

Cumulative incidence and changes in the prevalence of vertebral fractures in a rural Japanese community: a 10-year follow-up of the Miyama cohort

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Received: 28 April 2006 / Accepted: 17 July 2006 / Published online: 27 September 2006
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Abstract

Introduction The objective of this study was to clarify secular changes in the prevalence and cumulative incidence of vertebral fractures (VFX) among the general inhabitants of Miyama, a rural Japanese community.

Methods A cohort of 1,543 inhabitants aged 40–79 years was established using resident registration in 1989. Of these, 400 participants were selected and divided into four age strata, each containing 50 men and 50 women. Participants completed a self-administered questionnaire and anthropometric measurements were taken. In 1990, baseline radiographic examinations of the thoracolumbar spine in anteroposterior and lateral views were performed. Radiographic examinations of the same sites were performed on the same participants in 2000. All VFX were

diagnosed by the same experienced orthopedic surgeon. Evaluation of radiographic surveys after 10 years was completed for 287 of the 400 participants (132 men, 155 women; 71.8%).

Results The prevalences of VFX at follow-up in 2000 for subjects in their 50s, 60s, 70s, and 80s were 2.9%, 10.3%, 13.2%, and 25.0% for men, and 2.1%, 9.1%, 20.5%, and 54.2% for women, respectively. Prevalence in the present study was compared with that collected in 1990, showing improvements over time for men and women in their 50s, 60s, and 70s. Future problems with osteoporotic fractures in Japan might, thus, be less severe than previously predicted. Cumulative incidence over 10 years was detected by the number of incident cases divided by the number of all participants in the initial study. Incident cases were defined by two criteria. First, cases with prevalent VFX were excluded from both the numerators and the denominators. In this analysis, the cumulative incidence of the cases with first VFX was detected. Secondly, the cumulative incidence including the subjects who had suffered from the prevalent fractures and with an increased number of VFX in the following 10 years were calculated. This analysis clarified the cumulative incidence of cases with and without previous VFX. Incidence over 10 years of subjects in their 40s, 50s, 60s, and 70s with first VFX were 2.2%, 4.9%, 5.1%, and 10.8% for men, and 2.1%, 4.5%, 14.0%, and 22.2% for women, respectively. By contrast, if new cases were defined as including subjects who had suffered from previous fractures, but displayed an increased number of VFX, cumulative incidences for subjects in their 40s, 50s, 60s, and 70s were 2.1%, 8.3%, 10.0%, and 12.2% for men, and 2.1%, 6.1%, 18.0%, and 22.4% for women, respectively.

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Keywords Cohort study · Cumulative incidence · Japanese · Prevalence · Vertebral fractures

Introduction

Among fractures due to osteoporosis, vertebral fractures (VFX) are the most likely to lead to marked health problems. VFX are reportedly associated with functional impairment [1], back pain [2, 3], kyphosis [2, 3], esophageal reflux [4], depressive mood [5], respiratory dysfunctions [6], and decreased quality of life [7, 8]. In addition, VFX increase the risk of new VFX and hip fractures [9–13]. Furthermore, prevalent VFX reportedly increase mortality risk [14–16].

Although preventive strategies for VFX are urgently needed with the rapid aging of populations such as in Japan, little data is available concerning epidemiological characteristics in population samples, particularly in Japan [17–19]. A population-based cohort was established in 1990, comprising subjects aged 40- to 79-years-old from resident registrations for a village in a mountainous area, Miyama village, and follow-up studies were conducted in 1993, 1997, and 2000 [20–22]. In the previous report of the Miyama study, prevalences of VFX were ascertained for men and women [23].

In the present study, radiographic examination was performed on the same subjects after 10 years to assess secular trends of prevalence and to clarify cumulative incidences among different generations of VFX in a rural Japanese community.

Participants and methods

Epidemiological research based on populations has been conducted in Miyama village, Wakayama prefecture, Japan. As a detailed profile of the Miyama cohort has already been described elsewhere [20–23], a brief summary is provided here. In this village, a list of inhabitants aged 40- to 79-years-old was compiled based on resident registrations as of the end of December 1988. As a result, a cohort of 1,543 inhabitants (716 men, 827 women) was identified. A self-administered questionnaire survey regarding 125 daily activities such as smoking habits, alcohol consumption, and physical exercise was performed (baseline study).

From the above-mentioned cohort, 50 men and 50 women in each decade of age between 40 and 79 years (for a total of 400 participants) was recruited based on resident registrations for the purpose of measuring bone mineral density (BMD) in 1990 (initial BMD survey). In addition to BMD measurements using dual energy X-ray absorptiometry (DXA; Lunar DPX, GE Medical Systems, Madison, WI, USA), an interviewer-administered questionnaire, medical examination, and physical measurements were also performed for these participants. The questionnaire comprised 70 items, such as past history, history of

medication, smoking habits, alcohol consumption, calcium intake, physical exercise, history of childbirth, history of lactation, and menstrual status. Radiographic examination of the spine was performed on all participants in 1990. Anteroposterior and lateral images of thoracolumbar vertebrae Th5-L5 were used for diagnosis (initial X-ray survey). Radiographic examination was again performed on subjects who provided consent after 10 years. Lateral images of thoracolumbar vertebrae Th5-L5 were again used for diagnosis (2nd X-ray survey).

Lateral spinal radiographs were examined for the presence of one or more VFX between Th5-L5, using the criteria defined by the Japan Bone and Mineral Society (Fig. 1) [24]. According to these criteria, the measurement of anterior, middle, and posterior heights on lateral radiography of the thoracic and lumbar spine is required, in order to determine ratios defining the anterior wedge, biconcave, and compound dimensions of the vertebral bodies. Diagnosis of VFX on all radiographs was performed by the same experienced orthopedic doctor (HK).

Prevalence and cumulative incidence were calculated by utilizing the results of diagnosis. Cumulative incidence is defined simply as the ratio of incident cases to the at-risk population at the beginning of the observation period. In the present study, cumulative incidence over 10 years was, thus, detected as the number of incident cases divided by the number of all participants in the initial study. Incident cases were defined using two criteria. First, cases with previous VFX were excluded from both numerators and denominators. In this analysis, the cumulative incidence of cases with first VFX was detected. Secondly, the cumulative incidence including subjects who had suffered from previous VFX but who displayed an increased number of VFX over the following 10 years was calculated. This analysis clarified the cumulative incidence of cases with and without previous VFX.

In the above-mentioned formula to obtain the cumulative incidence, the denominator was defined as the total number of participants in the initial X-ray survey. Using this definition, drop-outs during the observed 10 years were considered as non-VFX. To avoid underestimation caused

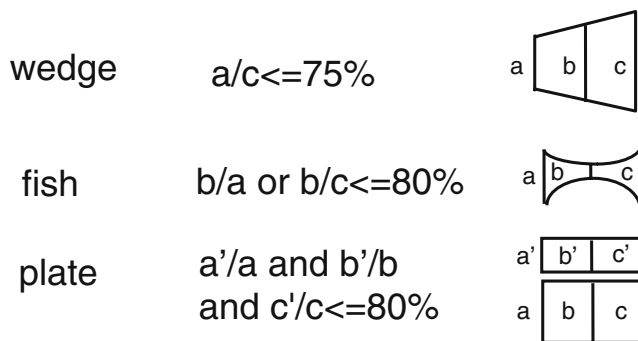


Fig. 1 Diagnostic criteria for vertebral fractures in Japan

by the inclusion of drop-outs, another cumulative incidence was calculated by excluding drop-outs during the study period from the denominator. This index was defined as the cumulative incidence among survivors.

All statistical analyses were performed using STATA statistical software (STATA Corp., College Station, TX, USA).

Results

We note our findings as follows:

1. Eligible participants

Among 400 participants of the initial BMD examination, 390 provided consent to participate in the radiographic examination (194 men, 196 women; 97.5%) of the initial survey. Radiographic evaluation after 10 years was completed for 287 of the 400 participants (132 men, 155 women; 71.8%). The loss of 101 participants was explained as follows: 55 participants died (37 men, 18 women); 16 participants moved (8 men, 8 women); 13 participants were ill (4 men, 9 women); 8 participants

were busy (8 men); 5 participants refused to participate further (5 men); and 4 participants were away from the area at the time of follow-up (1 man, 3 women). Another 12 participants (5 men, 7 women) were excluded from analysis due to the poor quality of radiographic images. Table 1 shows the physical characteristics of the remaining participants at the time of the initial X-ray survey.

2. Prevalence of VFX at the survey in 2000 and secular trends 1990–2000

The prevalences of VFX at follow-up in 2000 were calculated. The prevalences of VFX for subjects in their 50s, 60s, 70s, and 80s were 2.9%, 10.3%, 13.2%, and 25.0% for men, and 2.1%, 9.1%, 20.5%, and 54.2% for women, respectively.

Figure 2 shows a comparison of VFX prevalences in the same age group according to the results of radiographic examination in 1990. The prevalences were significantly decreased between the 50s to 70s age groups in 2000 compared to 1990 ($P < 0.05$) (Fig. 2).

3. Cumulative incidence of VFX

The cumulative incidences over 10 years for men and women sorted by the presence or absence of VFX at the

Table 1 Characteristics of participants of the initial survey, classified by sex and age stratum

Birth cohort	Age strata of participants	No. of participants	No. of participants	Anthropometric measurements at the initial survey (mean (SD))				BMD measurements on initial survey (mean (SD))	
				Age (yrs)	Height (cm)	Weight (kg)	BMI (kg/m ²)	BMD (L2-4)	BMD (femoral neck)
Men	Total	194	132	56.9 (10.3)	161.3 (6.7)	58.2 (9.2)	22.3 (2.5)	1.10 (0.19)	0.88 (0.14)
	1940–49	47	35	44.1 (3.2)	166.3 (5.8)	63.8 (8.4)	23.0 (2.3)	1.18 (0.17)	0.97 (0.15)
	1930–39	48	39	53.9 (2.6) ^a	162.0 (5.8) ^a	60.1 (8.2)	22.8 (2.4)	1.16 (0.18)	0.91 (0.12)
	1920–29	50	38	63.2 (2.8) ^{ab}	159.4 (5.4) ^a	56.1 (7.5) ^a	22.0 (2.4)	1.03 (0.19) ^{ab}	0.82 (0.12) ^{ab}
	1910–19	49	20	73.3 (2.8) ^{abc}	154.7 (4.8) ^{abc}	48.7 (6.8) ^{abc}	20.3 (2.5) ^{ab}	1.01 (0.16) ^{ab}	0.79 (0.10) ^{ab}
Women	Total	196	155	56.7 (10.1)	149.0 (6.0)	49.7 (8.1)	22.3 (2.8)	0.98 (0.23)	0.76 (0.14)
	1940–49	48	48	44.6 (3.0)	152.6 (4.7)	53.6 (8.3)	22.9 (2.8)	1.18 (0.16)	0.88 (0.12)
	1930–39	49	44	54.8 (2.7) ^a	149.9 (5.1)	50.1 (7.1)	22.3 (2.6)	0.99 (0.18) ^a	0.75 (0.12) ^a
	1920–29	50	39	64.3 (2.8) ^{ab}	147.4 (5.1) ^a	47.5 (6.9) ^a	21.9 (3.0)	0.85 (0.20) ^{ab}	0.69 (0.11) ^a
	1910–19	49	24	71.6 (1.8) ^{abc}	142.5 (5.6) ^{abc}	44.5 (7.6) ^{ab}	21.9 (3.0)	0.78 (0.16) ^{ab}	0.64 (0.09) ^{ab}

^a Significantly different from values of the birth cohort group born in 1940–49

^b Significantly different from values of the birth cohort group born in 1930–39

^c Significantly different from values of the birth cohort group born in 1920–29

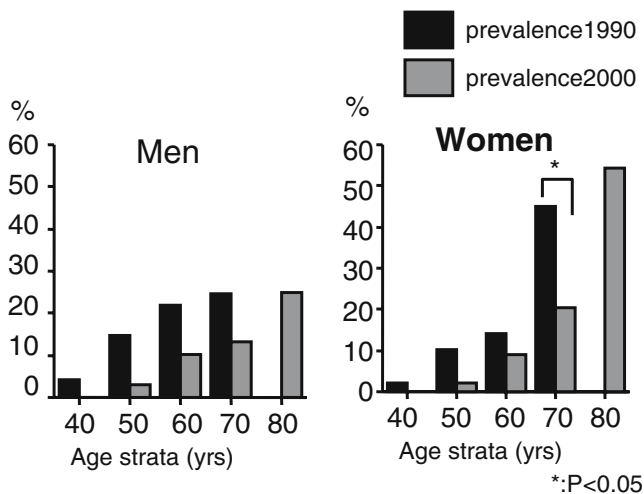


Fig. 2 Comparison of prevalence for vertebral fractures in the same age groups among participants in the X-ray examinations performed in 1990 and 2000

initial X-ray survey are shown in Table 2. The cumulative incidence of VFX including subjects with previous VFX after 10 years increased with age in both men and women.

After excluding cases with previous VFX, the cumulative incidence of first VFX for participants in their 40s, 50s, 60s, and 70s increased with age in both men and women, and was higher in women than in men in all age strata, except the 40s.

Although no association was identified between the cumulative incidence among participants with previous VFX and age in both men and women, cumulative incidence was significantly higher for subjects with previous VFX than for subjects without previous VFX in both men and women ($P < 0.05$).

4. Cumulative incidence of VFX among survivors

The cumulative incidences of VFX among cases with first VFX for survivors in their 40s, 50s, 60s, and 70s were 2.9%, 2.8%, 8.6%, and 21.1% for men, and 2.1%, 7.0%, 18.9%, and 31.3% for women, respectively. The cumulative incidence of the first VFX among survivors increased with age in both men and women, and was higher in women than in men in all age strata, except in the 40s.

The cumulative incidences of VFX including subjects who had suffered from previous fractures or with increased numbers of VFX after 10 years for survivors in their 40s, 50s, 60s, and 70s were 2.9%, 10.3%, 13.2%, and 30.0% for men, and 2.1%, 6.8%, 23.1%, and 45.8% for women, respectively, suggesting an increase with age. Cumulative incidence was, again, significantly higher for subjects with previous VFX than for those without previous VFX in both men and women ($P < 0.01$).

Discussion

The present study evaluated the prevalence and cumulative incidence of VFX over 10 years using radiographic diagnosis after the performance of two radiographic examinations of the thoracolumbar spine in anteroposterior and lateral views of population-based subjects.

Compared with hip fractures, epidemiological studies of VFX have been less available. One reason for this is the fact that VFX are not always symptomatic, and longitudinal

Table 2 Cumulative incidences over 10 years for men and women classified by the presence or absence of VFX at the initial survey

Birth cohort	Age strata at the initial survey (yrs)	All participants			With presence of VFX at the initial survey			With absence of VFX at the initial survey		
		No. of new VFX 1990–2000	No. of participants initial X-ray survey 1990	Cumulative incidence (%)	No. of new VFX cases 1990–2000	No. of participants initial survey 1990	Cumulative incidence (%)	No. of new VFX cases 1990–2000	No. of participants initial survey 1990	Cumulative incidence (%)
Men	Total	16	194	8.2	7	32	21.9	9	162	5.6
	1940–49	1	47	2.1	0	2	0.0	1	45	2.2
	1930–39	4	48	8.3	2	7	28.6	2	41	4.9
	1920–29	5	50	10.0	3	11	27.3	2	39	5.1
	1910–19	6	49	12.2	2	12	16.7	4	37	10.8
Women	Total	24	196	12.2	9	35	25.7	16	161	9.9
	1940–49	1	48	2.1	0	1	0.0	1	47	2.1
	1930–39	3	49	6.1	1	5	20.0	2	44	4.5
	1920–29	9	50	18.0	3	7	42.9	6	43	14.0
	1910–19	11	49	22.4	5	22	22.7	6	27	22.2

population surveys are required to clarify epidemiological indices. The present study used subjects representative of the whole town, and careful follow-up was performed. One limitation was the use of a relatively small number of fractures and small study population to determine the prevalence and cumulative incidence of VFX. However, few such longitudinal observational studies with high participation rates have been reported, representing the principal strength of this study.

Another reason for the scarcity of data on this subject is the lack consensus regarding the definition of VFX [25]. The present study utilized the diagnostic criteria of the Japan Bone and Mineral Society, as these criteria are the most popular and most commonly used in various clinical fields in Japan. These criteria seem to use the kind of semiquantitative (SQ) method advocated by Genant et al. [26]. From the perspective of grading in SQ methods, the criteria of the Japan Bone and Mineral Society correspond to grades above SD2. The epidemiological indices in the present study, thus, did not include cases classifiable as SD1, suggesting that the results obtained using Japanese criteria might represent an underestimation compared to other results diagnosed using SQ methods.

Population-based prevalence studies of VFX have been performed in Western countries, such as the USA [27], Denmark [28, 29], Finland [30], Sweden [31, 32], the Netherlands [33], England [34, 35], and France [36, 37], and in a multicenter study involving 19 European countries [38]. In Asia, the prevalence of VFX has been reported in Taiwan [39], China [40], and Japan [18, 19, 24]. Cummings et al. compared the results of population-based studies of radiological VFX prevalence in women ≥ 50 -years-old between Europe, Minnesota (USA), Hawaii (USA), Hiroshima (Japan), Taiwan, and Beijing (China) [41]. In this comparison, the prevalence of VFX among women >70 -years-old was higher in Japan than in the USA or China, and was as high as that in Europe. The prevalence of VFX in the mountainous area of the present study was much higher than that in Hiroshima, an urban area, suggesting that regional differences in VFX might be present in Japan.

The present study clarified differences in the prevalence of VFX for given age strata between birth cohorts. The comparison of data collected in 1990 and 2000 showed improvements over time for men and women in their 50s, 60s, and 70s. A previous report has already noted that BMD data from 1990 and 2000 showed significant improvements for men in their 60s and for women in their 50s [23], concluding that this might predict a decrease in the prevalence of osteoporosis in the near future in Japan. The present results suggest that the prevalence of not only osteoporosis, but also osteoporotic fractures, could decrease in the future in Japan. Fujiwara et al. assessed the effects of birth cohort on the incidence of VFX in Hiroshima, and

reported that incidence decreased by a factor of 0.5 in men and 0.6 in women with each succeeding birth decade [17]. The present data are consistent with these findings.

This low prevalence of VFX in more recent birth cohorts may reflect nutritional improvements, in addition to BMD improvements. As noted elsewhere [23], nationwide nutritional surveys have reported the mean calcium intake as 253 mg/day in 1946 (1st survey), 338 mg/day in 1955, 465 mg/day in 1965, and 552 mg/day in 1975 [42], representing a dramatic increase. These nutritional improvements might be expected to increase body build and BMD, and, thus, decrease the prevalence of VFX in all age groups. Another possible reason for the decreasing prevalence of VFX might be the effects of action taken for the prevention of osteoporosis by the government and academic societies. To achieve the prevention of osteoporosis, the Ministry of Health, Labour and Welfare recommended all that communities start BMD screening for middle-aged and elderly female residents in 1995. These examinations in various communities have enabled many women to become more aware of osteoporosis. Moreover, the Japan Osteoporosis Foundation and the Japan Osteoporosis Society, which were established in 1991 and 1999, respectively, have been conducting public education campaigns to inform the general population about osteoporosis and risk factors. These efforts by both the government and academic societies might have resulted in the decreasing prevalence of VFX.

Regarding the incidence of VFX, while some epidemiological studies have been reported from Western countries [43–45], the only investigation from Japan was reported by Fujiwara et al. [18]. Fujiwara et al. performed radiographic examinations in 1994–1995, with follow-up at an average of 4 years, and clarified the incidence of VFX in their Hiroshima cohort. Incidences of VFX in this Hiroshima cohort increased with age, particularly after about 60 years of age, and women displayed a nearly two-fold greater incidence of VFX than men. The incidence of VFX was much higher in individuals with previous VFX than in those without previous VFX. As cumulative incidence was clarified in the present study, rather than incidence, direct comparison of the present results with those of Fujiwara et al. might be incorrect. However, findings in the present study, such as age-dependency, sex differences, and higher incidence in individuals with previous VFX than in those without previous VFX, are consistent with the results of the previous investigation in Hiroshima.

The cumulative incidence for participants at the initial survey in the present study could be considered as a 10-year probability according to age. In this mountain village, 12.2% of men and 22.4% of women in their 70s will suffer new morphometric VFX in the next 10 years. Kanis et al. reported the 10-year probability of clinical VFX in female

participants in the European Prospective Osteoporosis Study [46]. The 10-year probability of clinical VFX among women with a 0 Z-score of amplitude-dependent speed of sound on quantitative ultrasound (QUS) measurements was 3.9% at 70 years old, 4.5% at 75 years old, and 4.9% at 80 years old. Although only about one-third of morphometric fractures reportedly come to medical attention [43], the present results indicated much higher incidences than those described by Kanis et al. Although the future incidence of VFX in the Japanese population could be predicted to decrease, elderly individuals seem to remain at high risk. To reduce the incidence and prevalence of osteoporosis and osteoporotic fractures, further epidemiological research is needed to facilitate the development of affordable strategies for the early identification of high-risk individuals.

Acknowledgments This work was supported by Grants-in-Aid for Scientific Research C16590512 from the Ministry of Education, Science, Sports, and Culture in Japan, H16-Chihou Kossetsu-021, H17-Meneki-009 from the Ministry of Health, Labour and Welfare, and the grant from the Japan Osteoporosis Society.

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