Hemorrhagic stroke in children caused by Bothrops marajoensis envenoming: a case report

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Abstract

According to the World Health Organization, snakebites are considered neglected diseases. Bothrops, the genus most frequently implicated in envenomations in Brazil, includes the species B. marajoensis Hoge, 1966, part of the complex B. atrox, which is found in the savannas of Marajó Island, Pará state, Brazil, a region that presents scarce epidemiological data. This work reports the first case of hemorrhagic stroke in a child, attributed to delayed medical care after snakebite envenoming by Bothrops marajoensis in Anajás city, Marajó Island, Pará, Brazil, which led to permanent hemiplegia as a sequela.

Keywords: Hemorrhagic stroke, Sequel, Bothrops marajoensis, Marajó Island

Background

Snakebites were included by the World Health Organization on the list of neglected diseases [1, 2]. An incidence of 5 million envenomings annually worldwide is estimated, with the highest incidence in developing countries with agricultural activities [3]. In Latin America there are approximately 129,084 cases annually, with the vast majority being caused by species of the family Viperidae [4]. In Brazil, from 2011 to 2014 there were 112,249 accidents, 72% caused by Bothrops [5].

The genus Bothrops has neotropical distribution, with the species B. marajoensis Hoge, 1966 [6] being found in the savannas of Marajó Island, Pará state, Brazil [7]. This species is part of the complex B. atrox, popularly known as “jararaca”, “surucucurana”, “combóia” and even by “surucucu” [8]. The Bothrops are implicated in a large number of envenomings in the Brazilian Amazon, where they affect mainly rural areas and are considered a public health problem. The epidemiological data on this species are scarce and underestimated in this region [9, 10].

Snake venom is a complex mixture containing biologically active peptides and proteins that can cause local inflammatory responses and changes of blood coagulation due to defibrination, disseminated intravascular coagulation and thrombocytopenia resulting in partial or complete blood incoagulability, leading to systemic bleeding that may provoke a hemorrhagic stroke [11–13]. But in children this picture is poorly described, which encouraged us to report the first case associated with snakebite envenoming by B. marajoensis [14, 15].

Case presentation

A male child younger than 10 years old, from Anajás city, Marajó Island, Pará (latitude 00°59’14” and longitude 49°56’25”) (Fig. 1), was bitten in an attack by a snake identified as B. marajoensis Hoge, 1966 [6] (Fig. 2), in the plantar region of the right foot, on July 14, 2013, while harvesting açaí.

After the bite, the patient reported local pain then local edema. His parent took him by motorized boat for medical assistance in Anajás. During the trip the boat malfunctioned, which delayed medical care for 25 h. During transport, the child became somnolent and stopped communicating. Upon admission at the hospital, he presented with edema on the entire right leg.
ecchymosis, blisters, hematuria and a comatose state, and was classified as severely poisoned.

The child received all four vials of antithropic antivenom available in the service unit. A transfer to Belém city, the capital of Pará state, was requested, and effected 2 days after to the Emergency Department Mario Pinotti by helicopter. He was admitted with the aforementioned clinical manifestations and the same severity, with eight antivenom vials having been infused to complete the number of ampoules required due to the gravity of the case.

He was transferred on 19 July to the University Hospital João de Barros Barreto in Belém, where he was admitted comatose, with right hemiplegia, labial commissure deviation.
to the left side (Fig. 3), heart and lung auscultation without abnormalities, swelling from the foot to the knee, blisters, normal urinary color and volume and signs of infection at the bite site.

He was evaluated by a neurologist who requested laboratory tests (Table 1) and a computed tomography of the brain that showed hemorrhagic lesions (Fig. 4). Conservative therapeutics and antibiotic therapy were applied. During the 19 days of hospitalization, the patient received medical and physiotherapist follow-up, receiving medical discharge with sequelae including right flaccid hemiplegia.

Discussion
A case of hemorrhagic stroke in a child associated with envenoming by *B. marajoensis* in Anajás city, Marajó Island, is herein reported. This type of accident in the pediatric age group is rarely described in the literature, but is more frequently reported in adults. Mosquera, Idrovo and Tafur [16] reported an incidence of 2.6 % of strokes in *Bothrops* victims in Ecuador, where as Thomas et al. [17] found 2.3 % in Martinique. Santos-Soares et al. [18] and Machado et al. [19] also reported cases of stroke following snakebites in Brazil.

*Bothrops* is the main genus involved in snakebites in Brazil [20]. In the Amazon, ophidic accidents are attributed mostly to *B. atrox* species complex, represented in the archipelago of Marajó by *B. marajoensis* [21, 22]. In all 144 cities of Pará state in the 2012–2014 period, there

| Exams                  | Results     | Reference values       |
|-----------------------|-------------|------------------------|
| Hemogram              | 3.61 millions/mm³ | 4.4 to 5.9 millions/mm³ |
| Hemoglobin            | 10.3 g/dL   | 13.8 to 18.0 g/dL      |
| Hematocrit            | 31.1 %      | 40 to 52 %             |
| Leukogram             | 6.54/mm³    | 5,000 to 10,000/mm³    |
| Segmented             | 90 %        | 40 to 75 %             |
| Platelet count        | 121,000/mm³ | 130,000 to 400,000/mm³ |
| APTT*                 | 20.3 s      | 25 to 34 s             |
| Urea                  | 27 mg/dL    | 10 to 40 mg/dL        |
| Creatinine            | 0.5 mg/dL   | 0.4 to 1.2 mg/dL      |
| Creatine kinase (CK)  | 915 IU/mL   | fem <145 l male < 171 |

*Activated partial thromboplastin time
were 14,842 snakebite accidents in total; of these, 86.84 % by *Bothrops*. Two thousand two hundred and seventy-one (2271, i.e., 15.3 %) of these envenomings occurred in the 16 cities of Marajó Island [5].

The venom of snakes of the genus *Bothrops* are complex mixtures of proteins and peptides that present various toxic activities leading to inflammatory, coagulant and hemorrhagic effects [12, 23]. Symptoms of envenoming are determined by the type of venom, the agent, the severity of the accident and by the conditions inherent to the victim, which may include local manifestations (pain, swelling and redness) and systemic manifestations (bleeding). Complications of envenoming with hemorrhagic stroke are more evident in patients with preexisting risk factors such as cardiovascular disease, particularly systemic hypertension, which was not detected in this case [19, 24].

The accident occurred in the countryside, when the child was descending an açaí palm tree (*Euterpe oleracea*), which bears a berry much appreciated by people and is a very popular food supplement. Snakebite is a common and frequently devastating occupational and environmental disease, especially in rural areas of tropical developing countries [25].

The accident site is about 7 hours away from medical care. Using a motorboat, however, the patient was only able to reach the hospital of the municipal headquarters 25 h after the event, due to engine failure on the boat that was transporting him. This is a common situation in the region, as the medical care is distant from where accidents normally occur. The precarious means of transport greatly aggravates the clinical condition of the victim. It is recommended by the Brazilian Ministry of Health that the initial care of the patient bitten by venomous animals should be obtained as soon as possible, with the number of vials based on the severity of the accident [20]. In this case, the initial treatment should consisted of 12 ampoules of antivenom. In Brazil, the antivenom used is liquid and requires electricity for its conservation, which delays treatment, since accidents occur in rural areas [20]; hence, if we had lyophilized serum in these regions, we believe that similar cases would not occur.

Hemorrhagic stroke is characterized by bleeding into brain tissue, the cerebellum, or brain stem, usually by rupture of a small vessel, causing the overflowed blood under pressure in the brain tissue to lead to clinical manifestations that depend on the location and the specific extent of bleeding [26]. The signs and symptoms observed in the present case included an acute onset without history of trauma and progressed to drowsiness and coma. In the United States, half of pediatric strokes are caused by bleeding in association with several other causes, but without references to envenoming by snakes [27]. In Brazil, Bucaretchi et al. [28] did not find any cases of stroke in 322 children envenomed by snakes. In our case, the clinical examination revealed right hemiplegia and deviation of the labial commissure to the left, while the cranial computed tomography scan showed bleeding lesion in the right frontal area, which characterized an acute neurological presentation of hemorrhagic stroke [16].

Intracerebral hemorrhage after snakebite is related to the occurrence of severe coagulopathy, which is not shown in this case due to the fact that the tests had been administered belatedly [29]. Pardal et al. [9] showed that *B. atrox* was responsible for 13.3 % of blood incoagulability while Ribeiro et al. [30] reported four cases of intracerebral hemorrhage caused by *Bothrops* that presented coagulopathy. These brain bleeding disorders have been linked with incoagulability and low platelet count associated with brain capillary endothelial damage caused by hemorrhages and possibly by other toxins found in the snake venom [12, 31]. Other laboratory exams such as blood count showed anemia, while total creatine phosphokinase appeared five times higher than the reference value, which is related to the inflammatory process at the bite site and brain injury in this case. This enzyme is mainly used to assess changes in the heart muscle, skeletal muscle and brain [31].

**Conclusion**

This is the first case report of hemorrhagic stroke in a child attributed to delayed medical care after snakebite envenoming by *Bothrops marajoensis* on the island of Marajó, which led to permanent hemiplegia as a sequel.

**Consent**

Written informed consent was obtained from the legal guardian of the patient for publication of this case report and accompanying images.

**Ethics committee approval**

This manuscript was approved by the Research Ethics Committee of the University Hospital João de Barros Barreto, document number 41157015.3.0000.0017.

**Competing interests**

The authors declare that there are no competing interests.

**Authors’ contributions**

All the authors observed the reported case and contributed to the design of the study and revision of the manuscript. All authors read and approved the final manuscript.

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