

Publication status: Preprint has not been submitted for publication

Clinical management and Hospital experience in Black Widow Bites: Case Series in Matagalpa, Nicaragua

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<https://doi.org/10.1590/SciELOPreprints.11216>

Submitted on: 2025-02-07

Posted on: 2025-02-13 (version 1)

(YYYY-MM-DD)

1 **1. CLINICAL MANAGEMENT AND HOSPITAL EXPERIENCE IN BLACK**
2 **WIDOW BITES: CASE SERIES IN MATAGALPA, NICARAGUA**

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35 **Running head: Black Widow Bites: Clinical Management**

36 **Funding/Support:** This research did not receive a specific grant from any
37 public, commercial, or not-for-profit funding agency.

38 **Financial disclosures:** No financial disclosures.

39

40 **Conflicts of interest**

41 The authors declare no conflict of interest.

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56 2. ABSTRACT

57 Introduction: Black widow spider (*Latrodectus mactans*) envenomation is a medical
58 challenge in regions with limited access to standardized treatments. This study analyzes a
59 series of 10 cases that occurred in Matagalpa, Nicaragua, highlighting the diagnosis and
60 treatment of patients.

61 Description of the cases: Over a nine-year period 10 hospitalized patients, mostly
62 adolescents (60%) and males (90%) from rural areas, were attended. Severe cases made up
63 30% of the total and included symptoms such as abdominal pain, neurogenic shock, and
64 vomiting. Laboratory findings showed leukocytosis and lymphopenia in 50% of the patients.
65 In terms of management, 60% received Aracmyn Plus antivenom, in addition,
66 corticosteroids, antihistamines, and non-steroidal analgesics were used in most cases. There
67 were no adverse reactions to antivenom treatment. All patients were discharged with no
68 mortality.

69 Discussion and conclusion: The lack of knowledge of this disease and the limited availability
70 of antivenom, poses significant challenges to the effective management of latrodectism in
71 the region. Although analgesic treatment was effective in most cases, some patients required
72 intensive care. This highlights the need for studies that reflect local experience in managing
73 this disease. Furthermore, this case series emphasizes the importance of publishing clinical
74 reports in areas where latrodectism has been little investigated. These data are fundamental
75 to generating evidence to optimize treatment and establish specific protocols in Nicaragua
76 and similar regions.

77 1. Keywords:

78 Spider Bites; Latrodectism; Attack; Poisoning; Hospitalization

79

80 3. INTRODUCTION

81 Black widow spiders (*Latrodectus mactans*) are among the few spiders capable of
82 envenoming humans (1). With the ongoing effects of climate change and globalization, these
83 spiders have invaded new ecological niches contributing to a growing global health issue.
84 This problem is particularly pronounced in developing countries, where access to treatment
85 is often limited, and standardized management protocols remain scarce (2).

86 Latrodectism, the envenomation syndrome caused by *Latrodectus* species, is the most
87 common severe spider envenomation with greater presentation in warmer parts of the
88 Americas, Europe, and Australia (3). Pain is the most prominent feature in all cases and is
89 the main reason a poisoning victim comes to a care unit (4). Black widow spider poisoning
90 is frequently reported to poison control centers and the American Association of Poison
91 Control Centers (AAPCC), with approximately 2,000 to 3,000 cases reported yearly (5). The
92 clinical effects of poisoning typically last an average of two days, with symptoms being more
93 severe in young children and older adults (6).

94 In Nicaragua, clinical documentation on latrodectism remains scarce, despite recent
95 discoveries of *Latrodectus mactans* in the region sparking interest in better understanding
96 this condition (7). This case series examines envenomation cases in northern Nicaragua to
97 provide insights into the clinical presentation and management of latrodectism. The study is
98 based on retrospective records obtained in a secondary care hospital, providing guidelines
99 for the early identification of the clinical presentation and the establishment of specific
100 management that improving patient outcomes.

101

102 4. PATIENT INFORMATION

103 1. Case Descriptions

104 Ten cases of latrodectism required hospitalization between January 2013 and December
105 2021, in a secondary regional care hospital unit. The highest incidence occurred in 2013 and
106 2018, with two cases reported each year. Ninety percent of patients were male from rural
107 areas, with a mean age of 15.9 years (SD: 10.2 years). Most of the bites occurred in the
108 morning (60%) and 50% of the patients were working at the time. The bites were most
109 frequently located on the upper limbs (60%) (Table 1).

110 Time to secondary hospital care had a median of 9.5 hours (RIC: 4-48 hours). Within the
111 hospital, 30% of patients were admitted to the intensive care unit (ICU), while 70% were
112 cared for in intermediate care. The median hospital stay in the ICU was 6 days (IQR: 3-14
113 days), while for all patients it was 4 days (IQR: 3-5 days). Thirty percent of the cases were
114 classified as severe poisoning, 20% as moderate, and 50% as mild.

115

116 2. Clinical findings

117 All patients presented throbbing pain and localized edema (100%). Systemic symptoms were
118 observed in 30% of patients, including vomiting, tachycardia, tachypnea, weakness,
119 sweating, and abdominal pain. Arterial hypertension and headache were observed in 20%,
120 while erythema and seizures occurred in 10% of patients (Figure 1).

121

122 3. Diagnostic assessment

123 The diagnostic studies revealed a median leukocyte count of 10,700 cells/ μ L (IQR: 9,700–
124 16,70 cells/ μ L), with a maximum value of 30,000 cells/ μ L observed in one patient.
125 Lymphopenia was documented in 50% of cases, with a minimum value of 725 cells/ μ L in
126 one patient. The mean random glucose level was 132.8 mg/dL (SD: 31.4 mg/dL). Total
127 creatine phosphokinase (CPK) levels were highest in one patient at 5,379 U/L. The mean
128 serum creatinine across all patients was 0.72 mg/dL (SD: 0.19 mg/dL) (Table 2).

129

130 4. **Therapeutic Intervention**

131 Aracmyn Plus antivenom was administered to 60% of patients. Among them, 83% received
132 one vial, while 17% required two vials. All severe cases were treated with antivenom, and
133 40% of mild cases also received it (Figure 2).

134 Supplemental oxygen was required by 30% of patients, including one case requiring invasive
135 mechanical ventilation and two cases using nasal prongs. In addition, 30% received tetanus
136 vaccination, and 80% were treated with analgesics, with non-steroidal anti-inflammatory
137 drugs (NSAIDs) being the most commonly used (88%). Corticosteroids and antihistamines
138 were used in 90% and 80% of cases, respectively. On the other hand, antibiotics were
139 prescribed in 50% of the patients, with crystalline penicillin being the most commonly used
140 antibiotic (80%) in the cases (Table 3).

141

142 5. FOLLOW-UP AND OUTCOMES

143 All patients were discharged without any mortality or allergic adverse events related to the
144 antivenom. However, complications associated with the arachnid stings were documented
145 in three cases: one patient developed a secondary skin and soft tissue infection during
146 hospitalization; another presented neurogenic shock probably caused by *Latrodectus*
147 *mactans* venom toxins, requiring vasoactive amines and invasive mechanical ventilation, but
148 responded satisfactorily to treatment with two vials of antivenom and presented no post-
149 discharge complications. Finally, one patient was discharged with antihypertensive therapy
150 after diagnosis of secondary chronic arterial hypertension; the other seven patients did not
151 present complications related to the envenomation or hospitalization.

152

153 6. DISCUSSION

154 The Black Widow spider, belonging to the genus *Latrodectus*, is one of the most common
155 venomous species affecting humans (8). The female *Latrodectus mactans* has a shiny black
156 coloration with a red marking on the abdomen that resembles an hourglass. The male is half
157 the size of the female and is brown. These spiders are often found in woodpiles, ground
158 cover, cracks, garages, barns, and latrines (9). Male and female black widow spiders are
159 venomous, usually, only the female has fangs long enough to poison humans (10).

160 Spider venom includes different peptides and substances that affect sodium, calcium, and
161 potassium channels in neurons, as well as glutamate and acetylcholine receptors. The venom
162 is composed of five latrotoxins, which are specific neurotoxins: alpha, beta, gamma, delta,
163 and epsilon (6). Three important syndromes are caused by spider bites: latrodectism,
164 loxoscelism, and funnel-web spider syndrome (11). The severity is determined by the
165 amount of venom administered and by factors such as the number of bites and the age of the
166 patient.

167 The venom potency is superior to that of rattlesnakes; it contains several toxins, and proteins
168 with affinity for nerve endings (12). All *Latrodectus* spiders are venomous and
169 approximately 33% of patients present with systemic symptoms and signs (13). This
170 correlates with our described case series, where 30 % of the patients presented with systemic
171 symptoms, probably due to delay in seeking medical attention and subsequent referral to
172 hospital, resulting in more advanced clinical pictures due to lack of timely treatment.

173 In the 10 cases, the most affected age group was adolescents (60%), followed by
174 schoolchildren and adults (20% each), all from rural areas. Half of the patients were working
175 in the field when they were bitten, while 30% were at home during the incident. The most
176 affected areas were the upper limbs (60%) and lower limbs (40%). These findings contrast
177 with a review of 70 records of children hospitalized in a pediatric hospital in Mexico between
178 1978 and 2014, where the prevalence of bites in the male gender was 61.4%, and infants
179 under one year of age accounted for 14.2%. 70% of the patients had contact with the arachnid
180 at home, with the lower limbs, neck, trunk, and abdomen being the most affected areas (12).

181 Latrodectism is a painful syndrome of muscle cramps produced by spider poisoning. Tests
182 used to assess the severity of latrodectism are diverse, pain severity scales are widely used
183 to evaluate pain and monitor treatment (4). The average duration of latrodectism syndrome
184 is 3 to 6 days. Untreated patients have presented with clinical signs for 7 days, but weakness,
185 muscle pain, and discomfort may persist for weeks (14).

186 In the first case reported in this series, the patient persisted with symptoms of envenomation
187 for 7 days, because the antivenom was not available in Nicaragua until he received the
188 antivenom and began to show improvement of all his signs and symptoms. The reason for
189 the prolonged symptoms is probably multifactorial, depending, among other factors, on the
190 patient, the species of *Latrodectus* involved, and the time to presentation (15) .

191 Latrodectism can be classified into three grades of severity. Grade 1 or mild envenomation
192 may be asymptomatic or characterized by local pain around the area of the bite. Grade 2 or
193 moderate envenomations include pain, which may extend regionally to areas other than the
194 bite site, and also include local diaphoresis. Grade 3 or severe envenomations are
195 characterized by generalized pain, diaphoresis, nausea, vomiting, headache, and altered vital
196 signs such as hypertension and tachycardia (2).

197 Three patients in this series presented with generalized abdominal pain. Latrodectism can be
198 misdiagnosed as an acute abdominal condition, so it should always be considered as a
199 differential diagnosis (1). The clinical manifestations of young infants can be confused with
200 scorpion bite poisoning due to irritability and constant crying, common signs in the bite by
201 this arthropod; however, the child who already speaks and suffered a bite by *Latrodectus* can
202 refer intense muscular pain, thoracic and abdominal pain, besides spasmodic muscular
203 crises, this helps in the differentiation with scorpion bite poisoning (12).

204 Intense and persistent pain is present in half to two-thirds of cases of latrodectism and is the
205 main target of treatments. In Australia, local pain radiating to the bitten limb or from the site
206 of the bite is typical, while in North and South America, back and abdominal pain
207 predominate (16).

208 Complications that may occur include compartment syndrome, severe life-threatening
209 hypertension, respiratory distress, myocardial infarction, priapism and premature delivery in
210 pregnant women (10). Four complications, including arterial hypertension, seizure,
211 secondary soft tissue infection, and neurogenic shock in one patient, occurred in the reported
212 cases. Cases of reversible myocarditis with cardiogenic pulmonary edema may occur after
213 black widow spider envenomation and may require mechanical ventilation. The mechanism
214 of myocardial injury is unknown and may include the direct toxic effect of the α -latrotoxin,
215 catecholamine fluctuation, or hypersensitivity response (13).

216

217 **1. Diagnostic Tests**

218 General laboratory methods are used to evaluate *Latrodectus* poisoning, as there are no
219 specific tests for its detection. Tests that may be useful for patient management include a
220 complete blood count, electrolytes, creatine phosphokinase, and urinalysis. Leukocytosis,
221 albuminuria, and an increased serum creatine phosphokinase level are common findings
222 (10). Leukocytosis and increases in serum creatine phosphokinase were present in our case
223 series. It is also important to also measure cardiac enzymes and monitor the
224 electrocardiogram, because of possible cardiac complications that may occur during
225 poisoning (12).

226

227 **2. Treatment**

228 Initial treatment measures after a mild envenomation include local measures such as gently
229 cleaning the bite with mild soap and water, oral analgesia as needed such as paracetamol,
230 ibuprofen, oxycodone or hydrocodone, oral muscle relaxants, and oral benzodiazepines may
231 be used. Still, evidence of their efficacy is lacking and there may be a risk of increased
232 adverse events. If indicated, tetanus prophylaxis should be administered. Antibiotics are
233 prescribed only if there are signs of infection, such as increased erythema, fluctuation, and
234 suppuration, and are rarely necessary (17).

235 In the case of moderate to severe envenomation, local wound care, and tetanus prophylaxis
236 if indicated, parenteral opioids for pain, parenteral benzodiazepines to reduce the frequency
237 and severity of muscle spasms, antiemetic therapy for nausea and vomiting should be given
238 (17).

239 The antivenom if indicated intramuscularly (IM) should be administered in the anterolateral
240 aspect of the thigh; and a tourniquet is applied proximal to the injection site if an adverse
241 systemic reaction occurs. In the case of intravenous (IV) administration, the antidote is
242 administered over 15 to 30 minutes; IV administration is preferred in severe cases with
243 shock. There appear to be no clinical differences in efficacy between IM and IV routes of
244 administration (17). In this series, all patients received the antivenom intravenously, without
245 any adverse event, with previous medication with antihistamines and corticoids.

246 Most toxicologists agree that the use of antivenom is indicated in cases of severe
247 envenomation that does not respond to standard therapy, because the administration of
248 antivenom involves the introduction of a foreign protein into the human immune system,
249 which can lead to early hypersensitivity reactions (5). In our case series, antivenom was
250 administered to all patients with severe envenomation and, in two cases, also to those with
251 mild envenomation. Antivenom should be reserved for patients with severe systemic
252 symptoms, inadequate pain control, and life-threatening symptoms induced by
253 envenomation, such as uncontrolled hypertensive emergencies or premature delivery (18).

254 Anti-poisons are available in several regions of the world, however, treatment in countries
255 where they are not available is largely supportive (19). Treatment is mainly based on
256 antitoxins. Early rehydration can dilute toxins, which may promote their metabolism and
257 prevent shock. Symptomatic treatment with nonsteroidal analgesics and, in severe cases,
258 benzodiazepines and morphine are usually effective, although the duration of symptoms may
259 require hospitalization for 1 or 2 days (6), most patients have a good prognosis without
260 sequelae (8). These findings are similar to ours in that the average hospitalization time was
261 4 days, with no sequelae at discharge, except for patient number 6 of the series, who was
262 discharged with chronic arterial hypertension.

263 At present, no conclusive controlled data are demonstrating that the antivenom is superior
264 to standard analgesics for the control of latrotoxicism symptoms. Based on the available
265 controlled data and retrospective analysis, the efficacy of the antidote as an analgesic in these
266 envenomations is relatively low and, at best, comparable to that of conventional analgesics.
267 There are other atypical analgesic agents, such as ketamine, pregabalin, and clonidine, which
268 may be considered in cases refractory to standard treatments and/or antivenom. Some
269 clinicians suggest that antivenom should be used with caution and only when other
270 therapeutic approaches have failed (19). In case of an anaphylactic reaction, the antivenom
271 infusion should be discontinued immediately and supportive treatment with antihistamines,
272 steroids, and epinephrine should be administered (20).

273 There are other treatments whose efficacy is questionable, as demonstrated by some
274 retrospective studies. In one case series, 96% of patients who received calcium gluconate
275 did not experience pain relief or resolution of other symptoms; these patients required
276 subsequent administration of opioids. In addition, the use of opioids appeared to be effective
277 in some cases, while other patients did not respond to this treatment (21). Therefore, the

278 choice of which analgesic treatment to use will depend on the intensity of the pain, its
279 characteristics, as well as its availability, being as individualized as possible.

280 Since 2014, the use of anti-venom has been reduced, no difference was found in the reported
281 reduction of pain in patients treated with analgesia alone compared with anti-venom
282 treatment, including patients who reported severe pain at presentation (19). Thus, it is
283 emphasized that hospitalization and administration of anti-venom should be limited to
284 patients with severe systemic symptoms or who have insufficient pain management, making
285 rapid pain relief the most crucial goal for all patients (22).

286 This case series describes and analyzes the results of the experience accumulated over
287 several years in the clinical management of black widow spider envenomations. Although
288 these cases are rare in this geographic region, their severity, together with factors such as
289 waiting times for care, length of hospital stay, variability in symptoms and lack of
290 standardization in pain management treatments, and their limited availability, make this
291 problem still relevant, so this descriptive analysis will contribute to optimizing patient care,
292 based on the best available resources.

293 Limitations of this case series include the lack of specific data on certain outcomes, such as
294 pain intensity, which was only recorded qualitatively. In addition, the identification of the
295 causative arachnids was based on testimonies of patients or relatives, who described
296 *Latrodectus mactans* according to its morphological characteristics, a method still commonly
297 used for its recognition. Only in the first case reported in 2013 in Nicaragua, the arachnid
298 was sent to an entomological center in the west of the country for precise identification,
299 which facilitated the importation of the antidote for the first time since it was not available
300 in the country due to the absence of previous reported cases of this type of envenomations
301 (23).

302 **3. Conclusion**

303 Enhancing the understanding of the pathophysiology of latrodectism is crucial for
304 developing more effective treatments. It is important to continue exploring alternatives that
305 optimize clinical management, establish standardized protocols, and conduct larger
306 randomized controlled clinical trials with new drugs. These studies will enable a more
307 accurate assessment of their safety and efficacy across diverse populations, generating high-
308 quality evidence to improve patient care.

309 7. ETHICS APPROVAL AND INFORMED CONSENT

310 This study was carried out with the approval of the Ethics Committee of the Hospital Escuela
311 César Amador Molina, as well as with the support of the Ministry of Health of Matagalpa,
312 Nicaragua (SILAIS-Matagalpa), for the collection of data from medical records.

313

314 8. AUTHOR CONTRIBUTIONS

315 J.F.G.-S. Contributed to the study design, conceptualization of the study, and drafting of the
316 manuscript.

317 L.M.L. Participated in data collection and contributed to the methodology section of the
318 study.

319 V.E.V.-V. Assisted in the analysis of data and contributed to the writing of the manuscript.

320 V.G.-H. Played a role in the drafting of the study and data analysis.

321 M.M.M. Contributed to the design and execution of the study and reviewed the manuscript.

322 D.S.L.-D. Provided expertise in the methodology and helped refine the analytical approach.

323 G.R.R. Contributed to data collection and participated in drafting the manuscript.

324 C.A.N. Led the study design, supervised the overall study process, and contributed to the
325 final review of the manuscript.

326 These contributions were made collaboratively to ensure a uniform approach to the design
327 and execution of the study.

328

329 9. ACKNOWLEDGMENTS

330 Department of Statistics of the Hospital Escuela César Amador Molina, Matagalpa,
331 Nicaragua, for their valuable collaboration in facilitating and organizing the medical records
332 used in this case series. We also thank the medical and nursing staff for the attention and
333 exceptional care provided to each patient during their stay at the hospital.

334

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400

Tables

Table 1. Baseline characteristics of the patients.

Patient	Sex	Origin	Age (years)	Year	Time of day of Spider Bite	Location of bite	Activity performed	Time of initial care (hours)	Care Level	Length of stay (days)	Severity of clinical presentation
1	Male	Rural	13	2013	Morning	Left wrist	Cutting firewood	3.5	Intensive care	14	Severe
2	Male	Rural	6	2013	Afternoon	Right hand	Manipulating the spider	48	Intermediate care	4	Moderate
3	Male	Rural	7	2014	Morning	Right foot	NA*	48	Intermediate care	4	Mild
4	Male	Rural	16	2016	Morning	Left foot	NA	4	Intermediate care	4	Mild
5	Male	Rural	12	2016	Afternoon	Right foot	Sleeping	96	Intermediate care	5	Mild
6	Male	Rural	21	2017	Night	Right thigh	Sleeping	3	Intermediate care	4	Moderate
7	Male	Rural	12	2017	Morning	Left hand	Walking in the field	6	Intermediate care	1	Mild
8	Male	Rural	12	2018	Afternoon	Right and Left hand	Stripping corn on the cob	12	Intensive care	6	Severe
9	Male	Rural	18	2018	Morning	Left forearm	Working the field	7	Intensive care	3	Severe
10	Female	Rural	42	2020	Morning	Left hand	Cutting coffee	24	Intermediate care	3	Mild

*NR: Not available.

Table 2. Diagnostic tests.

Patient	Leukocytes count (cells/μL)	Lymphocytes count (cells/μL)	Glycemia (mg/dL)	Total CPK* (U/L)	Serum Creatinine (mg/dL)
1	30000	1400	114	1337	1.0
2	9800	1245	127	NA*	0.7
3	16700	1787	77	NA	0.6
4	9800	725	186	NA	1.0
5	8700	2932	153	NA	0.7
6	9400	2303	97	867	0.8
7	11600	NR*	142	NA	0.4
8	13200	1320	158	1040	0.5
9	19000	760	129	5379	0.7
10	9700	2386	145	NA	0.8

*Abbreviations; CPK: Serum creatine phosphokinase; NA: Not available; NR: Not reported.

Table 3. Medications administered.

Medications	Number of patients	Percentage (%)
Tetanus toxoid	3	30
Corticosteroids	9	90
Diphenhydramine	8	80
Analgesic	8	80
NSAIDS*	7	88
Opioids	1	12
Parenteral hydration	10	100
Oxygen	3	30
Endotracheal intubation	1	33
Nasal prongs	2	67
Antibiotics	5	50
Crystalline penicillin	4	80
Cloxacillin	1	20
Aracmyn plus antivenin	6	60
2 bottles of antivenin	1	17
1 bottle of antivenin	5	83

*NSAIDs: Non-steroidal anti-inflammatory drugs.

Figures

Figure 1. Clinical manifestations of the patients.

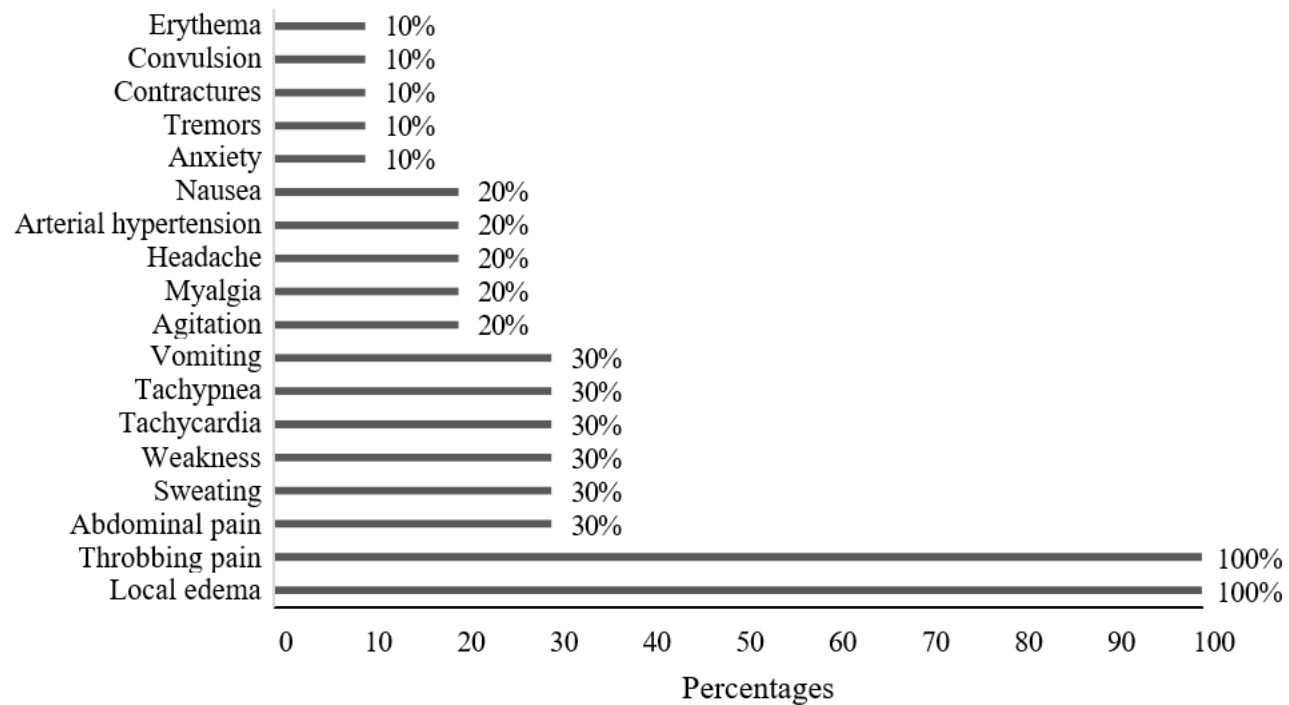
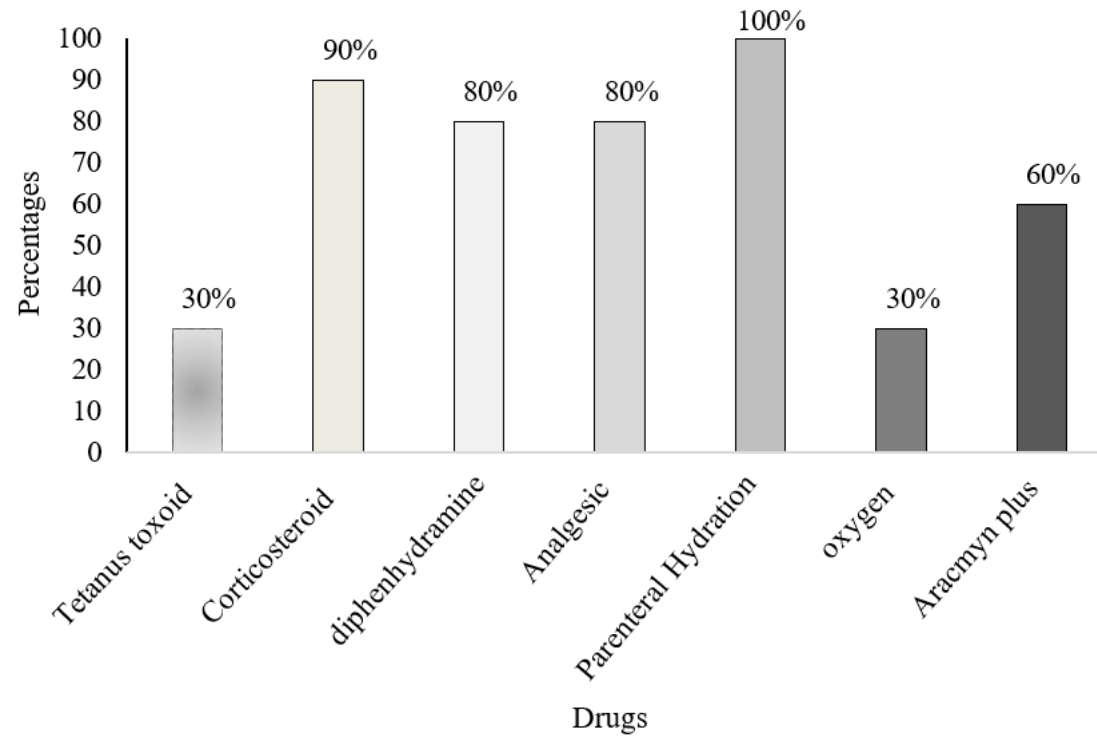


Figure 2. Medications administered to the patients.



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