European Journal of Medicinal Plants

Volume 34, Issue 1, Page 1-11, 2023; Article no.EJMP.94843 ISSN: 2231-0894, NLM ID: 101583475

Evaluation of the Anticonvulsant and Antidepressant Effects of the Aqueous Extract of the Leaves of Ascotheca paucinervia (T. Anderson ex C.B. Clarke) Heine in Mice

Bassoueka D'Avila Judicaël^{a*}, Ondele Radar^a, Omeka Ngassaki Gelvid^a and Abena Ange Antoine^b

 ^a Laboratory of Pharmacodynamics and Experimental Physiopathology (L2PE), Faculty of Sciences and Technology, Marien Ngouabi University, BP: 69, Brazzaville, Congo.
^b Biochemistry and Pharmacology Laboratory, Faculty of Health Sciences, Marien Ngouabi University, BP: 69, Brazzaville, Congo.

Authors' contributions

This work was carried out in collaboration among all authors. All authors read and approved the final manuscript.

Article Information

DOI: 10.9734/EJMP/2023/v34i11115

Open Peer Review History:

This journal follows the Advanced Open Peer Review policy. Identity of the Reviewers, Editor(s) and additional Reviewers, peer review comments, different versions of the manuscript, comments of the editors, etc are available here: https://www.sdiarticle5.com/review-history/94843

> Received: 21/10/2022 Accepted: 24/12/2022 Published: 02/01/2023

Original Research Article

ABSTRACT

The present study was undertaken to evaluate the anticonvulsant and antidepressant effects of *Ascotheca paucinervia* leaves on mice by using strychnine at 2.5mg/kg to induce convulsions and the forced swimming test to create a stressful situation, respectively. Concerning convulsions, only the 500mg/kg extract significantly increases (p<0.001) the time to onset of convulsions and it non-significantly reduces the duration of convulsions induced by strychnine. In addition, the extract

Euro. J. Med. Plants, 34, no. 1, pp. 1-11, 2023



^{*}Corresponding author: E-mail: basdavila@gmail.com;

reduces very significantly in a dose-dependent manner the time of immobility and it significantly increases the swimming time as well as the climbing time at both doses. At the same time, the estimation of the acute toxicity of the extract from the leaves of Ascotheca paucinervia according to guideline No. 425 of the OECD (2022) shows that the latter is weakly toxic and its LD50 is greater than 5000mg/kg. In addition, the evaluation of the sedative effect of this extract shows that it produces a dose-dependent sedative effects and at doses of 250m/kg and 500mg/kg, the extract significantly potentiates the sleep induced by phenobarbital. In summary, the results obtained suggest that Ascotheca paucineervia leaves extract possesses anticonvulsant and antidepressant effects.

Keywords: Anticonvulsant activity; antidepressant activity; epilepsy; Ascotheca paucinervia.

1. INTRODUCTION

Convulsions are seizures, being the clinical expression of which results in more or less violent and involuntary muscular contractures of one or more muscles, of one or more limbs or even of the whole body, of cerebral or medullary origin, provoked by "the hyperexcitation of a neuronal group [1]. Unlike other seizures, epileptic seizures are characterized by the occurrence of two or more spontaneous seizures manifested by brief episodes of involuntary tremors affecting a part of the body (focal seizures) or the whole body (generalized seizures) and they result from excessive electrical discharges in a group of brain cells [2]. Epilepsy suffers from a very negative brand image and is responsible for the suffering of approximately 50 million people worldwide, of whom 5 million are diagnosed each year, thus representing the second most common neurological disorder after migraine and with a mortality from 2 to 3 times higher than that of the normal population whose most common cause is sudden unexpected death. However, several psychiatric comorbidities are often associated with epilepsy, the most common of which is depression [3]. It is estimated that 6 to 30% of epileptics are affected by major depressive disorders [4,5]. More than 80% of the African population uses traditional medicine and medicinal plants for their primary health care [6]. Ascotheca Paucinervia is among one of the medicinal plants which treat epilepsy. Recently, an identified species in the Republic of Congo confirms that this plant has multiple virtues which can be scientifically tested. In African countries, 80% of the population uses traditional medicine and medicinal plants for their primary health care, because some of these plants contain

secondary metabolites with little or no exploration of effective activities, little or non-toxic [7] (which can constitute original series leads for the development of new drugs [8] and which remain an inexhaustible reservoir of new drugs. Indeed, numerous studies mention that approximately 400,000 plants species have been identified [9] but only 2,000 to 3,000 of them have been the subject of scientific, chemical or pharmacological studies [10] including Ascotheca paucinervia. Ascotheca paucinervia is a perennial plant up to one meter high, with young branches and petioles covered with brown pubescence and terminal inflorescences often axillary in frequently branching spikes [11]. Thus, Ascotheca paucinervia (T. Anderson ex C.B Clarke) Heine is a species of plants of Acanthaceae family and of the genus Ascotheca was before recognized in West Africa and Gabon. Likewise, it is a plant of the peatland zone of the large forests and can be domesticated easily by creating the necessary conditions. In addition, we found this plant in the "Cuvette department", Makoua district, precisely in the forest of the "Cité de Feu village" called Bamralondo. Traditionally, Ascotheca paucinervia is used in fishing for its ichthyotoxic effects and in traditional medicine where its use is still the prerogative of ancient traditional healers to treat epileptic seizures. The current work evaluates the anticonvulsant and antidepressant effects of Ascotheca paucinervia leaves in mice.

2. MATERIALS AND METHODS

This study was carried out in the laboratory of Pharmacodynamics and experimental physiopathology of the Faculty of Sciences and Techniques of Marien University - NGOUABI.

2.1 Plant Material

The plant material consisted of the leaves of *Ascotheca paucinervia* harvested in January 2021 in the "Cuvette" department, in Makoua district precisely in the forest of the village of Feu (48 km) and dried at room temperature at the Laboratory of Pharmacodynamics and Experimental Pathophysiology (L2PE).

2.2 Animal Material

The animal material was made up of mice of the Swiss albino strain with a body weight of between 20 and 25 g, of male and female sexes, provided by the IRSSA, reared at the animal facility of the Faculty of Science and Techniques under standard conditions ($25\pm5^{\circ}$ C, 40-70 HR), with a cycle of 12 hours of light and 12 hours of darkness. These mice had free access to tap water and standard food.

2.3 Preparation of the Aqueous Extract of Ascotheca paucinervia Leaves

The aqueous extract of the leaves of Ascotheca paucinervia was prepared by decoction at 10%, 50 g of powder of Ascotheca paucinervia leaves and mixed in 500 ml of distilled water and boiled for 15 minutes at 75°C. After cooling, the decoction obtained was filtered with absorbent collected cotton. The filtrate was then evaporated at reduced temperature (50-60°C.) for 48 hours. The dry extract obtained was used for acute toxicity evaluation and pharmacological tests.

2.4 Acute Toxicity Study

The acute toxicity of the aqueous extract of Ascotheca Paucinervia leaves was carried out according to the guideline N°425 of the OECD [12]. It consists of testing the aqueous extract orally at a single dose of 5000 mg/kg. The test was carried out on 6 female mice divided into two groups of 3 mice each. The first group of which (negative control) received distilled water at 0.5ml/100g and the second one received the extract at 5000mg /kg. After administration of products, the mice were placed in individual cages for observations for 4 h. These observations were related to parameters such as tremors, alertness, vocalization, stool state, of urine, reaction quantity to stimuli, aggressiveness and sleep. The mortality rate of animals per group was evaluated for 48 hours after administration of products. Each animal's body weight, water intake and food consumption were measured daily for 14 days.

2.5 Pharmacological Tests

2.5.1 Evaluation of the effects of Ascotheca paucinervia leaves against strychnineinduced seizures in mice (STR)

The method used was developed by Lehmann et al. in 1988 and taken up by Bassoueka (2016). It consists in inducing tonic convulsions within 10 minutes in mice by intraperitoneal administration of strychnine 2.5 mg/kg for one hour after all treatments. Four (4) groups of 5 mice each were formed and fasted for 18 hours before the experiment and treated as follows: Negative control group 1 received distilled water 0.5 mL/100 g, per-os. Positive control group 2 was treated with the reference molecule of diazepam 10 mg/kg, per os. Groups 3 and 4 were treated with the aqueous extract of Ascotheca paucinervia leaves at the respective doses of 250 and 500mg/Kg per os. One hour after convulsions treatments, were induced by intraperitoneal injection of strychnine 2.5 mg/kg. The animals were observed for 10 minutes. Those not presenting convulsions or presenting convulsions without dying during this period, were declared protected. The time to onset as well as the duration of seizures in each batch are determined [13].

2.6 Evaluation of Antidepressant Effects of *Ascotheca paucinervia* Leaves in Mice

The forced swimming test was applied according to the protocol established by [8], with some modifications. Four (4) groups of 4 mice were constituted and treated orally as follows: Group 1, negative control received distilled water 0.5 ml/100 g. Group 2, positive control, was treated with the reference molecule of Clomipramine at a dose of 15mg/kg. Groups 3 and 4 were treated with aqueous extract of Ascotheca Paucinervia leaves at doses of 250 mg/kg and 500 mg/kg respectively. One hour (1h) after the gavage, the mice were placed individually in a high glass cylinder 25cm with 20cm of diameter, containing 15cm of water, maintained at a temperature of ± 25°C for a period equivalent to 6 minutes. During this observation period, the durations of climbing, swimming and immobility are timed [14,15,16].

2.7 Evaluation of Sedative Activity of Aqueous Extract of Ascotheca paucinervia Leaves

2.7.1 Effect of aqueous extract of Ascotheca paucinervia leaves on motor activity

The method used was developed by Boissier and Simon in 1967 and taken up by Bassoueka (2016). It consists in assessing by using a cage with squared boards comprising 16 squares measuring 40×40 cm, based on the number of squares covered by a mouse in five (5) minutes. Four Groups (4) of 5 mice each were constituted and treated orally as follows: Group 1 of negative control received distilled water 0.5 mL/100 a. Group 2 of positive control was treated with the reference molecule of diazepam 10 mg/kg. Groups 3 and 4 were treated with the aqueous extract of Ascotheca paucinervia leaves at the respective doses of 250 and 500mg/Kg p.c. One hour after the administration of products, the animals were placed in turn in a squared cage, and the number of squares crossed by them after five (5) minutes is noted.

2.8 Effect of Aqueous Extract of Ascotheca paucinervia Leaves on Phenobarbital-Induced Sleep

The method used was developed [17]. By Lechat et al in 1964. Four (4) groups of 5 mice each were formed and fasted for 18 hours before the experiment and treated as follows. Negative control group 1 received distilled water 0.5 mL/100 g, per-os. Positive control group 2 was treated with the reference molecule of diazepam 10 mg/kg, per os. Groups 3 and 4 were treated aqueous with the extract of Ascotheca paucinervia leaves at the respective doses of 250 and 500mg/Kg per os. One hour after the administration of various products, the animals received phenobarbital intraperitoneally at a dose of 10mg/kg and the time to onset and sleep duration were determined for each mouse. Sleep duration is considered as the time between when the mouse loses the righting reflex and when the righting reflex reappears. The loss and/or appearance of the reflex is assessed by tickling the mouse's ear. The awake mouse reacts by moving the front paw on the stimulated side [18].

2.9 Determination of the Chemical Profile of Ascotheca paucinervia Leaves

To identify the different chemical groups or secondary metabolites (alkaloids, flavonoids,

steroids. tannins. terpenes. anthocvanins. anthraquinones, and holosides, saponosides, mucilages), a chemical screening of the dry leaves of this plant was carried out. For this, it was used classical phytochemical tests based on coloring and precipitation reactions. The different categories of secondary metabolites sought are chosen according to the common chemical nature shared by the major active principles to which they give access and their use in therapy. To search for alkaloids, 5 mL of aqueous extracts was placed in a test tube, adding 1 mL of 1N hydrochloric acid and a few drops of Dragendorff's or Mayer's reagent. In the presence of the alkaloids, a red precipitate is formed. So, for searching tannins, 5 mL of the aqueous extract contained in a test tube is used and 1 mL of an aqueous solution of iron III chloride is added. In the presence of tannins, a areenish or blue-blackish color is developed. Flavonoids are found from a test tube where 5 mL of extract solution is added, then 5 mL of hydrochloric acid (HCl), 1 mL of iso-amyl alcohol and some magnesium shavings are added as well. The appearance at the level of the supernatant layer of iso-amyl alcohol of a coloration can be determined from orange-pink which indicates the presence of flavones, purplish pink characterizes the flavanones, red indicates the presence of flavanols. To search for Anthocyanin in a test tube, 5 mL of the aqueous extract 3 mL of hydrochloric acid 20% is added as well. The appearance of a dark pink color when cold and orange-red is observed, hot indicates that the test is positive. For mucilages, 1 mL of the 10% decoction was essential in a test tube and 5 mL of absolute alcohol was added by obtaining a flaky precipitate ten (10) minutes later which indicates the presence of mucilage. For free anthraguinones, it is focused on 2.5 mL of the aqueous extract, with addition of 1 mL of 10% sodium hydroxide (NaOH) and the test is positive when a red color appears. While, steroids and terpenoids are searched from a test tube where 1 ml of acetic anhydride is added to 5 mL of the aqueous extract (non-chloroform) and a few drops of concentrated sulfuric acid are allowed to run down the wall of the test tube. The appearance of a green to red or purplish-blue color indicates that the test is positive.

2.10 Statistical Analyzes

Statistical analysis was done by using Excel software (Office 2010) and the results expressed as mean \pm SEM were subjected to a one-way analysis of variance followed by the Student-

Fischer t-test (p < 0.05, p < 0.01, p < 0.001) (Schwartz D.E., 1963).

3. RESULTS

3.1 Assessment of Acute Toxicity

After administration of a single dose of 5000 mg/kg of the aqueous extract of Ascotheca paucinervia leaves orally and observation for four hours (04 h), the general behavior of the treated animals in comparison with that of the controls remains normal, except at the level of a few parameters where there were certain changes at a given moment of the observation. Thus, it is from the fourth hour of observation of the animals that it was noted a reduction in mobility, vigilance as well as an appearance of light sleep in the group treated with the aqueous extract of Ascotheca paucinevia. In the end, no death was recorded in 48 hours and the notation of the water consumption, food and the weight of the animals were carried out for 14 days.

3.2 Effect of the Aqueous Extract of *Ascotheca paucinervia* on Weight Change

Fig. 1 shows the change in weight after single administration of the aqueous extract of *Ascotheca paucinevia* leaves orally at a dose of 5000 mg/kg of body weight for fourteen days (14 days). This presents three levels (03) of significance: a significant increase (p<0.001) in body mass on the second day (D2) of the

weighing, as well as on the third day (D3), the fifth day (D5) and the eighth (D8) day (p<0.01) and fourth (D4), sixth (D6) and seventh (D7) day (p<0.05). In general, the extract significantly increases body mass compared to distilled water.

3.3 Effect of Aqueous Extract of *Ascotheca paucinervia* on Food Consumption

Fig. 2 shows the effects of *Ascotheca paucinervia* leaves extract on food consumption after oral administration of the extract at a single dose of 5000 mg/kg body weight. The control and treated groups were fed with the standard food, i.e. 100 g for each group for fourteen (14) days. The amount of food remaining every two days was weighed and that consumed calculated. This figure shows an increase in the food consumption of the treated batch on the following days: D2, D4, D10 and D14 compared to the negative control.

3.4 Effect of Aqueous Extract of Ascotheca paucinervia on Water Consumption

Fig. 3 shows the effects of the aqueous extract of *Ascotheca paucinervia* leaves on water consumption after administration of aqueous extract of *Ascotheca paucinevia* leaves orally at a single dose of 5000 mg/kg. The control groups were treated and fed tap water for fourteen days. This shows an increase in water consumption on days: D0, D4, D12 and D14.

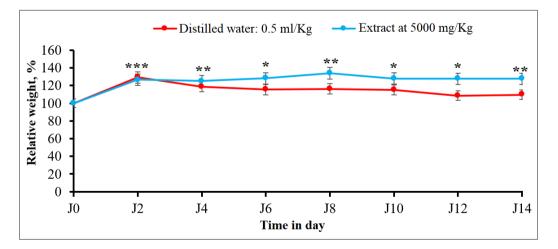
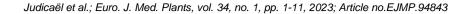
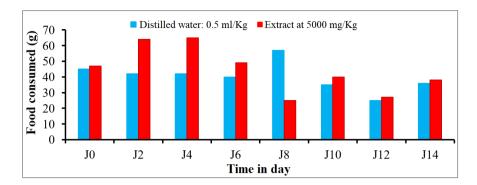


Fig. 1. Effect of the aqueous extract of Ascotheca pauinervia leaves on the evolution of body weight

The results are expressed as mean ± ESM error, n=3, *p<0.05, **p<0.01, ***p<0.001







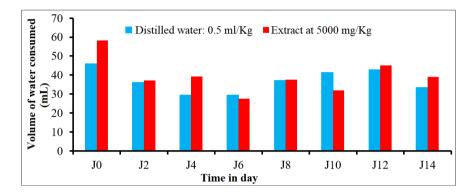


Fig. 3. Effects of the aqueous extract of Ascotheca paucinervia leaves on water consumption

3.5 Effect of Aqueous Extract of Ascotheca paucinervia Leaves against Strychnine-Induced Seizures

Table 1 shows the effects of the aqueous extract of the leaves of Ascotheca paucinervia at doses of 250 and 500 mg/kg of body weight against convulsions induced by strychnine. Compared to distilled water, at a dose of 250 mg/kg, the extract non-significantly increases and decreases the time to onset of convulsions and the duration of convulsions respectively. At a dose of 500 mg/kg, the extract very significantly increases the delay as well as it decreases the duration of convulsions in a non-significant way. Diazepam 10mg/kg used as a reference molecule led to a significant increase (p<0.001) in the time to onset of convulsions similar to that of the aqueous extract of Ascotheca paucinervia leaves at a dose of 500 mg/kg of body weight.

3.6 Antidepressant Effect of Aqueous Extract of Ascotheca Paucinervia Leaves in Mice

Table 2 shows the effects of *Ascotheca paucinervia* leaves extract on antidepressant activity in the forced swimming model. At a dose

of 250 mg/kg, the extract significantly reduces and increases (p<0.05) the immobility time, the climbing time and the time of swimming as well. At a dose of 500 mg/kg, the extract very significantly reduces and increases the time of immobility and the climbing time respectively, as well as it increases the swimming time in a nonsignificant way. Clomipramine used as a positive control produced effects to the aqueous extract of *Ascotheca paucinervia* leaves.

3.7 Evaluation of the Sedative Effect of the Aqueous Extract of Ascotheca paucinervia Leaves in Mice

3.7.1 Evaluation of the effect of the aqueous extract of *Ascotheca paucinervia* leaves on locomotor activity in mice

Table 3 shows the effects of aqueous extract of *Ascotheca paucinervia* leaves at doses of 250 and 500 mg/kg on locomotor activity. This shows that in comparison with distilled water, the aqueous extract of leaves of *Ascotheca paucinervia* significantly reduces the number of squares crossed at doses of 250 and 500 mg/kg and the effects are more pronounced at a dose of 500 mg/kg. Diazepam 10 mg/kg used as a

reference molecule caused a reduction in the number of squares crossed greater than that caused by the aqueous extract of *Ascotheca paucinervia* leaves at doses of 250 and 500 mg/kg of body weight.

3.8 Effect of Aqueous Extract of *Ascotheca paucinervia* Leaves on Barbiturate Sleep in Mice

Table 4 shows the effects of aqueous extract of *Ascotheca paucinervia* leaves at doses of 250 and 500 mg/kg on barbiturate sleep. Compared to distilled water; at a dose of 250 mg/kg of the aqueous extract, significantly decreases and increases respectively the time to onset and the duration of sleep. At a dose of 500 mg/kg of the aqueous extract, significantly decreases and increases respectively the time to onset and the duration of sleep. At a dose of 500 mg/kg of the aqueous extract, significantly decreases and increases respectively the time to onset and the

duration of sleep. Diazepam 5 mg/kg, used as a reference molecule, produced a more significant effect than that produced by the extract of *Ascotheca paucinervia* leaves.

3.9 Determination of Chemical Profile of Ascotheca paucinervia Leaves

Table 5 shows results of Chemical screening of *Ascotheca paucinervia* leaves

The chemical screening of aqueous extract of leaves of *Ascotheca paucinervia* revealed that the leaves of *Ascotheca paucinervia* contain six (06) chemical groups including three (03) in abundance (mucilages, anthocyanin, terpenes and steroids), two (02) few abundant (flavonoids and tannins) and one (01) in the form of traces (alkaloids).

Products	Doses (mg/kg)	Time to onset of seizures (Seconds)	Duration of seizures (Seconds)
Distilled water + Strychnine	0.5	85.75 ± 10.19	20± 4.81
Diazepam + Strychnine	10	207 ± 10.40 ***	342.5 ± 15,96 ***
A.paucinervia + Strychnine	250	101 ± 9.66 NS	39.5± 15,51 NS
A.paucinervia + Strychnine	500	201.25± 85.36***	23± 2.67 NS
The values are means + SEM	with $n = 4$: *** $n < 0$	001: significant difforance fr	rom and NS: non significant

The values are means \pm SEM; with n = 4; *** p < 0.001; significant difference from and NS: non-significant difference from negative control

Table 2. Antidepressant effect of aqueous extract of Ascotheca paucinervia leaves in mice

Treatments	Doses (mg/kg)	Immobility time	Climbing time	Swimming time
Distilled water	0.5	170 ± 10.33	85.75±10.01	102.75 ± 21.11
Clomipramine	15	70.5 ± 11.89 ***	200.75±34.29 **	69 ± 21.74 ^{NS}
A.paucinervia	250	99.75 ± 21.10 *	134.4 ± 18.92 *	61 ± 5.93 ^{NS}
A.paucinervia	500	47 ± 25.31 ***	190.75 ± 25.91 **	134.25 ± 28.11 ^{№\$}

Values are means ± SEM; with n=4; *p<0.05; **p<0.01; *** p < 0.001 significant difference from and NS: nonsignificant difference from negative control

Products	Doses (mg/kg)	Number of squares crossed (5 min)
Distilled water	0,5 (a)	119,40 ± 9,38
Diazépam	10	23,66 ± 3,17 ***
Ascotheca paucineria	250	80 ± 11,23 *
	500	37,66 ± 1,76 **

(a): in ml/kg; Values are means ± SEM; with n=5; *p<0.05; **p<0.01; *** p < 0.001 significant difference from

Table 4. Effects of aqueous extract of A. paucinervia leaves on barbiturate sleep

Products	Dosage (mg/kg)	Time to onset sleep time (min)	Duration of sleep (mins)
Distilled water + Ph (b)	0,5 (a)	38,33 ± 5,20	416,6 ± 22,82
Diazepam + Ph (b)	5	9,66 ± 0,31 ***	1360,2 ± 35,81 ***
A.paucineria + Ph (b)	250	18,66 ± 2,96 *	552,8 ± 14,04**
A.paucineria + Ph (b)	500	14,66 ± 1,20 *	501,4 ± 11,30*

Chemical groups	Results
Anthraquinone	-
Alkaloids	+
Flavonoids	++
Tannins	++
Anthocyanin	+++
mucilage	+++
Terpenes and Steroids	+++

Table 5. Chemical screening of Ascotheca paucinervia leaves

-: Absent; +: trace; ++: scarce; +++: Abundant

4. DISCUSSION

This present study made it possible to evaluate the anticonvulsant and antidepressant effects of the aqueous extract of Ascotheca paucinervia leaves at doses of 250 and 500 mg/kg of body weight in mice. In order to establish the safety of the aqueous extract of Ascotheca paucinervia leaves, the estimation of acute oral toxicity was carried out in accordance with OECD guideline No. 425 [12]. According to the latter, the level of overt toxicity is reached when at least one animal shows one of the following signs (from the day following the treatment): tremors, lethargy, irregular breathing or weight loss greater than 10% compared to the weight before treatment. However, the analyses of parameters highlighted to evaluate the acute toxicity of this extract at 5000 mg/kg in comparison with those of distilled water at 0.5ml/100g are globally identical during the four (04) hours of observation. Thus, this would suggest that the lethal dose 50 (LD50) of this extract would be greater than 5000 mg/kg because, no case of death was recorded within 48 hours as well as in the continuation of the observation (14 days) and according to the Globally Harmonized Classification System the latter could be classified in category 5 of LD50 between 2000 and 5000mg/Kg [12]. In addition, it noticed a little later (from 4 a.m.) some changes, in particular the reduction in mobility, the reduction in alertness, the appearance of very light sleep. These results could probably be attributed to the sedative or hypnotic effects of this extract. In addition, the analysis of weight change shows a significant increase in the weight of animals in the treated group. In relation to the quantities of food and water consumed, this weight increase suggests that this extract would have virtues on digestion particularly, it would be an appetite stimulant [18]. Our results are similar to those obtained by who worked on the aqueous extract of Cymbopogon densiflorus [19-23]. The anticonvulsant activity of the extract

studied usina strvchnine was as а convulsingenic agent, this substance produces spinal convulsions. The injection of strychnine into the animals first induces hyperexcitability, an increase in reflexes and then convulsions. Indeed, at doses of 250 and 500 mg/kg, the aqueous extract of the leaves of Ascotheca paucinervia increases respectively in a nonsignificant and very significant way (p<0.001) the time to onset of convulsions induced by strychnine and the significance of the extract at a dose of 500 mg/kg would appear to be equal to that of diazepam [24-28]. Likewise, terpenoids have been shown to activate the enzyme glutamic decarboxylase, confirming the potential of terpenoids for the treatment of epileptic seizures [29-34]. Concerning the duration of the convulsions, at doses of 250 mg/kg and 500 mg/kg the aqueous extract of the leaves of Ascotheca paucinervia insignificantly decreases the duration of the convulsions induced by strychnine. The inability of the aqueous leaf extract of Ascotheca paucinervia to significantly decrease the duration of strychnine-induced seizures suggests that the extract possesses a lower affinity than strychnine for glycinemediated inhibitory neurotransmission receptors. These results differ from those obtained by Bassoueka et al. [13] who worked on the anticonvulsant effect of the aqueous extract of the leaves of Crossopterys febrifuga. The forced swimming test (FST) was proposed by [8]. Thus, at doses of 250 and 500 mg/kg, the aqueous extract of the leaves of Ascotheca paucinervia leaves reduces respectively highly (p<0.01) and very highly significantly (p<0.001) the immobility time in mice. At a dose of 500 mg/kg, the effect of Ascotheca paucinervia leaf extract appears to be slightly greater than that of clomipramine, a molecule scientifically recognized as an antidepressant. In addition, the non-significant increase in swimming time and highly significant increase in climbing time of the treated batch suggests that the extract would act by using serotonergic and noradrenergic receptors [35-37]. Effects could be attributed to tannins which are present since, tannic acid has been shown in many published papers to be a non-selective MAO inhibitor, which increases levels of monoaminergic neurotransmitters [38,39]. It should be noted that the effects are more accentuated at the higher dose, which would suggest that the effects could be dosedependent. Results obtained are in agreement with those obtained by [40] who worked on the aqueous extract of cassia alata. Indeed, flavonoids, tannins, terpenes, anthocyanins and

mucilages could be responsible for the antidepressant effects of the extract [41]. Similarly, anticonvulsant effects of this extract could be attributed to terpenes and flavonoids [42-44].

5. CONCLUSION

The aqueous extract of Ascotheca paucinervia leaves is weakly toxic up to 5000 mg/kg in a single dose and LD50 > 5000 mg/kg. It significantly increases the time to onset of seizures and non-significantly decreases the duration of seizures in mice. Then, it very significantly reduces immobility time and increases swimming and climbing times in a nonsignificant and very significant way in mice, respectively. A dose-dependent reduces the mobility of mice where significantly decreases and increases respectively the time to onset and the duration of sleep. Classic chemical screening revealed the presence of flavonoids, terpenes, steroids, alkaloids, tannins, mucilages and anthocyanins.

CONSENT

It is not applicable.

ETHICAL APPROVAL

It is not applicable.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

REFERENCES

- VIDAL. Vidal. Accesson 2021 Available:www.vidal.fr>epilepsy>symptom s>: www.vidal.fr
- 2. Jost J. The determinants of the therapeutic deficit of epilepsy: Place of the quality of antiepileptics in sub-Saharan Africa, doctoral thesis, University of Limoges: Tropical Neuroepidemiology. 2018;306.
- WHO Action against epilepsy: A public health imperative, Geneva, CCBY-NC-SA3.0IGO; 2019.
- 4. Rasekhi TR, Kathryn ND, Joely AM, Nei M, Sprling MR, Asma B, Donmez M. Improving the prediction of sudden

unexpected death in epilepsy: SUDEP-7 to SUDEP-3, epilepsia, 2021;62: 1536-1545.

- 5. Guekht A, Brodie M, Secco M, Li S, Volkers N, Wiebe. The road to World Health Organization global action plan on epilepsy and neurological disorders, Epilepsia. flight. 2021; 62(5):1057-1063.
- 6. WHO. Main landmarks of epilepsy, World Health Organization. 2018.
- 7. Hingray C Biraben A psychiatric comorbidities and epilepsy, European psychiatry. 2015;30(8):S76.
- Kougan Nkwokap GB. Isolation and characterization of saponosides from three plants of the Araliaceae and Dracaenaceae families and evaluation of their cytotoxic activities on tumor cells, Cosupervised thesis for obtaining the degree of Doctor of the University, University of Burgundy. 2010;184.
- 9. Bachman S. State of the World's Plants Report, Royal Botanic Gardens, Kew. 2016;7184
- VIDAL. 2012 What are the origins of phytotherapy? Available:www.vidal.fr>originsphytotherapie Accesson August 2022).
- 11. Dobillard A, Chaletain C. Synonymous and bibliographic index of plants from North Africa. 2013;1-5.
- 12. OECD (2022) Test No. 425: Acute Oral Toxicity: Dose Adjustment Method, OECD Guidelines for the Testing of Chemicals, Section 4, OECD Publishing, Paris.
- Bassoueka DJ, Loufoua BAE, Etou-Ossibi AW, Nsondé-Ntandou, Ondelé R, ElionItou RDG, Ouamba JM, Abena AA. Anticonvulsant plants from the Congo, ethnobotanical approach, Phytotherapy. 2015;13:298-305.
- 14. Porsolt RD, Le Pichon M, Jalfre M. Depression: A new animal model sensitive to antidepressant treatments. Nature. 1977;266:730-732.
- 15. Chen L, Faas GC, Ferando I, Mody I. Novel insights into the behavioral analysis of mice subjected to the forcedswim test. Translational Psychiatry. 2015;5:1-9.
- Gong Y, Han T, Chen W, Dib HH, Yang G, Zhuang R, Chen Y, Tong X, Xiaoxy Yet al. Prevalence of anxiety and depression symptoms and related risk factors among physicians in China: A Cross-Sectional study. PLOS. 2014;1:32-42.

- Dhenain Marc. Primate models and therapeutic innovations against diseases of the central nervous system. 2020; 34-37.
- Nkundineza JC, Nsonde Ntandou GF, Bassoueka D'AJ, Boumba LS, Makambila MC, Abena AA. Anticonvulsant and Sedative Effects of Cassia alata (Fabaceae) in Mice. Galore International Journal of Health Sciences and Research. 2020;5(1):28-37.
- Reus-García MM, Sánchez-Campusano R, Ledderose J, Dogbevia GK, Treviño M, Hasan MT, Gruart A, Delgado-García JM. The claustrum is involved in cognitive processes related to the classical conditioning of eyelid responses in behaving rabbits. Cereb Cortex. 2021;31(1):281-300.
- CATALA A. characterization and training methods of assistance dogs to help people with epilepsy, doctoral thesis, University of Biology Bretagne Santé Loire, Rennes. 2019;1:235.
- 21. Ken Ho. Steroid medicine reduces functions of calprie-burning brown fat, EDO; 2016.
- 22. Cryan JF, Leonard BE. Depression: From Psychopathology to Pharmacotherapy. Switzerland: Karger. 2010;207-208.
- 23. Seibert JB, Viegas JS, Almeida TC, Amparo TR, Rodrigues IV, Lanza JS, Frézard FJ, Soares RD, Teixeira LF, de Souza GH, Vieira PM. Nanostructured systems improve the antimicrobial potential of the essential oil from *Cymbopogon densiflorus* leaves. Journal of Natural Products. 2019;82(12):3208-20.
- 24. Ahangar N, Mirfetros S, Ebrahimzadeh MA. Antidepressant activity of polyphenol fraction of Artimisia absinthium L. Pharmacologyonline, 2011;1:825-832.
- 25. Pocket Atlas of Anatomy, The Nervous System The Sense Organs, 5th Edition, La Voixier Medicine; 2015.
- 26. Begum A, Hossen A, Moly A, Bhuiyan M., and Shahed-Al-Mahmud, M. In Vivo Sedative and Anxiolytic Activities of Thunbergia erecta (Acanthaceae) Leaves Activate Gamma-Aminobutyric Acid (GABA) Mediated Hyperpolarization in Swiss albino mice. Pharmacology and Pharmacy, 2019;10:177-193
- 27. Brain Facts. The Neuron ; 2012.
- 28. Catala A. characterization and training methods of assistance dogs to help people with epilepsy, doctoral thesis, University of

Biology Bretagne Santé Loire, Rennes. 2019;1:235.

- 29. Graziella Cara. Discovery of a new mode of communication of the cells of our brain, presse.inserm.fr; 2021.
- 30. Hritcu L, Ionita R, Postu PA, Gupta GK, Turkez H, Lima TC. Anthocyanin on depression mice by increasing monoamine Neurotransmitter and up-regulating BDNF expression. J Function. 2017;1-18.
- 31. IBE. Annual report, 2019;26
- Kurada L, Bayat A, Joshi S, Chahine A, Koubeissi MZ. Antiepileptic effects of electrical stimulation of the piriform cortex. Exp Neurol. 2020;325.
- 33. Labrakakis C, Rudolph U, De Koninck. The heterogeneitv in GABAA receptor-IPSC mediated reflects Kinetics heterogeneity of subunit composition among inhibition and excitatorv interneurons in spenal lamina II. Frontiers in cellular Neuroscience. 2014;8(424):1-12
- 34. Michel N, Malvyne RD. the enteric nervous system and the digestive neuro-glioepithelial unit, Bulletin of the French Veterinary Academy. 2013;166(1):7-12.
- Olie JP. Psychotropic treatments. In: Guelfi JD, Rouillon F. Manuel de psychiatrie. 2nd edition. Paris: Elsevier Masson. 2012;571-574.
- Palazzolo J. Depression. In: Depression and anxiety: Understanding them better to better manage them. Paris: Elsevier Masson. 2007;5-60.
- Pfieger FW and Reber M. New insight into the mechanisms of neuron-glia communication, Med Sci (Paris). 2013;29(2):142-1444
- Rahman MM, Ichiyanagi T, Komiyama T, Sato S, Konishi T, 2020, Antidepressant flavonoids and their relationship with
- Stahl SM. Mood disorders. In: Essential psychopharmacology: Neuroscientific bases and practical applications. 2nd edition. Paris: Lavoisier. 2010;453-510.
- Sundaram R, Gowtham L, Nayak.BS. The 40. role of excitatory neurotransmitter glutamate in brain physiology and pathology, Asian Journal of Pharmaceutical and Clinical Research. 2012;5(2):1-7
- 41. Tchaleu Nguenkam BC, and Raharivelo A. Epidemiological profile of epilepsy in Central Africa and Madagascar, Medicine of Black Africa, August/September; 2011.

Judicaël et al.; Euro. J. Med. Plants, vol. 34, no. 1, pp. 1-11, 2023; Article no.EJMP.94843

- 42. Tricoire L, Hepp R, Lambolez B. The delta family of glutamate receptors, Med Sci (Paris). 2018;34:662-664.
- Yadav R, Subhash CG, Brandron GH, Bhatt JM, Stairs DJ, Shashank MD. Deletion of Glutamate Delta-1 Receptor in Mouse Leads to Aberrant Emotional and

Social Behaviors, PloS ONE. 2012;7(3): e32969:1-13

44. Yuzaki M. Two Classes of secreted synaptic organizers in the central nervous system. Ann. Rev. Physiol. 2018;80:243-262.

© 2023 Judicaël et al.; This is an Open Access article distributed under the terms of the Creative Commons Attribution License (http://creativecommons.org/licenses/by/4.0), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

Peer-review history: The peer review history for this paper can be accessed here: https://www.sdiarticle5.com/review-history/94843