

Hydrochemical anomaly of drinking waters in some endemic Kashin-Beck disease areas of Tibet, China

Qinghai Guo · Yanxin Wang

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Abstract Kashin-Beck disease (KBD) has been identified in some areas of Tibet, China. In this work, the Sangri, Nimu, Xietongmen and Gongbujiangda counties of Tibet were selected as case study areas to understand the relationship between KBD occurrence and chemical composition of drinking water. 30 drinking water samples were collected in the KBD-affected and KBD-unaffected villages of these four counties, and the hydrochemistry of endemic and non-endemic samples was compared. The results show that $\text{HCO}_3\text{-Ca}$, $\text{HCO}_3\text{-Ca-Na}$, and $\text{HCO}_3\text{-SO}_4\text{-Ca}$ are the major hydrochemical types of water samples from both KBD-affected and KBD-unaffected villages. Although Se deficiency in environment has been widely regarded as an important cause of KBD, the Se concentrations in the drinking water samples do not correlate with KBD occurrence in the study areas. However, there are significant differences between the concentrations of Al, Fe, Na, Mn, Cd, Co, Cu, Ba and Mo in the endemic drinking water samples and those in the non-endemic samples, indicating that these constituents may be related to the prevalence of KBD in the study areas.

Keywords Kashin-Beck disease · Hydrochemistry · Drinking water · Tibet

Introduction

Kashin-Beck disease (KBD), also known as “Big Bone Disease”, is a disorder affecting the bones and joints of the hands (including fingers), elbows, knees, and ankles of children and adolescents. Individuals stricken with the disease develop chronically stiff or deformed joints, and even short limbs and statures due to necrosis of the growth plates of bones and those of joint cartilage. Endemic KBD occurs mostly in China, southeastern Siberia and North Korea (Sokoloff 1989). China is the most seriously afflicted country in the world. It was reported that there are 13 provinces and 2 autonomous regions with prevalent KBD in China (Tan et al. 2002). The KBD patients in China are mainly farmers who live in remote and rural areas. According to the statistics of 1999, there are about 40 million people living in the KBD areas of China, and the total population of KBD patients is more than 1 million (Yang et al. 2006).

From the 1850s to the 1950s, Russian researchers investigated the prevalence of KBD in Russia (Hinsenkamp 2001). In China, the relevant research studies have been carried out since the 1950s. These studies focused mainly on environmental etiology of KBD, with an emphasis on the relationship between KBD prevalence and selenium levels in eco-environments or human body (Li et al. 1982, 1992; Deng et al. 1999; Guo et al. 1999; Xu et al. 2000; Zhang et al. 2001; Wang et al. 2003; Fan 2005), organic matter concentration in drinking waters (Wu et al. 1987; Peng and Xu 1988; Wang and Feng 1989; Zhang et al. 1990; Wang et al. 1991a, b) and severe contamination of grains by fungi-produced mycotoxin (Bi et al. 2001).

Due to the geographic distribution of the KBD-prevalent areas, there are few studies concerning KBD conducted by western researchers before the 1990s. Since the late 1990s,

Q. Guo (✉) · Y. Wang
State Key Laboratory of Biogeology and Environmental
Geology, School of Environmental Studies,
China University of Geosciences, Wuhan 430074,
Hubei, People's Republic of China
e-mail: qhguo2006@gmail.com

a group of Belgian researchers made a series of investigations on KBD in Tibet, China. Mathieu et al. (1997, 2001a, b) and Hinsenkamp et al. (2001) carried out clinical–radiological studies on the identification of KBD in Lhasa, Shannan, and Rikaze of Tibet. Pasteels et al. (2001) and Suetens et al. (2001) made histopathological studies of the KBD sufferers in Lhasa of Tibet, and considered that KBD is induced by the peroxidation of chondrocytes as the antioxidant system of human body is impaired. Moreno-Reyes et al. (1998, 2001, 2003) investigated the relationship among the serum selenium concentration, thyroid function, and KBD status in children with ages between 5 and 15 living in the villages around Lhasa, and evaluated the effects of selenium and iodine supplements on KBD control. Chasseur et al. (1997, 2001) and Haubruge et al. (2001, 2003) analyzed the impact of food crop contamination by fungi-produced mycotoxin on the KBD prevalence in Lhasa and Rikaze, and accordingly recommended some prevention and control measures for KBD.

From the above overview of previous work, one notes that much effort has been made to explain the prevalence of KBD in terms of environmental etiology and pathology. However, there are so far only a few systematic studies that have focused on the relationship between KBD occurrence and drinking water chemical composition. The aims of this study are: (1) to investigate the hydrochemistry of major and trace elements in drinking waters in four counties of Tibet as study areas where KBD is prevalent; and (2) to delineate any hydrochemical anomalies of the water samples collected from the study areas. This study will not only be a reference for future etiological studies of KBD, but help to guide the prevention and control of KBD in Tibet and other KBD-afflicted areas.

KBD in Tibet

Tibet is where KBD disease is the most widely distributed, serious and active in China (Institute of Kashin-Beck Disease of Harbin Medical University 2000), and there have been reports of KBD cases in all seven of its administrative divisions (Lhasa, Changdu, Linzhi, Shannan, Rikaze, Naqu and Ali). A survey made in the 1980s indicates that there were 13 KBD counties in Tibet (Wang et al. 1985) (Fig. 1). However, 20 new KBD counties were discovered during an investigation made in the late 1990s (Yang et al. 2003) (Fig. 1). Altogether, KBD disease was present in 379 KBD villages distributed in the above 33 counties (Gong et al. 2004a, b), and the number of individuals diagnosed with KBD was 18,000, equivalent to 0.69% of the total population of Tibet (Zaxi et al. 2003). The number of villages with KBD incidence rates greater than 40% reached 113 and accounted for 30% of the total

of KBD villages (Zaxi et al. 2007). Moreover, children and adolescents who were less than 25-year old accounted for 50% of the total KBD patients in Tibet (Gong et al. 2004a). The most serious KBD symptoms, such as shortened limbs and statures, were observed in children 5–10 years of age (Institute of Kashin-Beck Disease of Harbin Medical University 2000).

Sampling and analysis

The study areas selected for this investigation were the Sangri, Nimu, Xietongmen, and Gongbujiangda counties of Tibet. A total of 30 water samples were collected from springs, rivers, and streams that serve as drinking water sources for local residents. In each county, several KBD-affected and KBD-unaffected villages were chosen for drinking water sample collection and analysis. The sampling locations of all water samples were listed in Table 1.

All water samples were filtered through 0.45 μm membranes on site. Samples were collected in 300 ml polyethylene bottles, which had been rinsed with deionized water twice before sampling. For cation analysis, reagent-grade HNO_3 with molar concentration up to 14 M was added to one sample collected at each site to bring the pH below 1. Unstable hydrochemical parameters including water temperature, pH, and electrical conductance (EC) were measured on site using a portable pH/EC/temperature meter. Alkalinity was measured using the Gran titration method (Appelo and Postma 1996). The concentrations of SO_4^{2-} , Cl^- , and NO_3^- were determined on an unacidified sample by Ion Chromatography. Se was analyzed by Atomic Fluorescence Spectrometry, and other chemical constituents by ICP-MS within 2 weeks of sampling. The HCO_3^- , CO_3^{2-} and CO_2 concentrations of all samples were calculated from the measured alkalinity and pH using PHREEQC. The hydrochemical properties and major ion concentrations of all 30 water samples are listed in Table 2, and the trace element concentrations in Table 3.

Results and discussion

Major constituents

The hydrochemical ‘type’ of all collected water samples was determined from the concentrations of seven major constituents (HCO_3^- , SO_4^{2-} , Cl^- , Ca^{2+} , Mg^{2+} , Na^+ , and K^+), and are listed in Table 1. $\text{HCO}_3\text{-Ca}$, $\text{HCO}_3\text{-Ca-Na}$, or $\text{HCO}_3\text{-SO}_4\text{-Ca}$ are the hydrochemical types of almost all water samples, except for samples GBJD04-A, XTM01 and XTM02 from the KBD-affected villages, which are $\text{SO}_4\text{-Ca}$ and $\text{HCO}_3\text{-Ca-Mg}$ type, respectively. In other

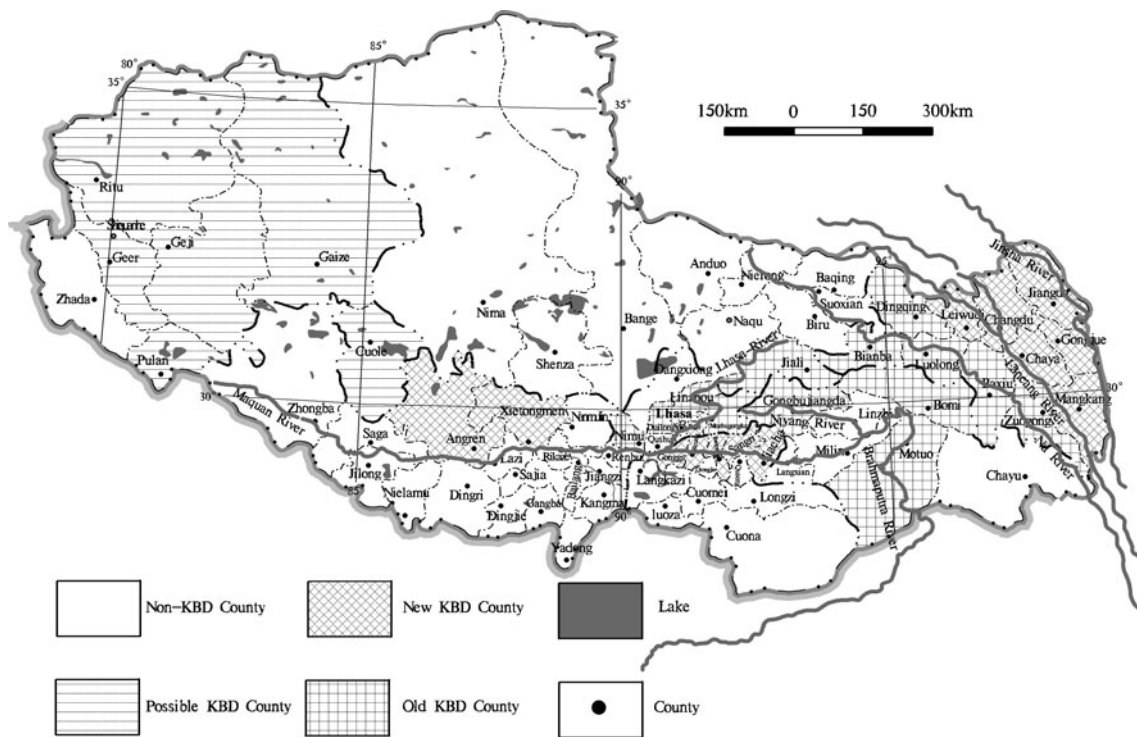


Fig. 1 Geographical distribution of the KBD counties in Tibet

Table 1 Sampling locations and hydrochemical types of water samples collected from the endemic and non-endemic villages of Sangri, Nimu, Xietongmen and Gongbujiangda counties

No.	Sampling location	Type of village	Longitude (°)	Latitude (°)	Elevation (m)	Hydrochemical type
SR01	Biba-III Village, Sangri County	Endemic area	92.034611	29.352056	3,871	HCO ₃ :SO ₄ -Ca
SR02	Biba-II Village, Sangri County	Endemic area	92.045111	29.319083	3,734	HCO ₃ :SO ₄ -Ca
SR03	Biba-I Village, Sangri County	Endemic area	92.041167	29.291361	3,628	HCO ₃ :SO ₄ -Ca
SR06	Zangga Village, Sangri County	Endemic area	92.210278	29.258611	3,581	HCO ₃ :SO ₄ -Ca
SR08	Qiamumai Village, Sangri County	Endemic area	92.327500	29.405833	3,946	HCO ₃ -Ca
SR09	Zengqi Village, Sangri County	Endemic area	92.364444	29.445000	4,105	HCO ₃ -Ca·Na
SR05	Jiangxiang-I Village, Sangri County	Non-endemic area	91.938861	29.280556	3,603	HCO ₃ -Ca
SR07	Quguosa Village, Sangri County	Non-endemic area	92.199444	29.336389	3,757	HCO ₃ :SO ₄ -Ca
SR10	Banggong Village, Sangri County	Non-endemic area	92.380000	29.483889	4,313	HCO ₃ -Ca·Na
NM02	Shangang Village, Nimu County	Endemic area	90.267778	29.569167	4,156	HCO ₃ -Ca
NM03	Old Hedong Village, Nimu County	Endemic area	90.262500	29.551667	4,068	HCO ₃ -Ca
NM04	Old Nixu Village, Nimu County	Endemic area	90.261111	29.515278	3,942	HCO ₃ -Ca
NM03-A	New Hedong Village, Nimu County	Non-endemic area ^a	90.266944	29.549167	4,072	HCO ₃ -Ca
NM04-A	New Nixu Village, Nimu County	Non-endemic area ^a	90.264167	29.520833	3,961	HCO ₃ -Ca·Na
NM01	Gonglang Village, Nimu County	Non-endemic area	90.274722	29.598889	4,305	HCO ₃ -Ca
NMS01	Shangang Village, Nimu County	Non-endemic area	90.271389	29.576389	4,201	HCO ₃ -Ca
NM05	Angang Village, Nimu County	Non-endemic area	90.265556	29.485000	3,917	HCO ₃ -Ca
NM06	Xumai Village, Nimu County	Non-endemic area	90.263333	29.468056	3,908	HCO ₃ -Ca·Na
XTM01	Old Lunzhuzi Village, Xietongmen County	Endemic area	88.580639	29.575917	4,158	HCO ₃ -Ca·Mg
XTM02	Old Duicuo Village, Xietongmen County	Endemic area	88.532972	29.507889	4,037	HCO ₃ -Ca·Mg

Table 1 continued

No.	Sampling location	Type of village	Longitude (°)	Latitude (°)	Elevation (m)	Hydrochemical type
XTM01-A	New Lunzhuzi Village, Xietongmen County	Non-endemic area ^a	88.577194	29.578000	4,182	HCO ₃ -Ca·Na
XTM02-A	New Duicuo Village, Xietongmen County	Non-endemic area ^a	88.533750	29.509222	4,036	HCO ₃ -Ca
XTM03	Renqinze Village, Xietongmen County	Non-endemic area	88.509611	29.487722	4,078	HCO ₃ -Ca
GBJD01	Old Xinsheng Village, Gongbujiangda County	Endemic area	92.961667	30.024444	3,666	HCO ₃ -Ca·Na
GBJD02	Nibi Village, Gongbujiangda County	Endemic area	93.022778	30.007222	3,631	HCO ₃ -Ca
GBJD03	Pang Village, Gongbujiangda County	Endemic area	93.043611	29.999444	3,652	HCO ₃ -Ca
GBJD04-A	Old Tangding Village, Gongbujiangda County	Endemic area	93.145278	29.934167	3,521	SO ₄ -Ca
GBJD01-A	New Xinsheng Village, Gongbujiangda County	Non-endemic area ^a	92.951667	30.030833	3,688	HCO ₃ -Ca
GBJD04	New Tangding Village, Gongbujiangda County	Non-endemic area ^a	93.145000	29.940000	3,501	HCO ₃ ·SO ₄ -Ca
GBJD05	Jin Village, Gongbujiangda County	Non-endemic area	93.163611	29.917222	3,463	HCO ₃ ·SO ₄ -Ca

^a Sampling area is a new non-endemic village whose residents were emigrated from an old endemic village

Table 2 Average, maximum, and minimum values of hydrochemical properties and major constituent concentrations of water samples (in mg/L except for water temperature, *T*, in °C, electrical conductivity, EC, in μS/cm and pH)

Type of sample locations	<i>T</i>	pH	EC	HCO ₃ ⁻	SO ₄ ²⁻	Cl ⁻	NO ₃ ⁻	CO ₃ ²⁻	CO ₂	SiO ₂	HBO ₂	Ca	Mg	Na	K	Al	Fe	Sr	TDS
Endemic villages of Sangri County																			
Average	11.7	7.78	83.4	27.5	11.8	2.2	2.8	0.08	1.16	4.6	0.17	12.3	1.3	2.4	0.7	0.23	0.73	0.06	54.6
Max	18.1	8.07	132.1	42.8	23.3	3.0	3.0	0.10	2.80	5.3	0.38	19.8	2.0	4.3	1.6	0.39	1.08	0.09	85.6
Min	7.1	7.46	56.7	18.5	4.0	1.7	2.6	0.04	0.34	3.4	0.00	7.0	0.9	1.0	0.2	0.06	0.40	0.04	41.2
Non-endemic villages of Sangri County																			
Average	15.6	7.94	145.2	72.5	10.3	2.6	2.7	0.53	1.79	7.3	0.09	21.5	2.1	4.6	1.2	0.08	0.30	0.09	91.5
Max	18.4	8.41	220.2	128.9	21.4	2.9	3.1	0.79	3.09	13.5	0.26	35.5	3.3	5.5	2.1	0.11	0.42	0.16	136.5
Min	13.1	7.57	95.7	38.3	4.8	2.2	2.1	0.08	0.35	3.1	0.00	11.9	1.6	3.4	0.7	0.04	0.18	0.06	63.4
Endemic villages of Nimu County																			
Average	13.4	7.67	52.7	25.5	3.1	2.6	3.1	0.06	1.04	4.2	0.04	8.0	1.3	2.9	1.0	0.45	0.95	0.05	41.4
Max	14.1	7.78	55.1	27.8	3.1	2.6	3.1	0.07	1.24	4.6	0.04	8.8	1.4	2.9	1.1	0.59	1.22	0.05	44.6
Min	12.9	7.57	51.4	24.2	3.1	2.6	3.0	0.04	0.77	3.8	0.03	7.6	1.2	2.8	0.9	0.32	0.74	0.04	39.8
Non-endemic villages of Nimu County																			
Average	15.1	7.70	116.0	61.9	5.2	2.8	3.3	0.25	2.10	5.1	0.15	15.9	1.5	6.0	2.2	0.10	0.34	0.12	75.8
Max	18.4	7.96	178.9	96.5	9.0	3.2	3.6	0.62	2.78	8.8	0.31	22.5	1.9	12.1	3.4	0.30	0.86	0.22	113.2
Min	10.7	7.48	42.9	17.4	2.5	2.6	3.0	0.03	0.76	3.5	0.02	6.1	1.1	2.4	0.7	0.01	0.08	0.04	33.3
Endemic villages of Xietongmen County																			
Average	11.9	8.33	86.7	85.1	8.7	2.7	3.2	1.16	1.10	14.3	0.01	17.8	7.3	3.4	4.5	8.25	14.2	0.08	129.2
Max	12.2	8.64	87.0	104.5	11.6	2.7	3.2	1.64	1.91	14.6	0.01	19.4	9.3	3.6	5.3	8.97	15.2	0.09	142.5
Min	11.6	8.01	86.3	65.8	5.8	2.6	3.2	0.69	0.29	13.9	0.01	16.2	5.4	3.2	3.6	7.53	13.3	0.08	115.9
Non-endemic villages of Xietongmen County																			
Average	14.5	8.20	106.0	53.0	6.4	2.6	3.2	0.57	1.34	6.5	0.29	14.3	2.3	4.7	1.9	0.25	0.44	0.08	71.3
Max	16.1	8.84	125.3	73.0	8.5	2.7	3.3	1.03	3.24	7.3	0.65	17.1	3.4	5.6	2.4	0.46	0.65	0.11	83.3
Min	13.4	7.62	77.5	28.8	3.9	2.6	3.1	0.18	0.08	5.3	0.01	10.0	1.3	4.0	1.3	0.02	0.08	0.06	56.3
Endemic villages of Gongbujiangda County																			
Average	16.6	8.07	54.9	19.6	8.5	2.6	3.0	0.12	0.35	2.5	0.10	8.3	1.1	2.1	0.7	0.39	0.44	0.03	40.0
Max	20.2	8.31	68.6	33.1	20.2	2.6	3.1	0.23	0.59	3.5	0.23	11.2	1.7	2.9	0.8	0.77	1.00	0.05	48.1
Min	14.3	7.72	42.5	2.6	2.8	2.5	3.0	0.03	0.02	1.9	0.03	6.8	0.8	1.7	0.6	0.05	0.09	0.03	34.3

Table 2 continued

Type of sample locations	<i>T</i>	pH	EC	HCO ₃ ⁻	SO ₄ ²⁻	Cl ⁻	NO ₃ ⁻	CO ₃ ²⁻	CO ₂	SiO ₂	HBO ₂	Ca	Mg	Na	K	Al	Fe	Sr	TDS
Non-endemic villages of Gongbujiangda County																			
Average	16.6	8.32	62.9	26.0	7.5	1.8	3.0	0.53	0.32	2.8	0.07	9.3	1.3	2.3	0.8	0.11	0.18	0.04	43.0
Max	18.2	8.78	76.6	36.2	8.1	2.7	3.1	1.26	0.73	4.9	0.13	10.7	1.7	3.4	1.1	0.16	0.26	0.04	54.3
Min	14.4	7.78	54.6	17.9	7.1	0.0	3.0	0.07	0.10	1.7	0.03	8.4	1.0	1.7	0.5	0.03	0.06	0.03	37.2

Table 3 Average, maximum, and minimum values of trace element concentrations of water samples (in µg/L)

Type of sample locations	Se	Mn	Cd	Co	Cu	Ba	Mo	Pb	Cr	Ni	V	Zn	As	Li
Endemic villages of Sangri County														
Average	0.70	22.1	0.04	0.43	2.0	11.6	2.0	9.6	1.3	8.2	19.6	16.8	5.0	2.2
Max	0.88	31.4	0.10	1.00	9.4	14.6	2.7	23.2	4.2	46.5	42.1	69.7	9.7	5.9
Min	0.51	10.5	0.00	0.00	0.0	5.1	1.1	0.5	0.0	0.0	0.5	0.2	1.0	0.6
Non-endemic villages of Sangri County														
Average	0.84	9.7	0.01	0.23	1.0	7.1	2.3	6.1	1.6	2.4	25.4	12.7	7.8	2.2
Max	1.12	12.7	0.02	0.41	1.8	9.5	4.1	16.5	3.0	5.7	70.6	17.8	10.7	3.1
Min	0.56	6.3	0.00	0.00	0.0	4.8	1.3	0.6	0.0	0.0	0.4	3.8	3.0	1.3
Endemic villages of Nimu County														
Average	0.22	45.7	0.01	0.86	2.8	7.3	1.9	1.6	3.1	2.9	1.3	20.6	1.7	2.4
Max	0.33	55.7	0.01	1.12	2.9	8.1	2.0	1.8	3.2	3.1	1.6	27.5	1.9	3.0
Min	0.07	33.2	0.01	0.66	2.7	6.5	1.8	1.4	2.9	2.7	1.0	17.0	1.6	1.9
Non-endemic villages of Nimu County														
Average	0.23	12.7	0.01	0.40	2.2	6.1	2.4	1.1	2.2	2.0	3.6	75.4	3.5	13.1
Max	0.48	32.5	0.01	0.57	3.9	8.5	6.1	2.7	3.0	3.9	12.1	365.2	8.6	29.3
Min	0.07	1.1	0.00	0.31	0.7	4.0	0.9	0.4	1.8	1.0	0.6	14.0	0.6	1.4
Endemic villages of Xietongmen County														
Average	1.03	879.6	0.08	12.2	24.7	127.3	1.6	25.3	8.2	11.5	24.4	98.0	7.6	28.4
Max	1.05	918.8	0.11	16.1	37.3	160.3	1.9	31.0	8.8	13.7	28.9	107.3	10.3	31.0
Min	1.00	840.3	0.05	8.26	12.2	94.2	1.3	19.6	7.6	9.3	19.9	88.7	5.0	25.7
Non-endemic villages of Xietongmen County														
Average	0.74	23.6	0.01	0.59	3.7	11.9	2.0	1.4	2.4	2.6	2.0	50.6	2.8	5.4
Max	1.05	41.5	0.02	0.84	6.6	16.2	3.0	2.3	2.5	3.7	2.6	93.4	6.0	11.2
Min	0.19	1.8	0.01	0.30	1.4	7.0	1.4	0.5	2.3	1.2	1.5	20.9	1.1	2.5
Endemic villages of Gongbujiangda County														
Average	0.27	26.1	0.06	0.83	31.6	6.9	0.6	1.1	3.1	4.7	0.6	29.0	0.6	1.2
Max	0.35	59.2	0.22	1.88	120.6	11.5	0.8	2.0	3.8	9.1	1.4	59.9	0.9	1.8
Min	0.13	5.0	0.01	0.38	1.4	4.5	0.5	0.6	2.1	1.7	0.2	14.9	0.3	0.6
Non-endemic villages of Gongbujiangda County														
Average	0.31	6.2	0.02	0.42	3.6	5.3	2.2	2.5	3.5	14.3	0.3	36.8	1.0	1.6
Max	0.48	10.0	0.03	0.48	6.4	6.6	3.4	5.9	3.6	28.9	0.4	63.0	1.4	2.3
Min	0.22	2.2	0.01	0.38	1.3	4.4	0.7	0.6	3.3	1.9	0.2	19.8	0.7	0.8

words, there is basically no difference in hydrochemical type between water samples from KBD-affected areas and those from KBD-unaffected areas. In fact, the concentrations

of most major constituents and the TDS values of the endemic samples do not show anomalies as compared to the non-endemic samples (see Table 2). For instance, the

average TDS values of water samples from the KBD-affected villages of Sangri and Nimu County are 54.6 and 41.4 mg/L, respectively, lower than those from the KBD-unaffected villages (91.5 and 75.8 mg/L, respectively). However, in the KBD-affected and KBD-unaffected villages of Xietongmen County, the average TDS values of the drinking water samples were 129.2 and 71.3 mg/L, respectively, the former being evidently higher than the latter. In Gongbujiangda County, the average TDS values in drinking water are almost the same between the KBD-affected and KBD-unaffected villages (40.0 and 43.0 mg/L, respectively). It is worth noting that a Russian geochemist Vinogradov (1949) proposed that KBD is caused by deficiency of calcium and excess of strontium in drinking water and other ingested media in the environment. However, this proposed idea is not supported by this study. In Sangri, Nimu and Gongbujiangda counties, the average concentrations of both Ca and Sr in the endemic samples were lower than those in the non-endemic samples, whereas in Xietongmen County, the situation was the reverse.

Although there are no significant differences in the concentrations of most major constituents between the endemic samples and the non-endemic samples, Na, Al, and Fe are exceptions. It can be seen from Fig. 2a that the average Na concentrations in the water samples from the KBD-affected villages of Sangri, Nimu, Xietongmen, and Gongbujiangda counties are all less than those from the KBD-unaffected villages of the corresponding county. Similarly, in every county, the average Al and Fe concentrations in the endemic samples are higher compared to the non-endemic samples (Fig. 2b, c). The average Al and Fe concentrations of the endemic samples collected from Xietongmen County are especially high and 33.3 and 32.3 times those of the non-endemic samples, respectively. It is possible that these elements may have relations to the KBD occurrence in the study areas. Interestingly, M. Aiiso and K. Hiyeda have also indicated that chronic intake of Fe from drinking water or food may result in KBD, but the mechanism by which excessive Fe intake induces KBD has not been determined (Guo 2008).

Selenium

Whether Se deficiency in the environment can cause KBD is still under debate. In 1972, Professor D. Mo of Xi'an Medical University of China reported for the first time that the selenium concentrations in drinking waters and food crops in some KBD areas were lower than those in KBD-unaffected areas (Mo et al. 1975). Since then, the relationship between KBD prevalence and selenium deficiency in environmental media or human organs has been intensively investigated in China. The most significant result of these studies is that KBD-affected areas of China

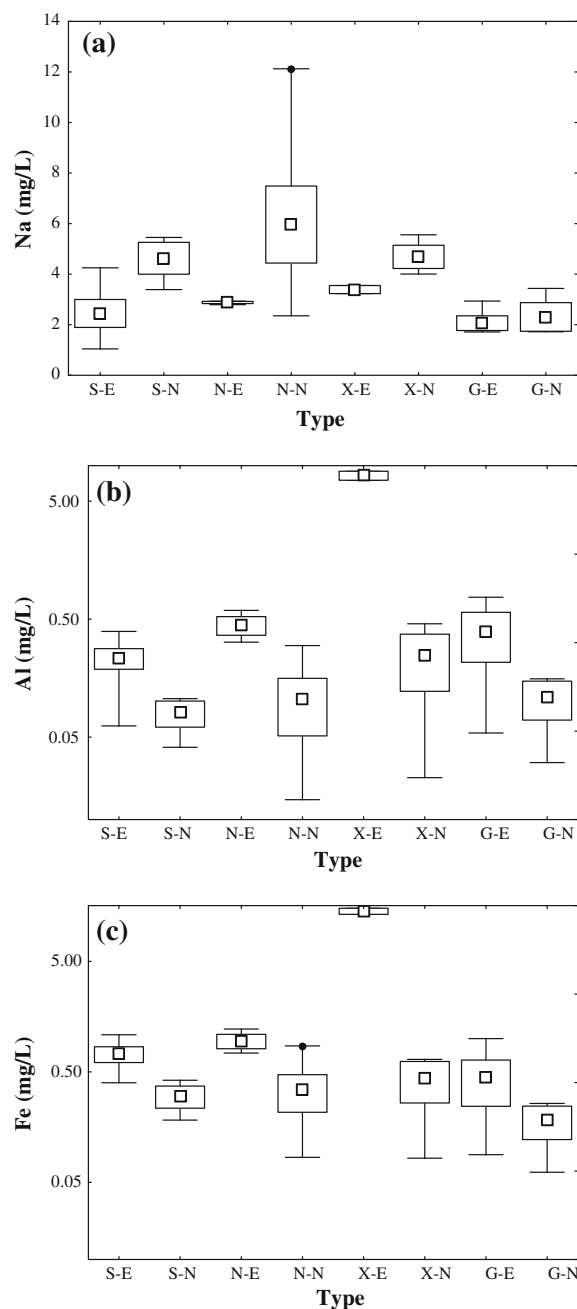


Fig. 2 Box and whisker plots of the Na (a), Al (b), and Fe (c) concentrations for water samples of different type. The meanings of type S-E, S-N, N-E, N-N, X-E, X-N, G-E and G-N are as follows. S-E samples collected from the endemic villages of Sangri County, S-N samples collected from the non-endemic villages of Sangri County, N-E samples collected from the endemic villages of Nimu County, N-N samples collected from the non-endemic villages of Nimu County, X-E samples collected from the endemic villages of Xietongmen County, X-N samples collected from the non-endemic villages of Xietongmen County, G-E samples collected from the endemic villages of Gongbujiangda County, G-N samples collected from the non-endemic villages of Gongbujiangda County. The boxes show the mean value minus standard error, the mean value, and the mean value plus standard error. The smallest and largest values are indicated by the small horizontal bars at the end of the whiskers. Bullet represents outlier values

are generally located along a northeast–southwest belt where the selenium concentrations of surface soils are usually lower than 0.125 mg/kg (Tan et al. 2002). However, there are many exceptions. For example, selenium levels are very low in Yulin and Luonan of Shaanxi Province, China, but there is no KBD occurrence in these two areas. Conversely, in many endemic KBD areas, such as Yidu of Shandong Province and Banma of Qinghai Province, soil selenium levels are relatively high.

To evaluate the relationship between selenium concentration of drinking water and KBD occurrence in Tibet, we compared the average selenium concentrations of the endemic water samples and those of the non-endemic water samples. The results indicate that although the average selenium concentrations in the water samples from the endemic villages of Sangri and Gongbujiangda counties (0.70 and 0.27 $\mu\text{g/L}$, respectively) are slightly lower than

those from the non-endemic villages (0.84 and 0.31 $\mu\text{g/L}$, respectively), the endemic samples from Xietongmen County (1.03 $\mu\text{g/L}$) have higher average selenium concentrations as compared to the non-endemic samples (0.74 $\mu\text{g/L}$). In Nimu County, there is little difference in average selenium concentration between the endemic samples (0.22 $\mu\text{g/L}$) and the non-endemic samples (0.23 $\mu\text{g/L}$). In fact, no significant statistical difference in selenium concentration was found between the endemic water samples and the non-endemic water samples in each of the four counties (the *P* levels are much higher than 0.05). So the selenium concentration of drinking water does not correlate with the prevalence of KBD in the study areas. Further study should focus on inspecting whether or not the prevalence of KBD in Tibet results from selenium deficiencies in other environmental media (such as soil and food crop).

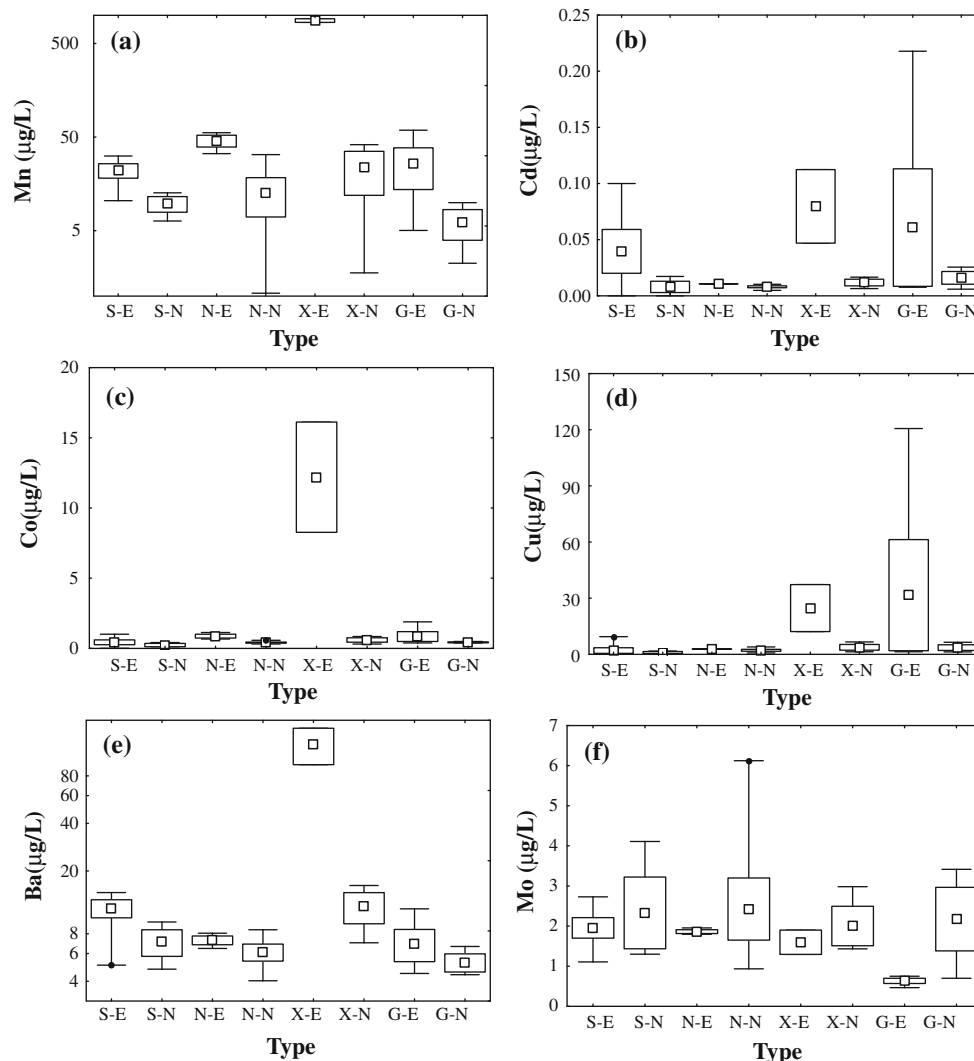


Fig. 3 Box and whisker plots of the Mn (a), Cd (b), Co (c), Cu (d), Ba (e) and Mo (f) concentrations for water samples of different type. The meanings of the types and the legend of the figure are the same as in Fig. 2

Other trace elements

Similar to Se, some other trace elements in drinking water, including Pb, Cr, Ni, V, Zn, As, and Li, do not correlate with KBD occurrence in the study areas. However, Mn in the drinking water samples is different. The average Mn concentration of the endemic samples is higher than that of the non-endemic samples (Fig. 3a). Specifically, the drinking water samples collected from the endemic areas of Sangri, Nimu, Xietongmen, and Gongbujiangda counties have average Mn concentrations of 2.3, 3.6, 37.3 and 4.2 times higher than those from the corresponding KBD-unaffected areas, which is in accordance with the research results of Voshchenko and Ivanov (1990). They concluded that manganese may play an important role in the etiology of KBD occurring in Russia, and proposed that excessive intake of manganese from drinking water can induce osteoclast activation, which in turn inhibits bone growth and causes KBD.

Besides manganese, the average concentrations of Cd, Co, Cu and Ba in the drinking water samples from the KBD-affected and KBD-unaffected villages were also compared, and the results are the same as for Mn. As Fig. 3b, c, d, and e shows, the water samples from the KBD-affected villages have higher average Cd, Co, Cu and Ba concentrations than those from the KBD-unaffected villages. Furthermore, contrary to Mn, Cd, Co, Cu and Ba, the average Mo concentration in the water samples from the KBD-affected areas of each county was lower than those from the corresponding KBD-unaffected areas (Fig. 3f). It is interesting that Cu and Ba were more enriched in drinking waters and soils in the KBD areas of Russia as compared to the background values as well (Muchkin 1967). However, whether or not the Cd, Co, Cu, Ba and Mo concentration anomalies in drinking water are related to the occurrence of KBD has not been ascertained through pathological studies. Thus, more investigations in other KBD areas are needed to further reveal the relationship between the prevalence of KBD and the Cd, Co, Cu, Ba and Mo concentrations in drinking water.

Conclusions

In Sangri, Nimu, Xietongmen and Gongbujiangda counties, there are no distinct differences in hydrochemical type between the drinking water samples from the KBD-affected villages and those from the KBD-unaffected villages. Almost all the water samples are of $\text{HCO}_3\text{-Ca}$, $\text{HCO}_3\text{-Ca-Na}$, and $\text{HCO}_3\text{-SO}_4\text{-Ca}$ types. Also most of the major constituents in drinking water, including HCO_3^- , SO_4^{2-} , Cl^- , NO_3^- , CO_3^{2-} , CO_2 , Ca, Mg, K, Sr, SiO_2 , and HBO_2 ,

are not correlated with KBD occurrence in the study areas. However, the Na, Al and Fe concentrations of drinking water in the KBD-affected villages of Sangri, Nimu, Xietongmen and Gongbujiangda were significantly different from that in the KBD-unaffected villages of the corresponding county, indicating that the anomalous concentrations of these constituents may be related to the prevalence of KBD.

The Se concentrations in drinking water are not correlated to the occurrence of KBD. In Xietongmen County, the average Se concentration of the endemic water samples is even higher than that of the KBD-unaffected samples. Among other trace elements, Mn, Cd, Co, Cu and Ba are of higher average concentration in the endemic samples than in the non-endemic samples, whereas the results for Mo are lower. In other words, the enrichment of Mn, Cd, Co, Cu and Ba and the deficiency of Mo in drinking water may be related to the prevalence of KBD in the study areas. However, the anomalous differences in these elements must be further substantiated in other areas, and pathological studies are required to substantiate any link between KBD disease and these elemental anomalies.

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References

- Appelo CAJ, Postma D (1996) *Geochemistry, groundwater, and pollution*. Balkema, Rotterdam
- Bi H, Zhang S, Luo Y, Feng J, Wu S, Kuang K, Shi S, Yang J, Wang Z, Zheng X (2001) A comparative investigation on the role of fungi and their toxins in food from diseased area of Shaanxi Province in both causes of Keshan disease and Kaschin-Beck disease. *J Xi'an Med Univ* 22(4):308–311 (in Chinese with English abstract)
- Chasseur C, Suetens C, Nolard N, Begaux F, Haubruge E (1997) Fungal contamination in barley and Kashin-Beck disease in Tibet. *Lancet* 350:1074
- Chasseur C, Suetens C, Michel V, Mathieu F, Begaux F, Nolard N, Haubruge E (2001) A 4-year study of the mycological aspects of Kashin-Beck disease in Tibet. *Int Orthop* 25(3):154–158
- Deng J, Li D, Liang D, Xia Z (1999) Role of selenium in the etiology of KBD: a case study on relationship between KBD and selenium in Sichuan Province. *Chin J Control Endemic Dis* 14(4):249–251 (in Chinese)
- Fan Z (2005) Relationship of Kashin-Beck disease and low selenium in environment. *Stud Trace Elements Health* 22(6):64 (in Chinese)
- Gong H, Zaxi S, Baima C, Gesang D (2004a) Prevention and control effect of iodized salt and selenite supplement on Kashin-Beck

- disease in Xietongmen County, Tibet. *Endemic Dis Bull* 19(1):61–62 (in Chinese)
- Gong H, Zaxi S, Xirao R, Gesang D (2004b) Report of sampling investigation of Kashin-Beck disease cases in Tibet. *Chin J Endemiol* 23(1):91 (in Chinese)
- Guo X (2008) Progression and prospect of etiology and pathogenesis of Kashin-Beck disease. *J Xi'an Jiaotong Univ (Med Sci)* 29(5):481–488 (in Chinese with English abstract)
- Guo X, Ding D, Zeng L, Yu Z, Chen F, Zhao Z, Wang J, Bi H (1999) The prospective study of relationship between low selenium and Kashin-Beck disease. *J Xi'an Med Univ* 11(1):1–7
- Haubruege E, Chasseur C, Deboucq C, Begaux F, Suetens C, Mathieu F, Michel V, Gaspar C, Rooze M, Hinsenkamp M, Gillet P, Nolard N, Lognay G (2001) The prevalence of mycotoxins in Kashin-Beck disease. *Int Orthop* 25:159–161
- Haubruege E, Chasseur C, Suetens C, Mathieu F, Begaux F, Malaisse F (2003) Mycotoxins in stored barley (*Hordeum vulgare*) in Tibet Autonomous Region (People's Republic of China). *Mountain Res Dev* 23(3):284–287
- Hinsenkamp M (2001) Kashin-Beck disease. *Int Orthop* 25:133
- Hinsenkamp M, Rypkens F, Begaux F, Mathieu F, Maertelaer VD, Lepeire M, Haubruege E, Chasseur C, Stallenberg B (2001) The anatomical distribution of radiological abnormalities in Kashin-Beck disease in Tibet. *Int Orthop* 25:142–146
- Institute of Kashin-Beck Disease of Harbin Medical University (2000) Investigative report on the prevalence condition of Kashin-Beck Diseases (KBD) in Tibet. *Chin J Endemiol* 19(1):41–43 (in Chinese with English abstract)
- Li J, Ren S, Chen D (1982) A study of Kashin-Beck disease associated with environmental selenium in Shanxi area. *Acta Sci Circumst* 2(2):91–101 (in Chinese with English abstract)
- Li J, Chen D, Ren S, Liu W (1992) Environmental factors causing the low level of Se in human body: a survey on a Kashin-Beck disease region in loess plateau, Shanxi Province. *Environ Sci* 13(6):16–22 (in Chinese with English abstract)
- Mathieu F, Begaux F, Zhang Y, Suetens C, Hinsenkamp M (1997) Clinical manifestations of Kashin-Beck disease in Nyemo Valley, Tibet. *Int Orthop* 21:151–156
- Mathieu F, Begaux F, Suetens C, Maertelaer VD, Hinsenkamp M (2001a) Anthropometry and clinical features of Kashin-Beck disease in central Tibet. *Int Orthop* 25:138–141
- Mathieu F, Suetens C, Begaux F, Maertelaer VD, Hinsenkamp M (2001b) Effects of physical therapy on patients with Kashin-Beck disease in Tibet. *Int Orthop* 25:191–193
- Mo D, Guo S, Liu Z (1975) Dietary hepatic necrosis of white rat induced by drinking water and food in a KBD area. *J Xi'an Med Univ* 39(4):76–85 (in Chinese)
- Moreno-Reyes R, Suetens C, Mathieu F (1998) Kashin-Beck osteoarthropathy in rural Tibet in relation to selenium and iodine status. *N Engl J Med* 339(16):1112–1120
- Moreno-Reyes R, Suetens C, Mathieu F, Begaux F, Zhu D, Rivera T, Boelaert M, Nève J, Perlmutter N, Vanderpas J (2001) Kashin-Beck disease and iodine deficiency in Tibet. *Int Orthop* 25:164–166
- Moreno-Reyes R, Mathieu F, Boelaert M, Begaux F, Suetens C, Rivera MT, Nève J, Perlmutter N, Vanderpas J (2003) Selenium and iodine supplementation of rural Tibetan children affected by Kashin-Beck osteoarthropathy. *Am J Clin Nutr* 78:137–144
- Muchkin NI (1967) Urov's disease in animals. *Veterinariia* 44(3):74–75 (in Russian)
- Pasteels JL, Liu FD, Hinsenkamp M, Rooze M, Mathieu F, Perlmutter N (2001) Histology of Kashin-Beck lesions. *Int Orthop* 25:151–153
- Peng A, Xu L (1988) Redox potential of selenium in Yongshou Kashin-Beck disease region and the influence of humic acid on Eh [Se (VI)/Se (IV)]. *Acta Sci Circumst* 8(2):164–171 (in Chinese with English abstract)
- Sokoloff L (1989) The history of Kashin-Beck disease. *N Y State J Med* 89:343–351 (Published erratum appears in *New York State Journal of Medicine* 89, 486)
- Suetens C, Moreno-Reyes R, Chasseur C, Mathieu F, Begaux F, Haubruege E, Durand MC, Nève J, Vanderpas J (2001) Epidemiological support for a multifactorial aetiology of Kashin-Beck disease in Tibet. *Int Orthop* 25:180–187
- Tan J, Zhu W, Wang W, Li R, Hou S, Wang D, Yang L (2002) Selenium in soil and endemic diseases in China. *Sci Total Environ* 284:227–235
- Vinogradov AP (1949) The etiology of Urovskoi Kashin-Beck disease. In: Borzenko SV (ed) *Transactions of the biogeochemical laboratory*, vol 9. Academy of Sciences of the USSR, Moscow, pp 5–29 (in Russian)
- Voshchenko AV, Ivanov VN (1990) Kashin-Beck disease in the USSR. In: *Proceeding on Kashin-Beck disease*, WHO, pp 152–196
- Wang W, Feng L (1989) An experiment of pathogeny of Kashin-Beck disease caused by humic compounds in water. *Environ Sci* 10(1):16–20 (in Chinese with English abstract)
- Wang W, Wang M, Zhu Z (1985) Geographical and epidemiological studies on Keshan disease and Kashin-Beck disease in Tibet. Science Press, Beijing (in Chinese)
- Wang W, Qi R, Peng A, Li X, Cai G (1991a) Identification of organic compounds in drinking water of Kashin-Beck disease area. *Environ Chem* 10(6):31–35 (in Chinese with English abstract)
- Wang Y, Peng B, Peng A (1991b) Influence of organics in water and soil in Kashin-Beck disease regions on the speciation and bioavailability of selenium. *Environ Sci* 12(4):86–89 (in Chinese with English abstract)
- Wang Z, Xiong Y, Chen J, Bi H, Yang Z, Tan X (2003) Research on relationship between KBD state and selenium as well as human parvovirus B19. *Chin J Endemiol* 22(z1):16–19 (in Chinese)
- Wu D, Ren S, Sun W, Zhang B, He P (1987) A study on relation between contents of soil humic and fulvic acids and Kashin-Beck disease. *Acta Sci Circumst* 7(2):166–171 (in Chinese with English abstract)
- Xu P, Zeng L, Schrauzer GN, Guo X (2000) Selenium, boron, and germanium deficiency in the etiology of Kashin-Beck disease. *Biol Trace Elem Res* 77(3):193–197
- Yang L, Li H, Wang W, Tan J, Li Y (2003) Study on the relationship between Kashin-Beck disease distribution and land use changes in Tibet. *Chin J Control Endemic Dis* 18(5):284–286 (in Chinese with English abstract)
- Yang L, Lv Y, Li H, Li Y, Li S, Wang W, Tan J (2006) Features of geographical environment of Kashin-Beck Disease (KBD) affected region in Tibet. *Sci Geogr Sinica* 26(4):466–471 (in Chinese with English abstract)
- Zaxi S, Gong H, Gesang D, Baima C (2003) Analysis of selenium concentrations of drinking water, food, soil, and human being hair in the Kashin-Beck disease areas of Tibet. *Chin J Control Endemic Dis* 18(3):177 (in Chinese)
- Zaxi S, Baima C, Xirao R, Gong H, Zhao S (2007) Prevention and control effect on Kashin-Beck disease in some experimental sites of Tibet. *Chin J Control Endemic Dis* 22(2):130–132 (in Chinese)
- Zhang L, Zhu Y, Ke F, Ju S, Yu D, Jiang P (1990) Study on relations between Kashin-Beck disease and content of selenium bounded by humic acids in soil in Northeast China. *Chin J Appl Ecol* 1(4):333–337 (in Chinese with English abstract)
- Zhang WH, Neve J, Xu JP, Vanderpas J, Wang ZL (2001) Selenium, iodine and fungal contamination in Yulin District (People's Republic of China) endemic for Kashin-Beck disease. *Int Orthop* 25:188–190