Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eMethods. Supplemental Methods

Exposure: Reproductive factors

During National Cancer Screening Program, information about age at menarche, age at menopause, parity, total lifetime breast feeding history, and use of hormone therapy (HT) / oral contraceptives (OCs) was collected by self-administered questionnaire. Age at menarche was categorized as \leq 12 years, 13-14 years, 15-16 years, and \geq 17 years, to be consistent with the distribution of age at menarche among Korean women. Age at menopause was categorized as \leq 40 years, 40-44 years, 45-49 years, 50-54 years, and \leq 55 years. Reproductive span was calculated as the interval between the age at menarche and the age at menopause. Parity was categorized as 0 (nulliparous), 1 (primiparous), or \leq 2 (multiparous). Total lifetime breast-feeding history was categorized as never, \leq 6 months, 6-11 months or \leq 12 total months. The duration of HT was categorized as never, \leq 2 years, \leq 5 years, or unknown. The duration of OC use was categorized as never, \leq 1 years, \leq 1 years, \leq 2 years, or unknown.

Study outcome: Fractures

The primary endpoint was newly diagnosed fractures during the follow-up period. Fractures were defined by ICD-10 codes as follows: (1) vertebral (S22.0, S22.1, S32.0, M48.4 and M48.5), (2) hip (S72.0 and S72.1), or (3) other fractures. Other fractures included clavicle (S42.0), upper arm (S42.2 and S42.3), wrist (S52.5 and S52.6), and ankle (S82.3, S82.5 and S82.6). Vertebral or other fractures were identified when subjects had two or more outpatient visits with relevant diagnosis codes within 12 months. Hip fracture was defined as one hospitalization with the relevant diagnosis code. Any fracture was defined as at least one of these three fracture types.

Covariates

Income level was based on the monthly insurance premium because insurance contribution is determined based on income level and not on health risk in Korea. Subjects were classified as never, ex-, or current smokers. Based on daily alcohol consumption, alcohol consumption was classified as none (0 g/day), moderate (<30 g/day), or heavy (≥30 g/day). Regular exercise was defined as performing >30 minutes of moderate physical activity at least five times per week or >20 minutes of strenuous physical activity at least three times per week. Subjects were categorized into five groups based on body mass index (BMI, kg/m²) according to the Asia-Pacific criteria of the World Health Organization as follows: underweight (<18.5 kg/m²), normal (18.5 to <23 kg/m²), overweight (23 to <25 kg/m²), obese (25 to <30 kg/m²), and severely obese (≥30 kg/m²). Systolic and diastolic blood pressure (BP) were measured in a seated position after at least 5 minutes rest. Blood samples for measurement of serum fasting glucose and lipid levels were drawn after an overnight fast. Baseline comorbidities (hypertension, diabetes mellitus, dyslipidemia, and cancer) were identified based on the combination of past medical history and ICD-10 and prescription codes.

eTable 1. Hazard Ratios and 95% Confidence Intervals of Any Fracture According to Reproductive Factors

| Reproductive factors | Subjects (N) | Events (n) | Follow-up duration (PYs) | Incidence rate (per 1,000 PYs) | Model 1 | Model 2 | Model 3 |
|--------------------------------|-----------------|---------------|--------------------------------|---|-------------------|-------------------|-------------------|
| Age at menarche, y | | | | , | | | |
| ≤12 | 12,580 | 1,285 | 99,083.4 | 13.0 | 1 (ref.) | 1 (ref.) | |
| 13-14 | 159,722 | 19,250 | 1,245,522.7 | 15.5 | 1.19 (1.12, 1.26) | 1.10 (1.04, 1.16) | |
| 15-16 | 500,445 | 70,016 | 3,853,210.6 | 18.2 | 1.40 (1.32, 1.48) | 1.16 (1.10, 1.23) | |
| ≥17 | 599,368 | 99,332 | 4,538,316.8 | 21.9 | 1.69 (1.60, 1.78) | 1.24 (1.17, 1.31) | |
| Age at menopause, y | | | | | | | |
| <40 | 21,101 | 3,748 | 157,201.0 | 23.8 | 1 (ref.) | 1 (ref.) | |
| 40-44 | 71,142 | 11,861 | 535,222.2 | 22.2 | 0.93 (0.90, 0.96) | 0.97 (0.93, 1.01) | |
| 45-49 | 345,678 | 51,249 | 2,648,260.5 | 19.4 | 0.81 (0.78, 0.84) | 0.94 (0.91, 0.97) | |
| 50-54 | 699,018 | 101,617 | 5,364,751.4 | 18.9 | 0.79 (0.77, 0.82) | 0.90 (0.88, 0.93) | |
| ≥55 | 135,176 | 21,408 | 1,030,698.4 | 20.8 | 0.87 (0.84, 0.90) | 0.89 (0.86, 0.93) | |
| Reproductive span, y | | | | | | | |
| <30 | 170,474 | 29,718 | 1,277,414.9 | 23.3 | 1 (ref.) | | 1 (ref.) |
| 30-34 | 529,589 | 80,796 | 4,043,760.9 | 20.0 | 0.86 (0.85, 0.87) | | 0.94 (0.93, 0.95) |
| 35-39 | 490,384 | 67,381 | 3,788,551.8 | 17.8 | 0.76 (0.75, 0.77) | | 0.89 (0.88, 0.90) |
| ≥40 | 81,668 | 11,988 | 626,405.9 | 19.1 | 0.82 (0.81, 0.84) | | 0.86 (0.84, 0.88) |
| Parity | | | | | | | |
| Nulliparity | 32,006 | 4,242 | 247,443.9 | 17.1 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| 1 | 79,411 | 9,446 | 619,475.1 | 15.2 | 0.89 (0.86, 0.92) | 0.96 (0.92, 0.99) | 0.95 (0.92, 0.99) |
| ≥2 | 1,160,698 | 176,195 | 8,869,214.4 | 19.9 | 1.16 (1.12, 1.20) | 0.99 (0.95, 1.02) | 0.98 (0.95, 1.02) |
| Duration of breast feeding, mo | | | | | | | |
| Never | 86,477 | 10,449 | 673,737.5 | 15.5 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <6 | 86,290 | 9,844 | 675,687.5 | 14.6 | 0.94 (0.91, 0.97) | 0.97 (0.94, 0.99) | 0.97 (0.94, 0.99) |
| 6-11 | 225,607 | 29,142 | 1,750,226.8 | 16.7 | 1.07 (1.05, 1.10) | 1.00 (0.97, 1.02) | 1.00 (0.98, 1.02) |
| ≥12 | 873,741 | 140,448 | 6,636,481.7 | 21.2 | 1.37 (1.34, 1.39) | 1.05 (1.03, 1.08) | 1.06 (1.04, 1.09) |

| Duration of HT, y | | | | | | | |
|-----------------------|-----------|---------|-------------|------|-------------------|-------------------|-------------------|
| Never | 1,023,519 | 156,924 | 7,801,630.3 | 20.1 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <2 | 118,146 | 15,351 | 922,759.7 | 16.6 | 0.83 (0.81, 0.84) | 0.99 (0.97, 1.01) | 0.99 (0.97, 1.00) |
| 2-4 | 48,812 | 6,113 | 382,048.7 | 16.0 | 0.80 (0.78, 0.82) | 0.93 (0.91, 0.95) | 0.93 (0.91, 0.95) |
| ≥5 | 37,581 | 4,723 | 294,392.8 | 16.0 | 0.80 (0.77, 0.82) | 0.85 (0.83, 0.88) | 0.85 (0.83, 0.88) |
| Unknown | 44,057 | 6,772 | 335,301.9 | 20.2 | 1.00 (0.98, 1.03) | 1.01 (0.98, 1.04) | 1.01 (0.98, 1.04) |
| Duration of OC use, y | | | | | | | |
| Never | 1,015,265 | 151,858 | 7,762,969.8 | 19.6 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <1 | 116,384 | 16,603 | 898,277.1 | 18.5 | 0.95 (0.93,0.96) | 1.00 (0.98, 1.02) | 1.00 (0.99, 1.02) |
| ≥1 | 77,581 | 11,863 | 594,891.8 | 19.9 | 1.02 (1.00, 1.04) | 1.02 (1.00, 1.04) | 1.03 (1.01, 1.05) |
| Unknown | 62,885 | 9,559 | 479,994.8 | 19.9 | 1.02 (1.00, 1.04) | 1.02 (1.00, 1.05) | 1.02 (1.00, 1.04) |

Model 1: crude model

Model 2: the full model included age, age at menarche, age at menopause, parity, duration of breast feeding, duration of HT, duration of OC use, alcohol consumption, smoking, regular exercise, income, body mass index, hypertension, diabetes, dyslipidemia, and cancer.

eTable 2. Hazard Ratios and 95% Confidence Intervals of Vertebral Fracture According to Reproductive Factors

| Reproductive factors | Subjects (N) | Events (n) | Follow-up duration (PYs) | Incidence rate (per 1,000 PYs) | Model 1 | Model 2 | Model 3 |
|--------------------------------|-----------------|---------------|--------------------------------|---|-------------------|-------------------|-------------------|
| Age at menarche, y | | | | , | | | |
| ≤12 | 12,580 | 357 | 99,083.4 | 3.6 | 1 (ref.) | 1 (ref.) | |
| 13-14 | 159,722 | 5,899 | 1,245,522.7 | 4.7 | 1.32 (1.18, 1.46) | 1.12 (1.01, 1.25) | |
| 15-16 | 500,445 | 25,166 | 3,853,210.6 | 6.5 | 1.82 (1.64, 2.02) | 1.26 (1.14, 1.40) | |
| ≥17 | 599,368 | 41,310 | 4,538,316.8 | 9.1 | 2.53 (2.28, 2.81) | 1.42 (1.28, 1.58) | |
| Age at menopause, y | | | | | | | |
| <40 | 21,101 | 1,764 | 157,201.0 | 11.2 | 1 (ref.) | 1 (ref.) | |
| 40-44 | 71,142 | 5,285 | 535,222.2 | 9.9 | 0.88 (0.83, 0.93) | 0.94 (0.89, 0.99) | |
| 45-49 | 345,678 | 20,155 | 2,648,260.5 | 7.6 | 0.68 (0.65, 0.71) | 0.88 (0.84, 0.93) | |
| 50-54 | 699,018 | 37,561 | 5,364,751.4 | 7.0 | 0.62 (0.59, 0.65) | 0.81 (0.77, 0.85) | |
| ≥55 | 135,176 | 7,967 | 1,030,698.4 | 7.7 | 0.69 (0.65, 0.72) | 0.77 (0.73, 0.81) | |
| Reproductive span, y | | | | | | | |
| <30 | 170,474 | 13,411 | 1,277,414.9 | 10.5 | 1 (ref.) | | 1 (ref.) |
| 30-34 | 529,589 | 31,978 | 4,043,760.9 | 7.9 | 0.75 (0.74, 0.77) | | 0.89 (0.87, 0.91) |
| 35-39 | 490,384 | 23,135 | 3,788,551.8 | 6.1 | 0.58 (0.57, 0.59) | | 0.79 (0.77, 0.81) |
| ≥40 | 81,668 | 4,208 | 626,405.9 | 6.7 | 0.64 (0.62, 0.66) | | 0.73 (0.71, 0.76) |
| Parity | | | | | | | |
| Nulliparity | 32,006 | 1,244 | 247,443.9 | 5.0 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| 1 | 79,411 | 2,579 | 619,475.1 | 4.2 | 0.83 (0.77, 0.89) | 0.90 (0.84, 0.97) | 0.90 (0.84, 0.97) |
| ≥2 | 1,160,698 | 68,909 | 8,869,214.4 | 7.8 | 1.55 (1.46, 1.64) | 1.07 (1.00, 1.14) | 1.06 (1.00, 1.13) |
| Duration of breast feeding, mo | | | | | | | |
| Never | 86,477 | 2,889 | 673,737.5 | 4.3 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <6 | 86,290 | 2,747 | 675,687.5 | 4.1 | 0.95 (0.90, 1.00) | 0.97 (0.92, 1.03) | 0.97 (0.92, 1.02) |
| 6-11 | 225,607 | 9,303 | 1,750,226.8 | 5.3 | 1.24 (1.19, 1.29) | 1.03 (0.99, 1.08) | 1.04 (0.99, 1.09) |
| ≥12 | 873,741 | 57,793 | 6,636,481.7 | 8.7 | 2.03 (1.96, 2.11) | 1.22 (1.17, 1.27) | 1.23 (1.18, 1.29) |

| Duration of HT, y | | | | | | | |
|-----------------------|-----------|--------|-------------|-----|-------------------|-------------------|-------------------|
| Never | 1,023,519 | 61,863 | 7,801,630.3 | 7.9 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <2 | 118,146 | 4,829 | 922,759.7 | 5.2 | 0.66 (0.64, 0.68) | 0.95 (0.92, 0.98) | 0.95 (0.92, 0.98) |
| 2-4 | 48,812 | 1,893 | 382,048.7 | 5.0 | 0.62 (0.60, 0.65) | 0.88 (0.84, 0.92) | 0.88 (0.84, 0.92) |
| ≥5 | 37,581 | 1,570 | 294,392.8 | 5.3 | 0.67 (0.64, 0.71) | 0.82 (0.78, 0.86) | 0.81 (0.77, 0.86) |
| Unknown | 44,057 | 2,577 | 335,301.9 | 7.7 | 0.97 (0.93, 1.01) | 0.98 (0.94, 1.03) | 0.98 (0.94, 1.03) |
| Duration of OC use, y | | | | | | | |
| Never | 1,015,265 | 58,466 | 7,762,969.8 | 7.5 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <1 | 116,384 | 6,039 | 898,277.1 | 6.7 | 0.89 (0.87, 0.92) | 1.01 (0.99, 1.04) | 1.02 (0.99, 1.04) |
| ≥1 | 77,581 | 4,552 | 594,891.8 | 7.7 | 1.02 (0.99, 1.05) | 1.06 (1.03, 1.09) | 1.06 (1.03, 1.10) |
| Unknown | 62,885 | 3,675 | 479,994.8 | 7.7 | 1.02 (0.98, 1.05) | 1.06 (1.02, 1.10) | 1.05 (1.01, 1.09) |

Model 1: crude model

Model 2: the full model included age, age at menarche, age at menopause, parity, duration of breast feeding, duration of HT, duration of OC use, alcohol consumption, smoking, regular exercise, income, body mass index, hypertension, diabetes, dyslipidemia, and cancer.

eTable 3. Hazard Ratios and 95% Confidence Intervals of Hip Fracture According to Reproductive Factors

| Reproductive factors | Subjects (N) | Events (n) | Follow-up duration (PYs) | Incidence rate (per 1,000 PYs) | Model 1 | Model 2 | Model 3 |
|--------------------------------|-----------------|---------------|--------------------------------|---|-------------------|-------------------|-------------------|
| Age at menarche, y | | | | | | | |
| ≤12 | 12,580 | 47 | 99,083.4 | 0.5 | 1 (ref.) | 1 (ref.) | |
| 13-14 | 159,722 | 886 | 1,245,522.7 | 0.7 | 1.50 (1.12, 2.01) | 1.18 (0.88, 1.58) | |
| 15-16 | 500,445 | 3,845 | 3,853,210.6 | 1.0 | 2.10 (1.58, 2.80) | 1.18 (0.89, 1.58) | |
| ≥17 | 599,368 | 6,375 | 4,538,316.8 | 1.4 | 2.96 (2.22, 3.94) | 1.23 (0.93, 1.64) | |
| Age at menopause, y | | | | | | | |
| <40 | 21,101 | 300 | 157,201.0 | 1.9 | 1 (ref.) | 1 (ref.) | |
| 40-44 | 71,142 | 905 | 535,222.2 | 1.7 | 0.89 (0.78, 1.01) | 0.95 (0.84, 1.09) | |
| 45-49 | 345,678 | 3,052 | 2,648,260.5 | 1.2 | 0.60 (0.54, 0.68) | 0.91 (0.81, 1.03) | |
| 50-54 | 699,018 | 5,622 | 5,364,751.4 | 1.0 | 0.55(0.49, 0.62) | 0.86 (0.77, 0.97) | |
| ≥55 | 135,176 | 1,274 | 1,030,698.4 | 1.2 | 0.65 (0.57, 0.74) | 0.88 (0.78, 1.00) | |
| Reproductive span, y | | | | | | | |
| <30 | 170,474 | 2,206 | 1,277,414.9 | 1.7 | 1 (ref.) | | 1 (ref.) |
| 30-34 | 529,589 | 4,964 | 4,043,760.9 | 1.2 | 0.71 (0.68, 0.75) | | 0.93 (0.88, 0.98) |
| 35-39 | 490,384 | 3,301 | 3,788,551.8 | 0.9 | 0.51 (0.48, 0.53) | | 0.88 (0.83, 0.93) |
| ≥40 | 81,668 | 682 | 626,405.9 | 1.1 | 0.63 (0.58, 0.69) | | 0.87 (0.80, 0.95) |
| Parity | | | | | | | |
| Nulliparity | 32,006 | 202 | 247,443.9 | 0.8 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| 1 | 79,411 | 439 | 619,475.1 | 0.7 | 0.87 (0.73, 1.02) | 1.13 (0.95, 1.34) | 1.12 (0.95, 1.34) |
| ≥2 | 1,160,698 | 10,512 | 8,869,214.4 | 1.2 | 1.45 (1.26, 1.66) | 1.01 (0.87, 1.19) | 1.01 (0.87, 1.18) |
| Duration of breast feeding, mo | | | | | | | |
| Never | 86,477 | 494 | 673,737.5 | 0.7 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <6 | 86,290 | 385 | 675,687.5 | 0.6 | 0.78 (0.68, 0.89) | 0.84 (0.73, 0.97) | 0.84 (0.73, 0.97) |
| 6-11 | 225,607 | 1,393 | 1,750,226.8 | 0.8 | 1.08 (0.98, 1.20) | 0.85 (0.76, 0.95) | 0.85 (0.76, 0.95) |
| ≥12 | 873,741 | 8,881 | 6,636,481.7 | 1.3 | 1.82 (1.66, 1.99) | 0.84 (0.76, 0.93) | 0.84 (0.76, 0.93) |

| Duration of HT, y | | | | | | | |
|-----------------------|-----------|-------|-------------|-----|-------------------|-------------------|-------------------|
| Never | 1,023,519 | 9,816 | 7,801,630.3 | 1.3 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <2 | 118,146 | 522 | 922,759.7 | 0.6 | 0.45 (0.41, 0.49) | 0.90 (0.82, 0.98) | 0.90 (0.82, 0.98) |
| 2-4 | 48,812 | 198 | 382,048.7 | 0.5 | 0.41 (0.36, 0.47) | 0.80 (0.69, 0.92) | 0.80 (0.69, 0.92) |
| ≥5 | 37,581 | 202 | 294,392.8 | 0.7 | 0.54 (0.47, 0.63) | 0.86 (0.74, 0.99) | 0.86 (0.75, 0.99) |
| Unknown | 44,057 | 415 | 335,301.9 | 1.2 | 0.99 (0.89, 1.09) | 0.98 (0.88, 1.09) | 0.98 (0.88, 1.09) |
| Duration of OC use, y | | | | | | | |
| Never | 1,015,265 | 9,159 | 7,762,969.8 | 1.2 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <1 | 116,384 | 776 | 898,277.1 | 0.9 | 0.73 (0.68, 0.79) | 0.98 (0.91, 1.06) | 0.99 (0.92, 1.06) |
| ≥1 | 77,581 | 637 | 594,891.8 | 1.1 | 0.91 (0.84, 0.99) | 1.06 (0.97, 1.15) | 1.06 (0.98, 1.15) |
| Unknown | 62,885 | 581 | 479,994.8 | 1.2 | 1.03 (0.95, 1.12) | 1.03 (0.94, 1.13) | 1.03 (0.94, 1.13) |

Model 1: crude model

Model 2: the full model included age, age at menarche, age at menopause, parity, duration of breast feeding, duration of HT, duration of OC use, alcohol consumption, smoking, regular exercise, income, body mass index, hypertension, diabetes, dyslipidemia, and cancer.

eTable 4. Hazard Ratios and 95% Confidence Intervals of Other Fractures According to Reproductive Factors

| Reproductive factors | Subjects (N) | Events (n) | Follow-up duration (PYs) | Incidence rate (per 1,000 PYs) | Model 1 | Model 2 | Model 3 |
|--------------------------------|-----------------|---------------|--------------------------------|---|-------------------|-------------------|-------------------|
| Age at menarche, y | | | | • | | | |
| ≤12 | 12,580 | 887 | 99,083.4 | 9.0 | 1 (ref.) | 1 (ref.) | |
| 13-14 | 159,722 | 12,530 | 1,245,522.7 | 10.1 | 1.12 (1.05, 1.20) | 1.10 (1.03, 1.18) | |
| 15-16 | 500,445 | 41,357 | 3,853,210.6 | 10.7 | 1.20 (1.12, 1.28) | 1.15 (1.07, 1.23) | |
| ≥17 | 599,368 | 52,121 | 4,538,316.8 | 11.5 | 1.28 (1.20, 1.37) | 1.19 (1.12, 1.28) | |
| Age at menopause, y | | | | | | | |
| <40 | 21,101 | 1,707 | 157,201.0 | 10.9 | 1 (ref.) | 1 (ref.) | |
| 40-44 | 71,142 | 5,728 | 535,222.2 | 10.7 | 0.99 (0.93, 1.04) | 1.00 (0.94, 1.05) | |
| 45-49 | 345,678 | 28,286 | 2,648,260.5 | 10.7 | 0.98 (0.94, 1.03) | 1.01 (0.97, 1.07) | |
| 50-54 | 699,018 | 58,917 | 5,364,751.4 | 11.0 | 1.01 (0.96, 1.06) | 1.03 (0.99, 1.09) | |
| ≥55 | 135,176 | 12,257 | 1,030,698.4 | 11.9 | 1.10 (1.04, 1.15) | 1.09 (1.03, 1.14) | |
| Reproductive span, y | | | | | | | |
| < 30 | 170,474 | 14,252 | 1,277,414.9 | 11.2 | 1 (ref.) | | 1 (ref.) |
| 30-34 | 529,589 | 44,244 | 4,043,760.9 | 10.9 | 0.98 (0.96, 1.00) | | 1.00 (0.98, 1.02) |
| 35-39 | 490,384 | 41,257 | 3,788,551.8 | 10.9 | 0.98 (0.96, 1.00) | | 1.00 (0.99, 1.02) |
| ≥40 | 81,668 | 7,142 | 626,405.9 | 11.4 | 1.02 (0.99, 1.05) | | 1.02 (0.99, 1.05) |
| Parity | | | | | | | |
| Nulliparity | 32,006 | 2,817 | 247,443.9 | 11.4 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| 1 | 79,411 | 6,463 | 619,475.1 | 10.4 | 0.92 (0.88, 0.96) | 0.94 (0.89, 0.98) | 0.93 (0.89, 0.98) |
| ≥2 | 1,160,698 | 97,615 | 8,869,214.4 | 11.0 | 0.97 (0.93, 1.00) | 0.93 (0.89, 0.97) | 0.93 (0.89, 0.97) |
| Duration of breast feeding, mo | | | | | | | |
| Never | 86,477 | 7,122 | 673,737.5 | 10.6 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <6 | 86,290 | 6,752 | 675,687.5 | 10.0 | 0.95 (0.91, 0.98) | 0.97 (0.93, 1.00) | 0.97 (0.93, 1.00) |
| 6-11 | 225,607 | 18,563 | 1,750,226.8 | 10.6 | 1.00 (0.98, 1.03) | 1.00 (0.97, 1.03) | 1.01 (0.98, 1.04) |
| ≥12 | 873,741 | 74,458 | 6,636,481.7 | 11.2 | 1.06 (1.04, 1.09) | 1.02 (0.99, 1.05) | 1.03 (1.00, 1.06) |

| Duration of HT, y | | | | | | | |
|-----------------------|-----------|--------|-------------|------|-------------------|-------------------|-------------------|
| Never | 1,023,519 | 86,001 | 7,801,630.3 | 11.0 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <2 | 118,146 | 10,066 | 922,759.7 | 10.9 | 0.99 (0.97, 1.01) | 1.02 (1.00, 1.04) | 1.02 (1.00, 1.04) |
| 2-4 | 48,812 | 4,051 | 382,048.7 | 10.6 | 0.96 (0.93, 0.99) | 0.98 (0.95, 1.02) | 0.98 (0.95, 1.01) |
| ≥5 | 37,581 | 2,968 | 294,392.8 | 10.1 | 0.92 (0.88, 0.95) | 0.92 (0.88, 0.95) | 0.91 (0.88, 0.95) |
| Unknown | 44,057 | 3,809 | 335,301.9 | 11.4 | 1.03 (1.00, 1.06) | 1.03 (0.99, 1.07) | 1.03 (0.99, 1.07) |
| Duration of OC use, y | | | | | | | |
| Never | 1,015,265 | 84,938 | 7,762,969.8 | 10.9 | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| <1 | 116,384 | 9,877 | 898,277.1 | 11.0 | 1.01 (0.98, 1.03) | 1.01 (0.99, 1.03) | 1.01 (0.99, 1.03) |
| ≥1 | 77,581 | 6,739 | 594,891.8 | 11.3 | 1.04 (1.01, 1.06) | 1.03 (1.00, 1.02) | 1.03 (1.00, 1.05) |
| Unknown | 62,885 | 5,341 | 479,994.8 | 11.1 | 1.02 (0.99, 1.05) | 1.01 (0.98, 1.04) | 1.01 (0.98, 1.04) |

Model 1: crude model

Model 2: the full model included age, age at menarche, age at menopause, parity, duration of breast feeding, duration of HT, duration of OC use, alcohol consumption, smoking, regular exercise, income, body mass index, hypertension, diabetes, dyslipidemia, and cancer.

eTable 5. Hazard Ratios and 95% Confidence Intervals of Fractures

| Variables | Any fracture | Vertebral fracture | Hip fracture | Other fracture |
|-------------------------|-------------------|--------------------|-------------------|-------------------|
| Income (quartile) | | | | |
| Q1 (lowest) | 1 (ref.) | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| Q2 | 1.00 (0.98, 1.01) | 1.00 (0.97, 1.02) | 1.04 (0.98, 1.11) | 0.99 (0.97, 1.01) |
| Q3 | 1.01 (0.99, 1.02) | 1.03 (1.01, 1.06) | 1.03 (0.97,1.09) | 1.00 (0.98, 1.02) |
| Q4 (highest) | 1.00 (0.98, 1.01) | 0.99 (0.97, 1.01) | 1.01 (0.96, 1.06) | 1.01 (0.99, 1.02) |
| Smoking status | | | | |
| Never | 1 (ref.) | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| Ex-smoker | 0.97 (0.93, 1.02) | 0.93 (0.86, 1.01) | 1.16 (0.98, 1.38) | 0.94 (0.89, 1.00) |
| Current smoker | 1.06 (1.03, 1.09) | 1.13 (1.08, 1.18) | 1.37 (1.24, 1.52) | 0.98 (0.94, 1.02) |
| Alcohol consumption | | | | |
| None | 1 (ref.) | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| Mild | 1.05 (1.04,1.07) | 0.98 (0.96, 1.01) | 0.91 (0.84, 0.98) | 1.08 (1.06, 1.10) |
| Heavy | 1.26 (1.19, 1.35) | 1.18 (1.05, 1.32) | 1.58 (1.19, 2.08) | 1.26 (1.17, 1.36) |
| Regular exercise | 0.98 (0.97, 0.99) | 0.90 (0.88, 0.92) | 0.92 (0.87, 0.97) | 1.05 (1.03, 1.06) |
| Body mass index (kg/m²) | | | | |
| <18.5 | 1.02 (0.98, 1.05) | 1.02 (0.98, 1.07) | 1.34 (1.21, 1.48) | 0.94 (0.90, 0.99) |
| 18.5 to <23 | 1 (ref.) | 1 (ref.) | 1 (ref.) | 1 (ref.) |
| 23 to <25 | 1.00 (0.99, 1.01) | 1.04 (1.02, 1.06) | 0.91 (0.86, 0.95) | 1.00 (0.99, 1.02) |
| 25 to <30 | 0.98 (0.97, 0.99) | 1.06 (1.05, 1.08) | 0.89 (0.85, 0.93) | 0.96 (0.94, 0.97) |
| ≥30 | 0.94 (0.92, 0.96) | 1.08 (1.04, 1.12) | 0.96 (0.87, 1.05) | 0.87 (0.85, 0.90) |
| Comorbidities | | | | |
| Hypertension | 0.97 (0.96, 0.98) | 0.96 (0.95, 0.98) | 1.18 (1.14, 1.23) | 0.96 (0.95, 0.98) |
| Diabetes mellitus | 1.05 (1.03, 1.06) | 0.86 (0.85, 0.88) | 1.71 (1.64, 1.79) | 1.12 (1.10, 1.14) |
| Dyslipidemia | 0.98 (0.97, 0.99) | 0.97 (0.95, 0.98) | 0.99 (0.95,1.03) | 1.00 (0.99, 1.02) |
| Cancer | 0.97 (0.95, 1.00) | 0.92 (0.87, 0.96) | 0.97 (0.86, 1.10) | 1.02 (0.98, 1.06) |

The full model included age, age at menarche, age at menopause, parity, duration of breast feeding, duration of HT, duration of OC use, alcohol consumption, smoking, regular exercise, income, body mass index, hypertension, diabetes, dyslipidemia, and cancer.

eFigure. Flow Chart of the Study Population

