The Level of Vitamin A and Selenium of Cystic Echinococcosis in Aba and Garze Tibetan Autonomous Prefecture, Sichuan, China

Cheng Li  
Affiliated hospital of Southwest Medical University  https://orcid.org/0000-0002-2398-7421

Yongqi Bai  
Affiliated Hospital of Southwest Medical University

Mingcai Zeng  
Ruoergai County People’s Hospital

Yong Luo  
Ruoergai County Center for Disease Control and Prevention

Chumu Luorong  
Daofu County Center for Disease Control and Prevention

Chuan Jin  
Affiliated Hospital of Southwest Medical University

Quilian Guo  
Affiliated Hospital of Southwest Medical University

Qian Wang  
Sichuan Provincial Center for Disease Control and Prevention

Wenjun Liu (lwjlyfy@sina.com)  
Affiliated Hospital of Southwest Medical University

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Abstract

Background

Cystic echinococcus (CE), endemic in Asia, causes disease resulting in a high socioeconomic burden.

Methods

Eighty-nine echinococcosis patients and 274 controls from the Aba and Garze Tibetan Autonomous Prefecture of Sichuan province in China were recruited. The serum levels of Selenium (Se) and vitamin A (VA) of the cases and controls were detected and stratified by age and gender.

Results

The results showed that the male to female ratio of the patients was 1:1.97. The echinococcosis patients were mainly adults. It was confirmed that the serum levels of VA and age were correlated. Compared with the healthy controls, the serum levels of Se and VA of the cases significantly declined in both children and adults.

Conclusions

It was confirmed that the serum levels of Se and VA were interrelated with the prevalence of echinococcosis.

Background

Cystic echinococcosis (CE), a hydatid disease, is a neglected yet serious threat to public health[1]. The parasitic disease is caused by the ingestion of tapeworm eggs from the species E. granulosus s.s. (G1) and E. canadensis (G6/7, G8, and G10)[2, 3]. These species of the echinococcus are highly endemic to China and especially prevalent in the western provinces including Sichuan[4–6]. On an individual level the disease presents a poor prognosis, with high morbidity and mortality rates and causes serious socioeconomic costs[7] often to already vulnerable populations. Infection is highly variable in endemic regions and dependent upon human behavioral risk factors[8].

CE is most commonly associated with the liver[3], an important metabolic organ. The liver stores and metabolizes trace elements including Vitamin A (VA) and Selenium (Se). VA is needed for normal growth and development and maintenance of regular reproductive ability. Variations in the levels of trace elements can influence immune function as well as the infection status of parasite[9]. Researchers have begun to study the relationship between trace elements and echinococcosis. Yet the serum levels of vitamin A and selenium in Sichuan are limited. In this study, we analyzed the serum levels of Se and VA in
echinococciasis patients and healthy controls in Aba and Garze Tibetan Autonomous Prefecture of Sichuan, China.

**Methods**

**Study location and screening of human CE cases**

In 2019, 89 CE patients were selected from Ruoergai County Center for Disease Control and Prevention of Aba Tibetan and Qiang Autonomous Prefecture and Daofu County Center for Disease Control and Prevention of Garze Tibetan Autonomous Prefecture, all the patients were confirmed through B-ultrasonic examination. The control group were the 274 healthy local residents. The case to control ratio was 1:3.08.

**Data Collection**

Blood samples (1 mL) were collected from the cases and controls with coagulant and centrifuged at 3000rpm for 10 min. The serum was collected and stored at -20 °C. Before detection, the standard curve was established. 20µL serum was mixed with 50 mL methyl alcohol (contain 0.5 µg/mL internal standard of Vitamin A acetate 3–4% CIS). After degeneration, precipitation, and mixture with trichloromethane, retinol was extracted completely. Finally, 180µL retinol was isolated and dried by nitrogen and dissolved with 100µL methyl alcohol (contain 5% dichloromethane). The liquid was used to detect VA by HPLC-MS/MS. Gradient ascent of national standard Se (GSB04-1751-2004) were mixed with 100µL serum for the Se test. After 15 minutes standing, the sample was diluted by Nitric acid solution and detected using ICP-MS.

**Data Analysis**

Location, sex, and age of the cases were analyzed with descriptive statistical methods. Spearman correlation analysis was performed to measure the statistical dependence between combinations of the variables using SPSS. The criterion for significance was $P < 0.01$.

**Results**

Eighty-nine CE patients and 274 healthy controls from the Aba and Garze Tibetan Autonomous Prefecture of Sichuan (Table 1). The average age of cases was 39.3 (22.2) while the average age of the controls was 16.6 (11.5). Of the 89 echinococciasis cases, 30 were males (33.7%) and 59 were females (66.3%). The male to female ratio of the cases was 1:1.97. Of the 274 controls 137 were males (50.0%) and 137 were female (50.0%). The male to female ratio of the cases was 1:1. The echinococciasis patients were divided into 2 age groups children and adults. Of the cases 29 were children (32.6%) and 60 were adults (67.4%). The ratio of adults to children was 1:2.1. Of the controls 205 were children (74.8%) and 69 were adults (25.2%). The ratio of adults to children was 1:3.0.
Table 1
Demographic factors for all echinococcosis patients and control

| Demographic factors | Patients | Control |
|---------------------|----------|---------|
| Age (Mean ± SD)     | 39.3 ± 22.2 | 16.6 ± 11.5 |
| Children            | 29       | 205     |
| Adults              | 60       | 69      |
| Gender (M/F)        | 30/59    | 137/137 |
| Selenium            |          |         |
| Deficiency          | 44       | 226     |
| Normal              | 45       | 48      |
| VA                  |          |         |
| Deficiency          | 47       | 89      |
| Subclinical deficiency | 35   | 108     |
| Normal              | 7        | 77      |

In this study 44 cases (49.4%) demonstrated Se deficiency while 45 cases (51.6%) presented normal serum Se levels. In the controls, 226 (82.5%) reported Se deficiency while 48 (17.5%) had normal Se serum levels. In this study, 47 cases (52.8%) had VA deficiency, 35 cases (39.3%) subclinical deficiency, and only 7 (7.9%) had normal serum VA levels. Among the controls, 89 (32.5%) demonstrated deficiency, 108 (39.4%) subclinical deficiency, and 77 (28.1%) presented normal serum VA readings.

When Se deficiency level was stratified by gender and age group, females and children had a higher number of Se deficiency cases as compared to males and adults without a significant difference (Fig. 1). Similarly, when VA deficiency level was stratified by gender and age, females and children had a higher number of VA deficiency cases as compared to males and adults with a significant difference among the age stratified comparison (Fig. 2).

The correlation between the variables of VA deficiency, age and gender demonstrates a significant effect only for VA deficiency and age (Table 2).
Table 2
Spearman correlation analysis among Selenium deficiency, VA deficiency and age/gender

| Variable                              | Spearman correlation | P value |
|---------------------------------------|----------------------|---------|
| Selenium deficiency and Gender        | 0.125                | 0.242   |
| Selenium deficiency and Age           | -0.103               | 0.337   |
| VA deficiency and Gender              | -0.06                | 0.605   |
| VA deficiency and Age                 | 0.281                | 0.008   |
| Selenium deficiency and VA deficiency | 0.014                | 0.895   |

When VA deficiency level was stratified by age among cases/controls, both children and adult cases had significantly higher VA deficiency as compared to controls (Fig. 3). When Se deficiency level was stratified by age among cases/controls, both children and adult cases had significantly higher Se deficiency as compared to controls (Fig. 4).

Discussion

There is an important relationship between trace elements and parasite infection. Many parasites can reduce the levels of trace elements of host, such as Se [10]. The body needs more trace elements while combating a parasite infection. Echinococcus cysts also absorb trace elements from the host to support their growth and development. Therefore, the serum level of Se and VA of an echinococcciasis patient will be lower than in echinococcus cyst fluid[11, 12]. This interdependency may support a possible cure for parasitic disease.

In this study, the data of 89 echinococcciasis patients in Aba and Garze Tibetan Autonomous Prefecture of Sichuan province were analyzed. The male to female ratio of the cases was 1:1.97 and gender is a known risk factor for infection [3, 8]. This known risk factor is often explained by cultural economic production habits, as women are often engaged in animal raising and the upkeep of the home increasing their exposure of the echinococcus cyst via infected dogs and livestock[3, 8].

The echinococcciasis patients were mainly adults (58%) where the controls skewed towards a lower age range. Age is a known risk factor for echinococcus infection[3, 8]. Adults have more opportunity to be infected with echinococcus cyst while in productive activities. Due to the lack of early diagnosis of echinococcciasis, young cases infected with echinococcus cyst are rarer. Most studies lack data on infections in children[13] while ours was able to account for their inclusion.

In this study, the serum levels of Se and VA of the echinococcciasis patients were significantly downregulated compared with the healthy people, which was consistent with the result of a previous
Animal studies also show similar results in the decline of VA in hydatid infected specimens [14]. Sufficient damage to the liver, due to the number and type of parasites, can cause acute and chronic hepatic insufficiency influencing the digestion, absorption, or metabolism of trace elements. Furthermore, progressive liver disease results in noted depletion of hepatic retinoid and VA content in humans[15].

The results suggest that Se and/or VA may be important trace elements for the treatment and prevention of echinococcosis. The change of trace elements might be due to host resistance to the echinococcus cyst or the growth of echinococcus cyst by absorbing trace elements. Research about the relationship between the trace elements and echinococcosis are limited especially in humans. Further studies are needed to delineate the role of trace elements in the survival, growth, and development of the echinococcus cyst. Also, research should clarify how the increase of trace elements in the host may prevent and/or treat echinococcosis. The influence of trace elements on host immune function is unknown. Furthermore, studies on the standard levels of trace elements in varying populations would assist in future evaluations.

Conclusions

Eighty-nine Echinococcosis patients and 274 healthy controls from the Aba and Garze Tibetan Autonomous Prefecture of Sichuan province in China were included in this study. The serum levels of Se and VA of the cases and controls was detected and stratified by age and gender. It was confirmed that the serum levels of VA and age were correlated and that VA was significantly downregulated in echinococcosis patients. Female infection was greater at 1:1.97 and the echinococcosis patients were mainly adults. Compared with the healthy controls, the serum levels of Se and VA of the cases significantly declined in both children and adults.

Abbreviations

CE
Cystic echinococcosis
VA
Vitamin A
Se
Selenium

Declarations

Ethics approval and consent to participate

This study protocols was approved by the Ethics Committee of Affiliated Hospital of Southwest Medical University(KY2019012).

Consent for publication
Not applicable.

**Availability of data and materials**

The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request.

**Competing interests**

The authors declare that they have no competing interests.

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**Authors' contributions**

CL conceived and designed the study. CL, YB, MZ, YL, CMLR performed the study. CL, CJ, QG, QW, WL analyzed and contributed reagents, materials, or analysis tools. CL wrote the paper. All authors reviewed the manuscript.

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Figures

Figure 1

Selenium deficiency distribution in age and gendersubgroup of patients A: Selenium deficiency and normal selenium patients distribution in female and male group. B: Selenium deficiency and normal selenium patients distribution in child and adult group.
Figure 2

VA deficiency distribution in age and gender subgroup of patients A: VA deficiency and normal VA patients distribution in female and male group. B: VA deficiency and normal VA patients distribution in child and adult group. **P<0.01, chi-square test.

Figure 3

Comparison of VA deficiency distribution in children and adults between patients and control A: VA deficiency distribution in children subgroup of patients and control. B: VA deficiency distribution in adults subgroup of patients and control. **P<0.01, chi-square test.
Figure 4

Comparison of Selenium deficiency distribution in children and adults between patients and control A: Selenium deficiency distribution in children subgroup of patients and control. B: Selenium deficiency distribution in adult's subgroup of patients and control. **P<0.01, chi-square test.