Epidemiology of Hemorrhagic Fever with Renal Syndrome in Korea, 2001-2010

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Hantavirus infection induces 2 different diseases, hemorrhagic fever with renal syndrome (HFRS) and hantavirus pulmonary syndrome (HPS), mainly in Korea, far east Russia, and China and in the United States, respectively (1, 2). The viruses are transmitted to humans by the inhalation of excreta of rodents infected with hantaviruses (3). HFRS was first recognized during the Korean War in the early 1950s, and although a hantavirus vaccine has been developed, the disease remains a serious problem in Asia and Europe (1, 4), and especially among soldiers in Korea (5, 6). A 2006 report describes HFRS epidemiology in endemic regions in Korea during a relatively short period, 1995-1998 (7); however, an updated epidemiological study is necessary to better understand the current status of HFRS in Korea. This study provides a comprehensive epidemiological overview of HFRS during the past decade in Korea.

Raw data were utilized from the National Notifiable Disease Surveillance System (NNDSS) website of the Korean Center for Disease Control and Prevention (KCDC) (2001-2010) (8), and “HFRS,” Incidence for Communicable Diseases (Category III), Yearbook of Health and Welfare Statistics by the Ministry of Health and Welfare, Republic of Korea (MHW, ROK), 2001-2010 (9). During the period 2001-2010, there were 3,953 patients and 40 fatal cases of HFRS (Table 1). The prevalence rate (PR) of HFRS per 100,000 population and case fatality rate (CFR) were estimated by the criteria established by the World Health Organization (WHO), and the upper and lower limits of the 95% confidence interval (CI) were calculated. Statistically significant differences between the epidemiological aspects were determined at P < 0.05 and P < 0.01 using the chi-square test or the paired t-test. The data analyses were carried out using the statistical system software included in Microsoft Excel 2007.

As shown in Table 1, the HFRS cases were analyzed according to the PR, CFR, gender, age, and occupation of the infected individuals. There were a total of 3,953 HFRS patients with an average PR of 0.81 (95% CI, 0.79-0.85) per 100,000 population and a total of 40 fatal cases with a CFR of 1.01% (95% CI, 0.7-1.3). A significantly higher number (P < 0.001) of men were infected than women (57% vs 43%). The CFR was also significantly higher (P < 0.05) in men than in women (1.3% vs 0.7%). The distribution of HFRS cases by age clearly showed a significantly higher incidence of HFRS in people older than 40 yr (P < 0.001). When the HFRS patients were analyzed by occupation, significantly higher (P < 0.001) number of infections occurred among farmers (35.6%) than among those in other job groups (38.7%).

HFRS cases were analyzed by season, geography, and residential area of individuals. The PR of HFRS in spring, summer, autumn, and winter was 7.0%, 9.1%, 61.4%, and 22.5%, respectively. The PR of HFRS in autumn was significantly higher than that in other seasons (P < 0.001). The HFRS cases predominantly occurred during the last quarter of the calendar year (October, November, and December). A significantly higher number of HFRS cases (P < 0.001) was reported in the western part (provinces of Gyeonggi, Chungnam, Jeonbuk, Jeonnam, and Jeju and major cities of Seoul, Incheon, Daejeon, and Gwangju, 68.9%)
than in the eastern part (provinces of Gangwon, Chungbuk, Gyeongbuk, Gyeongnam and major cities of Daegu, Ulsan, and Busan, 31.1%) of Korea (Fig. 1). Finally, HFRS cases were significantly higher ($P < 0.001)$ in rural areas (80.3%) than in urban areas (19.7%).

This study provides a more updated HFRS prevalence status in the whole population of Korea during the past decade. A previous report addressed annual prevalence and incidence rates of HFRS during the late 1990s only in endemic areas near the demilitarized zone (DMZ) between South and North Korea (7). The prevalence rate per 100,000 population of HFRS determined in this study was 0.81. In contrast, the annual prevalence of HFRS determined in the previous study was 40-64 per 100,000 military population in the DMZ areas. Therefore, the prevalence of HFRS could be seriously affected by the regions and populations examined.

This study confirmed that the incidence of HFRS significantly increased in the autumn. In the previous report, an almost identical pattern of HFRS incidence was identified in soldiers in endemic areas (7). Higher incidence of HFRS in the autumn, versus other seasons, could be attributed to the high numbers of hantavirus-infected rodents in Korea during this time (10, 11). Seroprevalence of hantaviruses in wild rodents in Korea confidently demonstrated that most wild rodent hantavirus infections occur in the autumn. Therefore, increased outdoor activity of people in the autumn, especially in rural areas, would increase the probability of hantavirus infection, leading to a higher incidence of HFRS in the autumn among farmers.

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### Table 1. Prevalence rate, case fatality rate, and other aspects of HFRS

| Parameters                  | No. of cases (%) | $P$-value* | 95% CI          |
|-----------------------------|------------------|------------|-----------------|
| Total No. of patients       | 3,953            |            |                 |
| Prevalence rate/100,000     | 0.81             | 0.79-0.85  |                 |
| Total No. of death          | 40               |            |                 |
| Case-fatality rate in %     | 1.01             | 0.7-1.3    |                 |
| Gender of patients          |                  |            |                 |
| Male                        | 2,252 (57.0)     | < 0.001    | 55.5-58.5       |
| Female                      | 1,701 (43.0)     |            | 41.5-44.5       |
| Total                       | 3,953            |            |                 |
| CFR (%) by sex             |                  |            |                 |
| Male                        | 29/2,252 (1.3)   | < 0.05     |                 |
| Female                      | 11/1,701 (0.7)   |            |                 |
| Age (yr)                    |                  |            |                 |
| < 39                        | 708 (17.9)       |            | 16.7-19.1       |
| > 40                        | 3,245 (82.1)     | < 0.001    | 80.9-83.3       |
| Total                       | 3,953            |            |                 |
| Occupation                  |                  |            |                 |
| Farmers                     | 1,407 (35.6)     | < 0.001    | 34.1-37.1       |
| Mistresses                  | 384 (8.8)        |            | 7.9-8.4         |
| White-collar jobs           | 301 (7.6)        |            | 6.8-8.4         |
| Blue-collar jobs            | 222 (5.6)        |            | 4.9-6.3         |
| Soldiers                    | 48 (1.2)         |            | 0.9-1.5         |
| Students                    | 98 (2.5)         |            | 2.0-3.0         |
| Other jobs                  | 1,529 (38.7)     |            |                 |
| Total                       | 3,953            |            |                 |

95% CI, Confidence interval of 95% of the rate; CFR, Case fatality rate.

*Chi-square analysis indicated a significant difference from the total value.

In this study, the HFRS patient numbers and CFR were significantly higher in men than in women. However, a recent study conducted in China reported that the incidence of HFRS is greater in men, but the CFR is higher in women than in men (12). The discrepancy of CFRs between Korean and Chinese HFRS patients is an interesting question that requires more thorough epidemiological studies to resolve. In addition, many more HFRS patients were identified in people older than 40 yr and in the western part of Korea. This could be partially explained by a larger population in the western part of Korea and the advanced ages of farmers.

In conclusion, HFRS still occurs commonly among men, in autumn, and in western rural area of Korea. These comprehensive epidemiological data would be valuable for developing HFRS control strategies in Korea.

### DISCLOSURE

The authors have no conflicts of interest to disclose.

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