	1
• Ever used	1.02 (0.87-1.21)
Tobacco Smoking/Nass chewing	
• Never	1
• Ever used	1.39 (1.25-1.54)
Opium Consumption	
• Never	1
• Ever used	1.67 (1.52-1.83)
Physical Activity (MET min/week)	
• 1 st Tertile	1
• 2 nd Tertile	0.83 (0.76-0.90)
• 3 rd Tertile	0.68 (0.62-0.75)
Body Mass Index (kg/m²)	
• <18.5	1.38 (1.20-1.59)
• 18.5-24.9	1
• 25-29.9	0.97 (0.89-1.06)
• ≥30	0.93 (0.84-1.03)
Vegetable/Fruit Consumption (g/day)	
• 1 st Tertile	1
• 2 nd Tertile	0.97 (0.89-1.06)
• 3 rd Tertile	0.88 (0.80-0.97)
High Blood Pressure	
• No	1
• Yes	1.50 (1.39-1.62)
Diabetes Mellitus	
• No	1
• Yes	2.39 (2.14-2.66)

*Exclude: age ≥70 years at baseline and history of ischemic heart disease, stroke, or cancer. **Adjusted for age at enrolment and all variables listed.

Table S5: Hazard ratios for associations between demographic and lifestyle risk factors and premature mortality based on sex, in the Golestan Cohort Study*

Variable	Men HR (95% CI) **	Women HR (95% CI) **
Ethnicity		
• Turkman	1	1
• Non-Turkman	0.94 (0.83-1.06)	1.03 (0.91-1.17)
Marital Status		
• Married	1	1
• Other	1.44 (1.13-1.84)	1.08 (0.96-1.23)
Residence		
• Urban	1	1
• Rural	0.84 (0.72-0.97)	0.96 (0.83-1.12)
Education		
• No formal schooling	1	1
• Primary Education	0.83 (0.73-0.94)	0.95 (0.75-1.20)
• Middle school or higher	0.87 (0.75-1.02)	0.41 (0.24-0.70)
Wealth Score		
• 1 st quintile	1	1
• 2 nd quintile	0.87 (0.76-0.99)	0.98 (0.85-1.13)
• 3 rd quintile	0.71 (0.62-0.82)	0.70 (0.60-0.82)
• 4 th quintile	0.72 (0.61-0.83)	0.75 (0.63-0.88)
• 5 th quintile	0.54 (0.45-0.64)	0.67 (0.55-0.81)
Alcohol Use		
• Never	1	1
• Ever used	1.05 (0.89-1.24)	0.58 (0.08-4.10)
Tobacco Smoking/Nass chewing		
• Never	1	1
• Ever used	1.42 (1.27-1.58)	1.45 (1.18-1.79)
Opium Consumption		
• Never	1	1
• Ever used	1.51 (1.36-1.69)	2.01 (1.74-2.32)
Physical Activity (MET min/week)		
• 1 st Tertile	1	1
• 2 nd Tertile	0.84 (0.74-0.95)	0.75 (0.66-0.85)
• 3 rd Tertile	0.83 (0.73-0.95)	0.55 (0.48-0.64)
Body Mass Index (kg/m²)		
• <18.5	1.47 (1.24-1.73)	1.27 (1.01-1.58)
• 18.5-24.9	1	1
• 25-29.9	1.00 (0.90-1.13)	0.90 (0.79-1.02)
• ≥30	1.06 (0.91-1.23)	0.82 (0.72-0.95)
Vegetable/Fruit Consumption (g/day)		
• 1 st Tertile	1	1
• 2 nd Tertile	0.92 (0.82-1.04)	0.99 (0.88-1.11)
• 3 rd Tertile	0.84 (0.75-0.95)	0.90 (0.79-1.03)
High Blood Pressure		
• No	1	1
• Yes	1.48 (1.34-1.63)	1.52 (1.36-1.69)
Diabetes Mellitus		
• No	1	1
• Yes	2.48 (2.12-2.91)	2.29 (1.98-2.63)

*Exclude: age ≥70 years at baseline and history of ischemic heart disease, stroke, or cancer. **Adjusted for age at enrolment and all variables listed.

Table S6: Hazard ratios for associations between demographic and lifestyle risk factors and premature mortality based on age, in the Golestan Cohort Study*

Variable	<55 years HR (95% CI) **	≥55 years HR (95% CI) **
Sex		
• Women	1	1
• Men	1.48 (1.30-1.70)	1.23 (1.06-1.42)
Ethnicity		
• Turkman	1	1
• Non-Turkman	1.01 (0.90-1.14)	0.98 (0.86-1.11)
Marital Status		
• Married	1	1
• Other	1.42 (1.20-1.67)	1.00 (0.86-1.16)
Residence		
• Urban	1	1
• Rural	0.94 (0.81-1.08)	0.87 (0.75-1.02)
Education		
• No formal schooling	1	1
• Primary Education	0.83 (0.72-0.94)	0.88 (0.73-1.05)
• Middle school or higher	0.81 (0.68-0.96)	0.90 (0.70-1.15)
Wealth Score		
• 1 st quintile	1	1
• 2 nd quintile	0.87 (0.77-1.00)	0.95 (0.82-1.10)
• 3 rd quintile	0.70 (0.61-0.80)	0.70 (0.60-0.81)
• 4 th quintile	0.74 (0.63-0.86)	0.70 (0.59-0.82)
• 5 th quintile	0.55 (0.46-0.66)	0.61 (0.50-0.75)
Alcohol Use		
• Never	1	1
• Ever used	0.99 (0.80-1.23)	1.10 (0.86-1.41)
Tobacco Smoking/Nass chewing		
• Never	1	1
• Ever used	1.53 (1.34-1.75)	1.23 (1.07-1.42)
Opium Consumption		
• Never	1	1
• Ever used	1.71 (1.52-1.93)	1.67 (1.47-1.91)
Physical Activity (MET min/week)		
• 1 st Tertile	1	1
• 2 nd Tertile	0.86 (0.76-0.96)	0.76 (0.67-0.86)
• 3 rd Tertile	0.68 (0.60-0.78)	0.67 (0.57-0.78)
Body Mass Index (kg/m²)		
• <18.5	1.53 (1.28-1.83)	1.26 (1.03-1.53)
• 18.5-24.9	1	1
• 25-29.9	1.04 (0.92-1.17)	0.87 (0.77-0.99)
• ≥30	1.00 (0.88-1.15)	0.82 (0.70-0.95)
Vegetable/Fruit Consumption (g/day)		
• 1 st Tertile	1	1
• 2 nd Tertile	0.97 (0.86-1.08)	0.94 (0.83-1.06)
• 3 rd Tertile	0.88 (0.78-1.00)	0.86 (0.75-0.98)
High Blood Pressure		
• No	1	1
• Yes	1.53 (1.39-1.69)	1.46 (1.31-1.62)
Diabetes Mellitus		
• No	1	1
• Yes	2.45 (2.10-2.84)	2.31 (1.99-2.68)

*Exclude: age ≥70 years at baseline and history of ischemic heart disease, stroke, or cancer. **Adjusted for age at enrolment and all variables listed.