7 – ORIGINAL ARTICLE WOUND HEALING

Schinus terebinthifolius raddi (Aroeira) and Orbignya phalerata mart. (Babassu) effect in cecorrahphy healing in rats¹

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ABSTRACT

PURPOSE: To evaluate the effect of *Schinus terebinthifolius* Raddi (aroeira) and *Orbignya phalerata* Mart. (babassu) in the healing process of cecorrhaphy in rats.

METHODS: Fifty four rats were used, distributed into three groups randomly: aroeira, babassu and control, which were divided into three subgroups (six animals) according to the time of the deaths (7, 14, 21 days). All underwent the same surgical procedure, cecotomy and cecorrhaphy. The animals in group aroeira and babassu received daily dose of 100 mg/kg of hydroalcoholic extract and 50 mg/kg of aquous extract respectively, by gavage. The control group received only saline solution. The parameters evaluated were: macroscopic changes, ,resistance test to air insufflations and histological changes.

RESULTS: All animals showed good healing without infection. All groups presented adhesions between cecum and neighboring organs. The resistance test insufflating of atmospheric air showed progressive increase of pressure according to the days in the aroeira group, and decrease in babassu group, without significant difference. Microscopy showed significant difference in the polymorphonuclear, hyperemia, angiogenesis, fibroblast proliferation and collagen histological variables in the 14th day.

CONCLUSION: Hydroalcoholic extract of aroeira and the aqueous extract of babassu favored the healing process in cecorrhaphy in rats.

Key words: Wound Healing. Colon. Phytotherapy. Plant Extracts. Rats.

Introduction

The healing process is a biological, chemical and physical complex event involving inflammation, chemotaxis, proliferation, differentiation and cellular remodeling. It can be divided into three phases: 1st or exudative inflammation; 2nd or proliferative/ fibroblastic (granulation tissue with deposition of extracellular matrix); 3rd or remodeling/maturation¹⁻⁶. The knowledge of each of these stages is important to guide appropriate adjuvant therapy to improve biological reaction⁶.

The key factors for good healing in the gastrointestinal tract are closely related to the layer of collagen in the submucosa, good vascularization of the bowel edges, contamination by intestinal contents, submucosal suture, and delicate handling of wounded tissues⁷.

The application of medicinal plants in healing of colonic lesions, as well as other tissues and organs, has been demonstrated in several studies⁸⁻¹⁰. Brazil has the largest diversified forest reserve on the planet. Many species are used for medicinal purposes with low or no evidence of its properties¹¹. The use of medicinal plants is not restricted to rural areas or regions devoided of medical and pharmaceutical care. They are also used extensively in the urban environment as an alternative or supplement to habitual medication. However, most of the data regarding the use of these plants is still empirical, based on popular culture. The interest in discovering new substances stimulated scientists from different areas to look for flora with medicinal properties normally used by the population. This fact can be observed in some studies that assessed the potential healing of plants capable of stimulating surgical repair¹².

The medicinal plant potentiality is based on the presence of active principles able to produce various pharmacological effects. Between the bioactive substances that can be found in different parts of a plant are highlighted alkaloids, saponins, tannins, glycosides, flavonoids and oils.

The pepper tree (*Schinus terebinthifolius* Raddi) is an Anacardiaceae native in Peru, kingdom Plantae, class Magnoliopsida, order Sapindales, family Arecaceae, genus *Schinus*, *Schinus terebinthifolius* species, with geographic distribution throughout the coastal Northeast, Southeast, South and Central Western Brazil, and natural location in the states of Paraná, Santa Catarina and Rio Grande do Sul. Known as toxic plant, is widely used in folk medicine, especially in the treatment of inflammatory processes in general. Many of its properties can be assigned to different polyphenols (defense mechanism), which are distributed unevenly in the various plant organs, such as bark, leaves, flowers, fruits and seeds¹³. Many studies have been done with aroeira: analysis of the morphology and anatomy of its seeds, bark and leaves; chemical and microscopic examination; bark development; fruits; mutagenic effect; antibiotic; antioxidant; antiulcerogenic; fungicide; as well as healing process on colon, bladder, skin, line alba and stomach^{8-10,14-16}.

Babassu (*Orbignya phalerata*) is another plant characterized by being a palm tree of kingdom Plantae, Magnoliophyta phylum, class Liliopsida, order Arecales, family Arecaceae, genus *Orbignya*, *Orbignya phalerata* species. It is found more frequently in the North, Northeast and Midwest regions of Brazil, and 54.2% of them are located in the state of Maranhão. It consists of four parts: exocarp, mesocarp, endocarp and seeds. The powder is derived from babassu mesocarp and features some proven anti-inflammatory and analgesic properties. Its popular use is indicated in the dysmenorrhea, ulcers, tumors, rheumatism, constipation, colitis and obesity¹⁷.

This study aims to analyze the healing effect of the hydroalcoholic extract of aroeira and babassu aqueous extract administered by gavage in cecorrhaphy in rats.

Methods

This study was performed at Laboratory of Experimental Surgery, Universidade Federal do Maranhão and Mother and Child University Hospital, and was approved by the Ethics Committee of the State University of Maranhão under nº 038/2012.

Forty four Wistar rats were used (*Rattus norvegicus albinus*, Rodentia mammalia) weighing on average 271.3 g, conditioned, acclimatized and fed by the standard form for the species. They were randomized during surgery in three groups of 18: aroeira group (GA), babassu group (GB) and control group (GC). Each group was divided into three subgroups of six animals, dead on the 7th, 14th and 21st days after the procedure.

Hydroalcoholic extract of aroeira

It was prepared at the Laboratory of Natural Products of Chemistry Department, FEPAR, Center of Exact Sciences and Technology from the dried inner bark of a single mature tree registered at Atico Seabra Herbarium of the same university named with the number 488. The guidelines of the Brazilian Pharmacopoeia II were adopted following the known methodology to prepare the extract. The plant material (8 kg of dry bark) was placed to dry at room temperature, crushed with knife and placed for 24 h in oven at temperature between 45-50°C to remove moisture. It was then subjected to the milling process to obtain the powder which resulted in 2.400 g. This amount was deposited in a glass container with addition of 70% alcohol solution at a ratio of 1:3 powder. The resulting mixture was left for 12 h and stirred for 5 min every 2 h; then, through two simple filtration under reduced pressure in a cotton glass funnel, was obtained 3450 ml of purple crude extract. The residual powder was discarded. This extract was concentrated on a rotary evaporator under reduced pressure at temperature between 55° C and 60° C for complete removal of solvent forming a paste of 596.85 g, from which was withdrawn a sample (2.8 g) of the hydroalcoholic crude extract (ethanol) diluted in 28 ml saline solution resulting in concentration of 100 mg/ml.

Aqueous extract of babassu

The babassu mesocarp was obtained from mature coconut that felt naturally on the ground. After drying for three days, it was placed in oven with temperature of 45-50°C for 24 h to remove the moisture and through grinding process was obtained a powder. To obtain the aqueous extract was used 10 g of powder, diluted with 400 ml of distilled water resulting in a concentration of 25 mg/ ml. This final extract was placed in a glass in a refrigerator at a constant temperature of 10°C.

Operative procedure

Before anesthetic induction, the animals were weighed using a digital electronic scale; all of them, but one, were identified on their tails using colored dermographic markers (black, purple, blue, green and red). After seven acclimatization days, they were carried out to surgery preceded by food suspension of 6 h, but with free access to water. Intramuscular anesthesia with 20 mg/kg ketamine hydrochloride 5% and 10 mg/kg xylazine 2% was used in all animals

After anesthesia, they were submitted to ventral abdominal epilation followed by site disinfection with povidoneiodine solution and sterile fenestrated drape covering of area to be operated. No bowel or antibiotic preparation was performed. Median longitudinal laparotomy was carried out over 3 cm caudally with exposure, identification and exteriorization of the cecum, in which was held a 1 cm long transverse incision in the anti-mesenteric border, followed by a suture in full single plane with inversion of the mucosa with blue monofilament polypropylene 6-0. Four separate sutures were done with knots out from the intestinal lumen and overlapping the edges of the cecum. Thereafter the abdominal wall synthesis was performed in two layers with 5-0 monofilament nylon sutures, and subsequent daily IM analgesia with Dipyrone (25 mg/kg/dose) for five days.

Postoperative

After anesthetic recovery, the animals were randomly divided into three groups of 18. The GA received by gavage 100 mg/kg of aroeira, GB 50 mg/kg of babassu oil and the same amount of saline solution 0.9 % in the GC, with daily administration.

After gavage, the animals were accommodated in cages containing six animals each, according to the groups and date of death. Clinical evaluation (surgical site, motor activity, provision for food), specific protocol filling, body weight (even in the scheduled death day) were noted down daily. The animals had free access to water and standard chow until the death day.

Macroscopic evaluation

After the scheduled death of the animals, the surgical scars were inspected looking for signs of infection, dehiscence of the wall, hematoma and abscess. Then, laparotomy was performed with two parallel transverse incisions (cranial and caudal) and a 1 cm left paramedian one, parallel to the median and perpendicular to the transverse incisions. Abdominal findings suggestive of infection, collections, fistulae and adhesions by Nair score were inspected. Surgical specimens were removed 2 cm above and below the suture (cecum, ascending colon and terminal ileum), not undoing the structures and organs attached to it in order to not compromise the inflation test. The specimen, after being washed with water, was identified individually with the animal number and subgroup, and then referred to inflation resistance by air insufflation test (Figure 1).



FIGURE 1 - Resistance to air insufflations. **A)** Probe Polzin[®] conected to silicone catether; **B)** Clear glass container with running water showing the moment of rupture of colonic anastomosis, evidenced by water bubbling with the pressure break marked in mmHg.

Histological evaluation

After insufflation test, surgical specimens containing colonic segment involving the suture line were fixed in a polystyrene plate and sent for standard hematoxylin-eosin and Masson's trichrome staining, evaluated by the same pathologist, blinded to the group identification. The data were classified according to the intensity transformed into quantitative variables by a score index given to histological findings (0-absent, 1-mild, moderate-2 and severe-3). The presence of polymorphonuclear, vascular congestion and edema were indicative of acute inflammation (acute phase). The presence of mononuclear cells, angiogenesis, fibroblast proliferation, fibrosis (collagen deposition) and collagen were indicative of chronic inflammation (chronic phase).

Statistical analysis

Data were analyzed using the statistical program SPSS 20.0 Statistics[®] (2011). Initially, the numerical variables were assessed using the normality of Liliefors test, and was found that none had a normal distribution. Therefore, these variables were evaluated in relation to the effect of group and days within each group using the Kruskal Wallis and post hoc Dunn tests. In histological variables and the degree of adherence effect of group within each day and the day within each group were also performed by these tests. For the association of the variables of the macroscopic evaluation and the rupture site the chi-square and Pearson tests were used. In all, the level of significance (α) was 5%, which is considered significant when p<0.05.

Results

The surgeries of all animals were uneventful and there were no deaths. The daily clinical evaluations showed adequate recovery, as good physical activity and good dietary compliance.

Macroscopic evaluation

The abdominal wall healing was considered adequate in all animals, with no infection, dehiscence, hematoma, abscess, except in one rat of GA, in which partial skin dehiscence occurred without clinical importance. Into abdominal cavity there were not abscess, hematoma, fistula and/or dehiscence.

Adherence gradation (Nair criteria)

The adherence on day 7 was in grade I in all three groups, two in grade II in GA and GB and one in grade III in GB, with no significant difference in the 7^{th} and 14^{th} days. However, significant difference was observed at day 21 (p=0.014, Table 1).

TABLE 1 – Inter-group evaluation in different post operative days concerning to adherence.

Day	Nair ³⁷		Group		р
		Control	Aroeira	Babassu	
7	0	0 (0.0)	1 (16.7)	0 (0.0)	0.638
	Ι	3 (50.0)	3 (50.0)	3 (50.0)	
	II	3 (50.0)	2 (33.3)	2 (33.3)	
	III	0 (0.0)	0 (0.0)	1 (16.7)	
14	Ι	1 (16.7)	2 (33.3)	2 (33.3)	0.126
	II	2 (33.3)	4 (66.7)	4 (66.7)	
	III	3 (50.0)	0 (0.0)	0 (0.0)	
21	0	1 (16.7)	1 (16.7)	0 (0.0)	0.014
	Ι	4 (66.7)	0 (0.0)	0 (0.0)	
	II	1 (16.7)	5 (83.3)	6 (100.0)	
In	0	1 (5.6)	2 (11.1)	0 (0.0)	0.195
general	Ι	8 (44.4)	5 (27.8)	5 (27.8)	
	II	6 (33.3)	11 (61.1)	12 (66.7)	
	III	3 (16.7)	0 (0.0)	1 (5.6)	

Test: Kruskal Wallis. Significance level p<0.05

Adhesions between an organ and abdominal wall or between two organs were found in eight GC rats and in five GA and GB (Figure 2).



FIGURE 2 – Examples of adhesions between organs and abdominal wall. **A**) Grade I; **B**) Grade III.

In GB one rat showed widespread adhesion without adhering to the abdominal wall (Nair III) on day 7 (Figure 3B). At day 21 was evidenced grade II in all GB animals and in great amount on GA, while in the GC it was observed in only one animal, with significant difference (p=0.014).

Resistance evaluation (air insufflation test)

The mean values of burst pressure were evaluated by chi-square test, observing progressive increase in GA according to the days and decreasing in GB. However, no significant difference between the groups was achieved (Figure 3).



FIGURE 3 - Mean values of burst pressure in the colon.

The resistance test to air insufflation was performed in all rats, uneventfully. In control animals, on day 7, the break point was within the suture in five; four animals on day 21 had rupture outside the suture. In GB there was no difference compared to days. In the GA on the 21st day, the break was out of the suture in all animals, showing significant difference (p=0.049) (Table 2).

 TABLE 2 – Breaking point according to post operative day (intra-group).

Days	Group	Rupture	р	
		Out of the suture	On the suture	
Control	7	1	5	0.135
	14	4	2	
	21	4	2	
Aroeira	7	4	2	0.049
	14	2	4	
	21	6	0	
Babassu	7	4	2	1.000
	14	4	2	
	21	4	2	

Test: chi-square - Pearson. Level of significance p<0.05

Microscopic evaluation

In Kruskal Wallis histological variances in relation to the day within the groups (inter-group) there was significant difference on day 14 (Table 3).

TABLE	3 - Variances related to	histology, according to
the post operative	day within inter-group.	

Dav			Group		р
		Control	Aroeira	Babassu	•
7	Polymorpho- nuclear	Mod./ Accent.	Mod./ Accent.	Accent.	0.385
	Monomor- phonuclear	Mod./ Accent.	Accent.	Mod.	0.057
	Edema	Mod.	Mod.	Mod.	0.322
	Congestion	Mod./ Accent.	Accent.	Accent.	0.150
	Angiogenesis Fibroblast proliferation	Mod. Mod./ Accent.	Accent. Accent.	Mod./ Accent. Accent.	0.533 0.123
	Fibrosis	Slight	Slight	Abs	0.189
	Collagen	Slight	Slight	Abs/ Slight	0.284
14	Polymorpho- nuclear	Mod. a	Slight/ Mod. ab	Abs b	0.004
	Monomor- phonuclear	Accent.	Mod./ Accent.	Slight/ Mod.	0.104
	Edema	Mod.	Slight/ Mod.	Slight/ Mod.	0.809
	Congestion	Accent. a	Mod. b	Slight c	0.004
	Angiogenesis Fibroblast proliferation	Mod. a Accent. a	Mod./ Accent. a Accent. a	Slight b Mod. b	0.011 0.030
	Fibrosis	Slight	Slight	Abs	0.056
	Collagen	Slight a	Slight a	Abs b	0.033
21	Polymorpho- nuclear	Abs/Sli- ght	Slight	Slight	0.414
	Monomor- phonuclear	Mod.	Mod.	Mod.	0.087
	Edema	Slight	Slight/ Mod.	Mod.	0.533
	Congestion	Mod.	Mod.	Mod.	0.812
	Angiogenesis	Slight/	Slight/	Slight	0.757
	Fibroblast	Mod.	Mod.	Accent.	0.298
	proliferation	Accent.	Accent.	4.1	0.402
	F1bros1s	Abs/ Slight	Abs	Abs	0.492
	Collagen	Abs/ Slight	Slight	Abs	0.533

Test: Kruskal Wallis. Legend: a, b and c refer to p<0.05; Accent.=accentuated; Mod.=moderate; Slight= slight; Abs=absent

Regarding the presence of polymorphonuclear, it was observed significant difference on day 14th between the GC and GB (Table 4).

	Day/ Group	GC	GA	GB	р
	7	Mod./ Accent.	Mod./Ac- cent.	Accent.	0.385
Polymor- phonuclear	14	Mod. a	Slight/ Mod. ab	Abs b	0.004*
	21	Abs/ Slight	Slight	Slight	0.414
	7	Mod.	Mod.	Mod.	0.322
Edema	14	Mod.	Slight/ Mod.	Slight/ Mod.	0.809
	21	Slight	Slight/ Mod.	Mod.	0.533
	7	Mod./ Accent.	Accent.	Accent.	0.150
Congestion	14	Accent. a	Mod. b	Slight c	0.004
	21	Mod.	Mod.	Mod.	0.812
M	7	Mod./ Accent.	Accent.	Mod.	0.057
Monomor- phonuclear	14	Accent.	Mod./ Accent.	Slight/ Mod.	0.104
	21	Mod.	Mod.	Mod.	0.087
	7	Mod.	Accent.	Mod./ Accent.	0.533
Angioge- nesis	14	Mod. a	Mod./ Accent. a	Slight b	0.011
	21	Slight/ Mod.	Slight/ Mod.	Slight	0.757
Fibroblast	7	Mod./ Accent.	Accent.	Accent.	0.123
prolifera- tion	14	Accent. a	Accent. a	Mod. b	0.030
	21	Accent.	Accent.	Accent.	0.298
	7	Slight	Slight	Abs	0.189
Fibrosis	14	Slight	Slight	Abs	0.056
11010010	21	Abs/ Slight	Abs	Abs	0.492
	7	Slight	Slight	Abs/ Slight	0.284
Collagen	14	Slight a	Slight a	Abs b	0.033
	21	Abs/ Slight	Slight	Abs	0.533

TABLE 4 - Detailed inter-group histological analysis.

Test: Kruskal Wallis. Legend: a, b and c refer to p<0.05; Accent.=accentuated; Mod.=moderate; Slight= slight; Abs=absent

Polymorphonuclear cells were found in the GC and GB with moderate and absent intensity, respectively. Edema was present with moderate intensity in all groups except GC on day 21 that was discrete, with no significant difference between groups (Table 4).

Congestion was observed with increased intensity in the three groups on day 7 with no significant difference between groups at 7 and 21 days. However, significant differences between the three groups were found on day 14 (p=0.004, Table 4).

The mononuclear cells were observed with increased intensity on day 7 in GA and GC on the 14th day, being discrete in GB on the 14th day, with no significant difference between groups (Table 4).

Regarding angiogenesis, a significant difference was evident at 14 days between the GC, and GA with GB, with no difference between the first two (Table 4), with mild and moderate/ severe intensity in GA and GB, respectively).

Fibroblast proliferation had significant difference at 14 days between the GC and GB, with severe and moderate intensity, respectively (p=0.03) (Table 4). On the other days, the variable appeared with increased intensity.

Fibrosis was observed discrete or absent in all groups and days, showing no significant difference between them (Table 4).

Regarding the presence of collagen was similar to fibrosis, but significant difference was observed on day 14 between GC and GA with GB (p=0.033), with no difference between the first two (Table 4). The collagen was evident in GC and GA discreetly and absent in GB.

Discussion

This study was done using only male rats to avoid influences of the hormonal cycle of females¹⁸. The days of death were chosen based on the studies of Goes *et al.*¹⁹ on the healing of colonic anastomoses in rats killed at intervals of 3, 7, 14 and 21 days. Baldez *et al.*¹⁸ and Ferreira *et al.*²⁰ chose as death periods 3 and 7 days postoperatively. Considering these differences, were used in this study 7, 14 and 21 days for scheduled death, since between the 3rd and the 7th days the healing process is quite similar, and at 21 days there is real surgical healing, allowing true comparison.

Herbal choice and route of administration

The aroeira has popular use in home treatment; may be administered orally or topically for curing various diseases. The reason that influenced its choice was easy extraction, preparation, drying, grinding, storage and application²¹, and also because its current use as medicine herbal, though without scientific confirmation^{8,16}.

Preclinical studies in mice showed that the extract of babassu mesocarp had beneficial effect on several circumstances²². It is used as food and in the treatment of various diseases; it has a wide distribution in Maranhão state in Brazil.

The dose chosen was based on other studies referring better results with 50 mg/kg of babassu extract²³ and 100 mg/ kg of the aqueous extract of aroeira¹⁶. Already the oral route of administration by daily gavage^{24,25} differs from other papers^{23,26} using it parenterally, intraperitoneally and even topically in different dosages, proving that this herbal can be used by different administration routes. Was considered by the authors of this study that daily administration would give better action opportunity and could contribute to the literature with alternative higher dosage.

Macroscopic evaluation and adherence gradation

Macroscopy, regarding complications (infection, dehiscence, fistula, hematoma, abscess, both on the wall and in the abdominal cavity), showed no complications except for one GC rat with wound partial dehiscence; however, this fact did not demonstrate significant difference between groups^{8,27}.

Adherence can be observed as a result of trauma to the peritoneal surface of the intra-abdominal organs promoting inflammatory reaction, followed by increased vascular permeability and releasing rich fibrin exudate. Adhesions were observed in all groups and in all days, but only significant on day 21 in GB wherein all rats had adhesion type II, but without significant difference. But Lucena *et al.*⁹ found significant differences more often in GA than in the GC on the 7th day. In this research, although without significance, on the 14th day, it can be inferred that there was favoritism for GA and GB, as these groups had lower adhesion degrees compared to GC.

Tensiometric evaluation (resistance to air insufflation)

According to Santos *et al.*¹⁰, the mechanical strength of the scar through the resistance to air insufflation test is one of the main parameters for assessing the integrity of the anastomoses in the early postoperative days. As hollow viscus, this test is to reproduce physiological pressure vectors that normally are transmitted to the wall, with clinical simulation, since the break-up will occur depending on the distention.

The burst pressure is significantly lower in the early postoperative days, compared to 7 and 14 days, since the initial anastomose resistance is more dependent on sutures, being the late stage to collagen deposition.

In this research, was observed a progressive increase in burst pressure values in GA according to days, decreasing in GB. In GC, this pressure increased on day 14 and decreased on day 21, all without significant difference. Despite studies of Lucena *et al.*⁹ and Coutinho *et al.*⁸ did not show significant difference, the burst pressure increased with the days. This result may be justified due to the fact that these authors evaluated only until the 7th day.

Petroianu *et al.*²⁸ emphasized that anastomose strength in the late period, after the 14th day, is best assessed by linear traction dynamometers with precision in the range of isolated tissue.

Unlike the study by Santos *et al.*²⁵ in which the rupture site on insufflation test occurred in specific point of the suture line (weakest point), it can be inferred that in this site young collagen predominated; in this study, rupture occurred both inside and outside of suture site, with a significant difference in GA at day 21, when all were outside.

Histological evaluation

In this study, histological assessment used the technique of hematoxylin and eosin staining, where different histological variables are quantified and classified according to the presence and intensity, establishing the classification of healing in acute and chronic inflammation. The presence of vascular congestion, edema and polymorphonuclear were indicative of acute inflammation (acute or inflammatory phase of the healing process). The presence of mononuclear cells, angiogenesis, fibroblast proliferation, fibrosis (collagen deposition) and collagen were indicative of chronic inflammation (chronic or proliferative phase of the healing process). To evaluate the presence of collagen fibers in the area of healing, the technique of Masson trichrome staining was used, although it is not free of criticism for not differentiate collagen from the point of view of quality but quantity. These techniques were used because they are consolidated^{19,20,29}.

The tissue injury healing by first intention causes death of epithelial cells and connective tissue, promoting continuity solution on epithelial basement membrane. The area of the lesion is filled with blood clots, fibrin and white blood cells, approximately 24 h with a peak of up to five days, migrating neutrophils (polymorphonuclear leukocytes) to the edge of the incision to remove crusts - the phagocytosis. The intestinal wall becomes increasingly thickened by increasing vascular permeability (congestion) and causing edema, and the migration of large numbers of acute inflammatory cells.

Regarding the presence of polymorphonuclear cells, significant difference was observed on day 14 between the GC and GB, being moderate and absent, respectively. On the 7th day, groups showed be moderate to severe in most animals, contradicting the study of Coutinho *et al.*⁸ and Santos *et al.*¹⁰ that referred discrete to moderate.

Similar to other papers^{8,27} for the presence of edema, here it was found to be moderate in all groups except on GC day 21 (discrete) without significant difference. However, in other studies^{18,30} it was seen sparingly on the 7th day of GB, with significant difference. Importantly, in all three compared studies the evaluation was on 7th day.

Another important acute inflammatory variable is the histological vascular permeability or congestion. It performed with marked intensity on day 7, consistent with the initial acute phase, there was no significant difference between groups. However, on day 14 there was a significant difference between the GC and GB, where the latter was discrete^{18,27}.

Acute inflammation to some degree is necessary for good response of tissue repair (wound healing); however, intense inflammatory reaction may decrease the blood supply with consequent inhibition of fibroblastic proliferation²⁹.

Acute exudate is gradually replaced by mononuclear cell infiltration (macrophages and lymphocytes) and granulation (angiogenesis and fibroblasts), and deposition of collagen produced by fibroblasts in the margins of the incision, initiates the chronic inflammatory process.

With respect to mononuclear cells, was observed in this study to be moderate to marked in all groups and in all days, with marginal significant difference noted on day 7 between GA and GB. Studies of Lucena *et al.*⁹ and Santos *et al.*²⁵ showed intensity on day 7 of chronic inflammation favoring the experimental group with a significant difference in angiogenesis variable. In the present study, there was significant difference between groups. Batista *et al.*²⁹ have published significant trend for the experimental group (babassu), suggesting that longer observation could point to advantages of using herbal medicine.

Some degree of inflammation is necessary; however, if intense is harmful, by impairment the microcirculation and also inhibiting cell proliferation (formation of fibroblasts)⁹. In relation to fibroblast proliferation, there was a significant difference at 14 days between the GC and GB, favoring GC and GA, similar to other studies^{24,25}. Another important variable to assess the chronic inflammatory process is fibrosis or collagen deposition. In this research it was observed to be missing/discrete in all groups without significant difference; other studies showed significant differences favoring the experimental group^{26,27} and, in contrast, favoring the GC²⁴. In the gastrointestinal tract, collagen is synthesized by fibroblasts and smooth muscle cells, predominantly in the scar collagen type I (68%) that plays an important role in the tensile strength of the scar¹⁸. In this study it was evaluated quantitatively, showing significant difference at 14 days, confirming prior studies^{27,30}.

Several are the medicinal actions of aroeira and babassu. Continuity of experimental studies must be based on the use fractions of these herbal, as well as use of other routes of administration, different dosages, longer observation and use of immunohistochemistry in order to better assess their healing actions.

Conclusion

Hydroalcoholic extract of *Schinus terebinthifolius* Raddi (aroeira) and the aqueous extract of *Orbignya phalerata* Mart. (babassu) favored the healing process in cecorrahphy in rats.

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