

Host status of medicinal plants to root-knot nematodes

.ABSTRACT

Medicinal plants are described such as those produce substances capable of provoking reactions in the human body leading to the cure of diseases. Like as cultivated species, medicinal plants can be attacked by various pests and diseases, affecting the qualitative and quantitative characteristics of their curative properties, as well as productivity. Phytonematodes are one of the main factors limiting the productivity of cultivated plants. In medicinal species this pathogens group have been caused damage in the sanity of the plants interfering in the quality of the compounds produced. Among them, due to the high parasitism degree, the species of the genus *Meloidogyne*, popularly known as root-knot nematodes. Among the management strategies of these phytopathogens, biological and cultural controls have low efficiency reports. Likewise, chemical control is not indicated due to the high cost of the products, besides the high toxicity and risk of environmental pollution. Therefore, the most effective control method is the use of resistant species or cultivars. Once these species are identified, they can be used as antagonists or incorporated into the soil, aiming to decrease the nematode population in infested areas. The use of resistant medicinal species allows little or no reproduction of *Meloidogyne* spp. in the field, providing effective control. Other characteristics are the reduction of production costs, and the protection of the environment against pollution caused by chemical waste.

Keywords: Meloidogyne incognita; Meloidogyne javanica; Meloidogyne enterolobii; Phytonematodes; Parasitism; Resistance sources; Gall; Traditional medicine; Herbal medicines

1. INTRODUCTION

Plants have been used as medicines for thousands of years in the form of raw drugs such as tinctures, teas, poultices, and other herbal formulations. Medicinal plants can be described as any plant which has in one or more organs substances that can be used for therapeutic purposes or that are precursors of semisynthetic drugs [1]. Knowledge about the use of medicinal plants has been passed down through generations among traditional populations. Over the years, the use of plants as a drug began to be carried out from the isolation of active compounds, starting with the isolation of morphine and opium in the early 19th century [2]. The isolation and pharmacological characterization of the active compounds of medicinal plants to be continue carried until the present day.

The consumption demand of medicinal plants and herbal medicines has increased worldwide. The World Health Organization (WHO) reports about 80% of the world population makes use of some type of medicinal herb for the treatment of the most diverse diseases [3]. The use of medicinal plants has even received incentives from WHO itself. It is estimated that the market for natural products has increases of approximately 22% per year [4]. Among the factors contributed to this increase are the high cost of synthetic drugs and the

34 resistance of pathogens to synthetic drugs. This growing increase in the use demand for
35 medicinal plants meant that the production of some species was not made on extractivism
36 form anymore, but now it is carried out in the form of a large-scale cultivation.

37
38 Like other cultivated species, medicinal plants can be attacked by pests and diseases which,
39 in addition to causing damage, can affect both qualitatively and quantitatively the production
40 of its active ingredients. Among the pathogens that may affect medicinal plants are
41 phytonematoids, [5], such as those of the genus *Meloidogyne*. Species of this group are
42 popularly known as root-knot nematodes, they are highly polyphagous sedentary organisms
43 penetrate the plant roots and when they set up their feeding cycle they cause the plants to
44 form galls at their roots, once installed their feeding cycle pass to parasitize the host plant.
45 The damage severity caused will depend on the susceptibility of the parasitized species /
46 variety, on the environmental conditions, on the presence of other pathogens and on the
47 population density of these pathogens in the soil [6].

48
49 In general, the root-knot nematodes weaken the parasitized plants as a result of the removal
50 of their nutrients. In addition, wounds, lesions and the formation of feeding sites near
51 vascular tissues reduce the roots ability to absorb water and nutrients. As a result, the aerial
52 part does not develop normally, causing reduction in the production and, in some cases,
53 culminating in the plant death. The medicinal species jaborandi (*Pilocarpus microphyllus*)
54 produces the alkaloid pilocarpine, which is synthesized in the roots and translocated to the
55 leaves, from where it is extracted and marketed as eye drops for the glaucoma treatment.
56 According to [7], commercial plantations of jaborandi, in the state of Maranhão - Brazil,
57 under conditions of high temperature, sandy soil and irrigation, were severely impaired by *M.*
58 *javanica*. The attack symptoms were the low development of the plants and the decrease in
59 the pilocarpine content of the collected leaves, compromising the drug production.

60
61 The control of *Meloidogyne* spp. is essential for the success of plant species cultivation,
62 since the root-knot nematodes can cause losses of up to 100% in the production, depending
63 on the infestation intensity of the area and the planted cultivars. Different control methods
64 can be used against the root-knot nematode, among them the chemical, cultural, biological,
65 but some of them stand out as inefficient and others can cause environmental damage,
66 compromising the sanity of the people who manipulate them such as pollution of the
67 agricultural environment. In view of the difficulty of nematode control by the previously
68 mentioned methods, the use of resistant cultivars is a widely accepted alternative, due to its
69 low cost and high efficiency [8]. Medicinal plants produce substances in their secondary
70 metabolism that may decrease or even inhibit the reproduction of field nematodes by
71 decreasing the population of these pathogens. In this sense, it is necessary to identify
72 species with these properties, aiming to make them a viable alternative to the
73 meloidogynosis management.

74

75 **2. GENERAL ASPECTS OF MEDICINAL PLANTS**

76

77 Medicinal plants are characterized as those have active principles capable of restoring the
78 homeostasis of the organism affected by diseases [9]. The natural properties of medicinal
79 plants are exploited by humans to meet their needs, especially those with therapeutic
80 actions and effects, and are often the only resources available for the treatment of diseases
81 in underdeveloped communities, rural communities and people living in isolated groups [10].
82 Traditional medicine involves the use of plant extracts or their active principles, research and
83 scientific proof of the effectiveness of these species used, can serve as a source of raw
84 material for modern medicines. In developing countries the use of medicinal plants
85 contributes by replacing imported drugs, thus increasing the economic self-sufficiency of
86 these countries [11].

87 Traditional medicine has a long history of serving people throughout the world, it is believed
88 the use of plants in the treatment of diseases has been carried out since the emergence of
89 early civilizations [12]. Fossil records indicate the use of plants as medicines since the
90 Paleolithic period. Evidence of this initial association was found in the tomb of a Neanderthal
91 man buried 60,000 years ago, where pollen analysis indicated that all plants buried with the
92 corpse had medicinal value [13].
93

94 The Sumerian Clay Table is the earliest known medical document with 4,000 years old, and
95 in that document are records plant remedies for various diseases. At the time of the ancient
96 Egyptian civilization, there was already information about the great wealth of medicinal
97 plants. Among the many remedies prescribed were man's drake for pain relief and garlic for
98 the treatment of circulatory and cardiac disorders. This information, along with hundreds of
99 other remedies, has been preserved in the Ebers papyrus written about 3,500 years ago
100 [13]. In ancient China there is also information about the first medicinal uses of plants.
101

102 In Brazil, the history of the use of plants in the treatment of diseases has strong influences
103 from African, indigenous and European cultures [14]. The contribution of the African peoples
104 to the tradition of the use of medicinal plants in Brazil came from the plants that the slaves
105 brought with them for use in religious rituals and also for their use of the pharmacological
106 properties, empirically discovered. The brazilian indian people who lived in countless tribes
107 used a great quantity of medicinal plants and, through the shamans, this knowledge of the
108 local herbs and their uses was transmitted and improved with each generation.
109

110 Medicinal plants, herbal remedies and isolated natural products represent a market that
111 moves billions of dollars in both developed and developing countries [16]. According to the
112 Organization for Economic Cooperation and Development [17], drug use in general is
113 increasing, and part of this increase is due to the growing need for medicines for the
114 treatment of chronic diseases and those related to aging.
115

116 In the group of drug products highlighted in the above survey, the following categories of
117 drugs have the highest growth rates, antihypertensive, antidepressant, antidiabetic and
118 antitumor drugs. In the survey conducted between 2000 and 2015 in the OECD countries,
119 consumption almost doubled for drugs used as antihypertensives, antidiabetics and
120 antidepressants, and almost quadrupled for drugs used to control cholesterol.
121

122 These data demonstrate how the pharmaceutical market is consolidated among the existing
123 pharmaceutical formulations, most of them are derived from plants or synthetic analogs
124 derived from plants. Based on the known curative capacity present in several plant species,
125 a large part of the world population, mainly from developing countries, still makes use of
126 medicinal plants for their daily health care needs and depend on plants for use as medicines
127 [16].
128

129 The World Health Organization also states that the consumption of medicinal plants has
130 increased considerably over the last few years, as well as the number of species used, [3].
131 Datas of the United Nations indicate the world market for natural products is growing by 7%
132 a year. Involving more than 32 thousand species, of which 900 are cultivated or are in the
133 process of domestication, and moves around 400 thousand tons of plants per year. In Brazil,
134 it is estimated that sales of pharmaceutical products derived from plants reach US \$ 550
135 million [18]. According to Coutinho [19], only the Amazon, in 2050, will be able to produce
136 500 billion dollars in medicines and cosmetics, from medicinal and aromatic plants.
137

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146

147 Brazil has one of the most interesting, important and diverse floras of the planet, distributed
148 in its limits in six bioactive phytogeographical domains: Amazon, Cerrado, Caatinga, Atlantic
149 Forest, Pampa and Pantanal [20]. There is a growing interest in scientifically studying and
150 documenting this diversity, including new technologies for study involving both academy and
151 pharmaceutical companies that promote studies for commercial exorcination of plant
152 resources for the development of new medicines.
153

154 Due to the growing interest in the use of medicinal plants, there is a notable growth in the
155 field of research involving these species, in addition to the adoption by health systems in
156 developed countries of traditional medicine as a medical practice integrating their culture.
157 Historically, all drugs were derived from plants, either in the simple form, through raw
158 vegetable materials or in the refined form of crude extracts, mixtures, oils, etc., which
159 demonstrate the important role of research and development of new drugs. Since 2002, the
160 World Health Organization (WHO) has been encouraging the rescue of data on plants used
161 in the practice of ancient medicine, since they are considered potentially useful in the
162 development of new drugs [22] Recent estimates suggest that thousands of plants are
163 known with medicinal applications in various cultures [23], and are still used today to treat
164 diseases [24].
165

166 Several studies about the chemical composition of medicinal plants allowed an association
167 between different species and their respective biological activities based on observation,
168 description and experimental research. Such studies have largely contributed to the
169 discovery of bioactive natural products [25]. Medicinal plants produce a broad spectrum of
170 chemical compounds, apparently without direct contribution to their growth and
171 development, termed secondary metabolites [26]. Such as those outlined below, divided into
172 three main classes of compounds of interest in plants: flavonoid, terpenes and phenolic
173 compounds. Terpenes are found in families such as Rhamnaceae, have several curative
174 actions, such as purgative, energetic, and antifebrile action, being widely used because of its
175 effectiveness and its low cost [27]. Flavonoids are present in families such as Passifloraceae
176 that represent a chemical class of ubiquity in the group of phenolic substances of plant origin
177 [28], presenting a wide range of biological activities, such as antitumor, anti-inflammatory,
178 antioxidant, antibacterial action, antiparasitic, antiviral, among others [29]. And, the phenolic
179 compounds found easily in plants of the family Lamiaceae and Asteraceae that may exhibit
180 antimicrobial and anti-inflammatory effects [30].
181

182 For the plants these compounds have different biological properties and in phytotherapy are
183 used in the treatment of a wide variety of diseases, including cancers [31], neurological
184 disorders 26, chronic inflammations and lesions [32], diabetes [33], atherosclerosis [34],
185 cardiovascular diseases and skin diseases [35]. Many botanical families have in their
186 compounds healing properties, derived from substances produced by their secondary
187 metabolism.
188

189 **2.1 Acanthaceae**

190
191 Acanthaceae is a family of dicotyledonous plants, of the order Lamiales, which contains 250
192 genera and about 2,500 species, mostly composed of tropical herbs, shrubs, creepers or

193 epiphytes, mainly distributed in Indonesia, Malaysia, Africa, Brazil and Central America [36].
194 In Brazil, the family is represented by 41 genera and 432 species [37], of which *Justicia*
195 *gendarussa* (black adusa) and *Justicia pectoralis* var. *stenophylla* (freshcut) are some
196 examples of medicinal plants of this family. Some species are used as ornamental and
197 forage, and are commonly found in both wet and semi-arid forests [38]. Phytochemical
198 studies indicate some species of this family have potential for pharmacological use.

199
200 The medicinal plants of this family are used for the treatment of fever, rheumatism,
201 headache, abdominal cramps, lung inflammations, cough, and also as expectorant, sudorific,
202 anti-inflammatory and aphrodisiac [15]. The main constituents present in these plants
203 responsible for these curative effects are coumarins, umbelliferae, dihydroxycoumarin,
204 acetylated orthohydroxy-transcinnamic acid, betasitosterol, C-glycosylphoflavones-0-
205 methoxylated eswertisin, 2"-0-ramnosilesterthylaponine, betaine and lignan justicidina B.

206 207 **2.2 Amaranthaceae**

208
209 The Amaranthaceae family is composed of 169 genera and about 2,360 species, being the
210 largest genera being *Atriplex* (300 spp.), *Gomphrena* (120 spp.), *Salsola* (120 spp.),
211 *Alternanthera* (100 spp.), *Chenopodium* (100 spp.), *Ptilotus* (100 spp.), *Suaeda* (100 spp.),
212 *Iresine* (80 spp.), *Amaranthus* (60 spp.), *Corispermum* (60 spp.) e *Celosia* (50 spp.), mainly
213 distributed by *Ptilotus* (100 spp.), *Suaeda* (100 spp.), distributed in regions of tropical climate
214 and temperate regions. In Brazil, there are 145 species belonging to 19 genera, being 71
215 endemic species from different regions and Brazilian biomes [40]. The plants of the
216 Amaranthaceae family are predominantly herbaceous and sub-shrub.

217
218 About 20 species have food and / or medicinal uses, mainly species of the genus
219 *Alternanthera*, *Amaranthus* and *Gomphrena*. Among the medicinal properties of these plants
220 are the antitumor, diuretic, digestive, depurative, vermifuge, emmenagogue, tonic, bitter,
221 immunostimulating, astringent, antidiarrheal, antitussive, laxative and antibleorrhagic
222 activities [15]. These medicinal properties are derived from the presence of flavonic
223 compounds, palmitic, oleic and linoleic acids, as well as carotenoids, saponins, ecdysterone,
224 spinasterol and ascaridiol.

225
226 The species of the family Amaranthaceae are able to tolerate highly arid habitats and very
227 saline soils due to a series of specific adaptations. These include extremely high seed
228 production, varying between 13,000 and 50,000 seeds per plant, depending on soil richness
229 and an efficient seed dispersal mechanism [41]. Both mechanisms are intended to ensure
230 the survival of plants during arid periods and in hostile environments. Some species may
231 accumulate high concentrations of arsenic in different parts of their organism, without
232 showing decrease of the growth, contributing to their use as fixatives in highly polluted soils
233 [41].

234 235 **2.3 Apiaceae**

236
237 The family Apiaceae (formerly known as the Umbelliferae family), is one of the great families
238 among dicotyledons, comprising approximately 450 genera and 3,700 species [42]. This
239 family has great economic value, because it has edible species, spices, as well as used in
240 perfumery or as essences in alcoholic beverages. Besides the gastronomic importance,
241 these plants are sources of gums and resins of great medicinal use as sedatives,
242 antispasmodics, stimulants, and even poisons, being possible to emphasize *Anethum*
243 *graveolens* (dill), *Anthriscus cerefolium* (garden chervil), *Angélica* spp. (*Angelica*), *Carum*
244 *carvi* (caraway), *Coriandrum sativum* (Coriander), *Cuminum cyminum* (cumin), *Foeniculum*
245 *vulgare* (sweet fennel), *Ferula gummosa* (galbanum) and *Pimpinella anisum* (anise) [15].

246

247 The plants of this family usually have the pungent odor due to the presence of essential oil
248 or oleoresin and several other compounds with many biological activities [43]. Some of the
249 main properties are the ability to induce apoptosis, antibacterial action, anti-inflammatory,
250 excitatory, depurative, depurative, expectorant, febrile, aphrodisiac, hepatoprotective, vaso-
251 relaxing, diuretic effects, cyclooxygenase inhibitor and antitumor activities 44. The essential
252 oils of Apiaceae are employed in different fields such as food and beverages, cosmetics and
253 pharmaceuticals, [45]. In addition to the essential oil these plants may contain alkaloids,
254 saponins, flavonoids, quercetin, amino acids, minerals, pectins, tannins, mucilage, acetic
255 and oxalic acids, coryandrol, limonene, linalool, terpinene and sugars. They are also
256 considered promising candidates for biocides to control parasites and vectors [46].
257

258 **2.4 Apocynaceae**

259

260 The family Apocynaceae is composed of species usually characterized by the presence of
261 latex, about 5,000 species distributed in 550 genera [47]. They are found predominantly in
262 the tropics and subtropics [48]. In Brazil, there are more than 400 species in 41 genera, of
263 which 32 are found only in the Amazon [49].
264

265

266 Many of the tree species of this family present high quality in their woods [50]; in addition,
267 their bark is commonly used in the infusions form for the treatment of a number of diseases
268 due to their low toxicity and the absence of contraindications [50]. As examples, the species
269 *Aspidosperma ramiflorum* (guatambu amarillo) is employed in the treatment of
270 leishmaniasis; *Aspidosperma nitidum* (brasilian name: "carapanaúba") is used as a
271 contraceptive, in the inflammation treatment of the uterus and ovary, in diabetes, in stomach
272 problems, against cancer, fever and rheumatism; *Catharanthus roseus* (madagascar
273 periwinkle) is used as vasodilator, sudorific, diuretic, hypoglycemic, antileukemic and
274 febrifuge and; *Calotropis procera* (roostertree) that has analgesic, antirheumatic, proteolytic,
275 anti-inflammatory, odontogenic and tranquilizing properties [15].
276

277

278 Phytochemical studies point to the frequent occurrence of alkaloid structures, biologically
279 they act on the opiate, GABAergic, cholinergic, muscarinic, serotonergic and dopaminergic
280 neurotransmitter systems [50]. Their are widely used as arterial hypotension, sympatholytic,
281 diuretic, peripheral vasoconstrictor, respiratory stimulant, anesthetic, adrenergic blocking
282 agent, intestinal spasmogen, sedative and skeletal muscle relaxant [51]. Besides these
283 substances, methyl ether, coumarins, sugars, triterpenoids, betulinic acid, cardioactive
284 glycosides and lactose iridoids can also be found [15].
285

286

287 **2.5 Asteraceae**

288

289 The Asteraceae family is predominantly composed of herbaceous individuals, but includes
290 some woody species and arboreal types. This botanical family has approximately 25,000
291 species distributed in more than 1,100 genera [52], being characterized as the second
292 largest family of the vegetable kingdom. The family members have a remarkable ecological
293 and economic importance, including ethnobotanical, phytochemical, and medicinal purposes
294 [52].
295

296

297 Several plants of this family are edible and are used in traditional medicine. The species of
298 this family are generally a source of many biologically active compounds such as essential
299 oils, organic compounds [53], flavonoids [54], terpenoids [55], lignans [54], alkaloids,
300 saponins, stilbenes and sterols [56], polysaccharides [57], antioxidants, lapatin, fuquinone,
301 glycosides, mucilages, antibiotic principle and B-complex vitamins conferring anesthetic,
302

298 bactericidal, anti-inflammatory, antimycotic, contraceptive, cancerous and ulcer inhibitory
299 action, antipyretic, hepatoprotective and immunostimulatory activity [15].
300

301 **2.6 Brassicaceae**

302

303 The Brassicaceae family (syn. Cruciferae) has relevant importance economic and
304 scientific, because it includes a large number of cultivated plants, mainly olive, medicinal and
305 ornamental species, weeds and some model organisms, such as *Arabidopsis thaliana* [58].
306 The family has 338 genera containing 3,709 species [59], among the main medicinal plants
307 belonging to this family, it is possible to highlight *Brassica rapa* (mustard), *Coronopus*
308 *didymus* (lesser swinecress) and *Nasturtium officinale* (watercress).
309

310 In traditional medicine, plants belonging to the family Brassicaceae are used as a revulsive
311 or rubefacient drug, blood depurative, diuretic, expectorant, febrifuge, stomachic, tonic and
312 as a stimulant of the digestive organs. Externally, they can be used in the treatment of
313 rheumatic pain and injury [15]. These medicinal properties are due to the presence of
314 essential oils, glycosides, carotenes, thioglucosides, minerals, vitamins, triglycerides,
315 proteins, sinapine and phenylpropanoide.
316

317 **2.7 Costaceae**

318

319 The Costaceae family belong to about 110 to 115 species, distributed in the genera *Costus*,
320 *Monocostus*, *Dimerocostuse* and *Tapeinocheilas*, being generally found in tropical and
321 subtropical areas, tropical forests and other humid environments. *Costus*, the largest genus,
322 has a pantropical distribution, with its greatest diversity in the neotropical region (about 40
323 spp.), The species of this family present great variability in terms of leaf and flower
324 morphology [61]. Economically, the family still has little importance, being cultivated mainly
325 for ornamental and medicinal purposes [62].
326

327 Phytochemical studies in plants of the Costaceae family show mainly the presence of
328 flavonoids, quercetin, tamarixetine, steroids, alkaloids, saponins, inulin, tannins, sistosterol,
329 mucilages, sapogenins, pectins and calcium oxalate [63]. In addition to these compounds,
330 diosgenin has also been isolated from rhizomes, which has attracted the attention of many
331 researchers as a new precursor source of steroidal hormones [64]. These compounds confer
332 depurative, astringent, diuretic, antimicrobial, antioxidant, anti-inflammatory and
333 cardiovascular protective properties [65].
334

335 **2.8 Crassulaceae**

336

337 Crassulaceae is a family of dicotyledonous plants, consisting of 35 genera with
338 approximately 1,400 species, usually herbaceous and succulent plants, occasionally sub-
339 shrub or shrub, mainly distributed in the Northern Hemisphere and Southern Africa [66].
340 Several species of this family are used in the landscaping as forraces, creating borders or
341 massifs in the sunny spaces, being ideal for compositions of rocky gardens, some plants are
342 cultivated in pots, including in hanging pots, making use of pending plants or prostrate
343 habits. They are also ideal for the constitution of minijardins, according to the species. [66].
344

345 Among the species belonging to the family Crassulaceae, the species *Bryophyllum pinnatum*
346 (airplant) presents a wide use in folk medicine for the treatment of arthritis and dyspepsia
347 [66, 15]. The said species also has anti-inflammatory, antitumor, antiviral, antimicrobial,
348 emollient and healing properties and can be used for the relief of pertussis, bronchitis,
349 asthma and various respiratory tract infections; external use for the treatment of burns,
350 dermatoses, bruises, cuts, dermatitis and boils; acnes, pimples, calluses and chicken pox;

351 itching for no apparent reason; heartburn, gastritis, ulcer and stomach or abdominal
352 discomfort; headaches and migraines; dysentery and diarrhea; cramps and menstrual
353 disorders; balance diabetes; eliminate or reduce kidney stones [67]. Regarding chemical
354 composition, the literature reports the presence of polysaccharides, anthocyanidins,
355 flavonoids, bufadienolides, aglycone, patuletin, quercitrin and isoquercitrin [68].
356

357 **2.9 Lamiaceae**

358
359 The Lamiaceae family is one of the most diversified in terms of use in traditional medicine,
360 having its medicinal value based on the high concentration of volatile oils [69]. It is one of the
361 largest families among dicotyledons, being composed of more than 240 genera, many
362 species belonging to this family are highly aromatic, due to the presence of glandular
363 structures producing volatile oils [70]. These oils are important in the composition of
364 pesticides, pharmaceuticals, flavorings, perfumes, fragrances and cosmetics [71].
365

366 Species of Lamiaceae family have a wide range of biological and pharmacological activities,
367 including essential oils, linalool, limonene, citral, citronellol, flavonoids, mucilages, esters,
368 coumarins, tannins, saponins, diterpenes, stachyhyd, glycosides, fatty acids, edridiol, resins
369 and bitter substances [15].
370

371 These compounds are present in the species of this family with antimicrobial, spasmolytic,
372 carminative, anticancer, antivomitive, anti-infective, anti-dyspeptic and other properties [15,
373 70]. Some medicinal herbs of this family such as *Mentha arvensis* (wild mint), *Mentha*
374 *piperita* (peppermint) *Thymus vulgaris* (garden thyme) and *Salvia officinalis* (kitchen sage),
375 have expressive amounts of phenolic acids and are therefore considered promising sources
376 of natural antioxidants [72].
377

378 **2.10 Moraceae**

379
380 The Moraceae family comprises 37 genera and more than 1,100 species worldwide,
381 distributed in tropical forests of Central and South America [73]. In general, the plants in this
382 family are trees and shrubs adapted to a wide range of tropical rainforest habitats, including
383 terra firme forests, seasonal and flooded forests [74]. Species of the family Moraceae have
384 important economic value, producing quality wood for use in the civil construction, naval and
385 furniture industries, in addition to its recognized medicinal value. These species are widely
386 recognized as sources of bioactive secondary metabolites, among them flavonoids,
387 stilbenes, triterpenoids and xanthenes [75].
388

389 Among the medicinal plants of the family Moraceae stand out: *Brosimum gaudichaudii*
390 (mamica-de-cadela) used in the treatment of vitiligo and other skin patches, rheumatic
391 diseases, cold flu and bronchitis [76]; *Dorstenia cayapia* (*Dorstenia*) for treatment of typhoid
392 fever and other infections of the digestive tract, of respiratory airways and respiratory atony
393 [76]; *Ficus carica* (edible fig) widely used in traditional medicine as a pectoral and laxative
394 emollient, for treatment of constipation, bronchitis, coughs, flu, colds, mouth and throat
395 inflammations [77] and; *Ficus insipida* (chamate) that has purgative, aphrodisiac and memory-
396 stimulating action [76]. The literature reports that these medicinal properties are due to the
397 presence of alkaloids, sterols, triterpenes, coumarins, flavonoids, furanocoumarins, organic
398 acids, mucilages, pectin, sugars, dorstenin, bergapten and psoralen.
399

400 **2.11 Phytolaccaceae**

401
402 The family Phytolaccaceae comprises about 17 genera and 120 species pantropical, widely
403 distributed throughout the American territory [78]. Among the species of Phytolaccaceae, the

404 most popular is *Petiveria alliacea* (guinea henweed), a perennial shrub with a rigid and
405 straight stem, reaching a height of up to 150 cm [79]. The medicinal use of *P. alliacea* occurs
406 in several regions of the world, mainly in the American continent. the same has curative and
407 mystical purposes, is used in religious ceremonies in Brazil, with reports from the time of
408 slavery, where slaves used *P. alliacea* for its toxic and sedative effects [80]. According to
409 indigenous medicine, the root, powder and leaf of *P. alliacea* has several therapeutic
410 properties, such as diuretic, antispasmodic, emenagogic, analgesic, anti-inflammatory,
411 antileukemic, antirheumatic, anthelmintic, antimicrobial and depurative properties.
412

413 In addition, different preparations of *P. alliacea* are used for their activities in the central
414 nervous system (CNS) as anticonvulsants, anxiolytics, mnemonics, anesthetics and
415 sedatives [81]. In the chemical composition of *P. alliacea* are found coumarins, saponins,
416 flavonoids, tannins and organic sulfides [82].
417

418 2.12 Rutaceae

419 About 180 genera and 1300-1600 species belong to the Rutaceae family. Plants of this
420 family have great economic value and some of them have highly fragrant flowers and are
421 used in the commercial production of essential oils. Some constituents of essential oils, such
422 as citronella and bergamot, are obtained by distillation of plants of this family [83]. Genres
423 like *Citrus*, *Poncirus* and *Fortunella* produce fruits with high commercial value; *Citrus* has
424 aromatic plants; *Murraya*, *Atalantia* and *Citrus* have ornamental plants; and *Zanthoxylum*,
425 *Citrus*, *Phellodendron*, and *Evodia* have species used for medicinal purposes [84].
426
427

428 Among the medicinal plants belonging to the Rutaceae family, it is possible to emphasize
429 *Citrus aurantium* (bigarade) being considered aromatic, bitter, digestive, expectorant, diuretic
430 and hypotensive [85]; *Ertela trifolia* has emmenagogue, diuretic, tonic, sudorific and
431 stimulant properties [86]; *Pilocarpus microphyllus* used in the treatment of bronchitis, dry
432 skin, and as hair tonic, in addition to possessing sweat, febrifuge and stimulant properties
433 [15]; and *Ruta graveolens* is used in the treatment of menstrual disorders, skin
434 inflammations, ear pain, toothache, cramps, liver diseases and verminoses [87]. In the
435 phytochemical composition of Rutaceae, pectin, hesperidin, bitter substances, narigenin,
436 sugars, limonene, linalool, linalyl acetate, saponin, lignan, alkaloids and flavonic glycosides
437 stand out [15].
438

439 2.12 Solanaceae

440 The Solanaceae family has 3,000 species, distributed in 84 genera [88]. They are species of
441 cosmopolitan distribution, in Brazil there are 32 genera and 350 species, among
442 herbaceous, shrub and arboreal plants [15]. This family has several species of economic
443 importance such as potato (*Solanum tuberosum*), eggplant (*S. melongela*), tomato (*S.*
444 *lycopersicum*), cucumber (*S. muricatum*), peppers in general and pepper (*Capsicum* spp.).
445 Some plants of the family Solanaceae have been studied for use in the pharmaceutical
446 industry because they possess chemicals of the class of alkaloids in general, emphasizing
447 tropical alkaloids and ale-steroids. Many species are widely used for medicinal,
448 hallucinogenic and for steroid synthesis, used in the pharmaceutical industry [15].
449
450

451 This family has a wide range of medicinal plants, among them *Datura innoxia* (prickly burr)
452 used to cure cough and asthma [90], treatment of swollen limbs, use as repellent and
453 vermicide [91]. *Solanum nigrum* (black nightshade) is an antispasmodic, diuretic, analgesic,
454 sedative, narcotic, expectorant, anaphrodisiac and laxative plant [92]; *Solanum surattense*
455 (yellow-fruit nightshade) is used in the treatment of phlegmatic cough, asthma, chest pain
456 and fever [93]; and *Withania somnifera* (withania) is useful in fighting chronic fatigue,

457 weakness, dehydration, brittle bones, impotence, premature aging, weight loss, weakness,
458 muscle tension, tumors, inflammation, psoriasis, bronchitis, asthma, ulcers, scabies and
459 insomnia [94]. The main active constituents responsible for the healing properties of the
460 Solanaceae family are fatty acids, alkaloids, triterpenoids, saponins, steroids, essential oils
461 and quassinoids [15].
462

463 **2.13 Verbenaceae**

464

465 The Verbenaceae family consists of trees, shrubs and herbs found mainly in the tropical
466 regions of the world [95]. The family includes about 1,200 species distributed in 35 genera.
467 Plants of the Verbenaceae family are well known for their uses in traditional medicinal
468 systems in several countries [95]. A large number of plants in this family have bioactive
469 phytochemicals with important pharmacological effects, among them the diterpenoids,
470 having antimicrobial and antiparasitic properties [96], tannins, mucilages and alkaloids (97,
471 15).

472
473 Some species of medicinal plants of this family are *Lippia alba* (bushy lippia) is a specie
474 commonly used throughout Brazil, has high concentrations of citral, myrcene, limonene and
475 carvina, these constituents give it calming, analgesic, anxiolytic, sedative and mucolytic [98];
476 *Lippia gracilis* (brasilian name: "alecrim de chapada") is a species of the semi-arid region of
477 the Northeastern region. Its chemical composition contains thymol, carvacrol, flavonoids and
478 quinones, which give it antimicrobial, fungicidal and molluscicidal activities, as well as being
479 used in the treatment of acne, scabies, white cloth, impingens, dandruff and bad smell in the
480 armpits, feet and groin [15]; and *Stachytarpheta cayennensis* (cayenne porterweed) is a
481 plant widely used in traditional Brazilian medicine for the treatment of fevers, dyspepsia,
482 hepatitis, constipation, colds, flu, bronchitis, wounds, bruises and skin conditions.
483

484 **3. ROOT-KNOT NEMATODES**

485

486 Nematodes probably arose early in the Cambrian Period [99], although some researchers
487 have suggested that their origin dates back to a billion years ago [100]. Free-living
488 nematodes, present in terrestrial and aquatic environments, have developed parasitic
489 relationships with other eukaryotes on several independent occasions [101]. Molecular and
490 palaeontologic evidence suggests land plants originated from 425 to 490 million years [102],
491 indicating that land plants and nematodes coexisted in the earth soil for an extended period
492 of time, developing morphological and physiological characteristics specific to feed from the
493 roots of plants in the rhizosphere [103].
494

495 The productive and economic losses caused by nematodes on cultivated species are difficult
496 to establish accurately. It is estimated that global averages of income loss are around 10%,
497 with this figure reaching 20% for some crops; in monetary terms, world losses exceed US \$
498 175 billion annually [6]. Most of the damage is caused by a relatively small number of
499 nematode genera that attack the crops, especially the sedentary root-knot nematodes
500 (*Meloidogyne* spp.), the cyst nematode (*Globodera* and *Heterodera* spp.) and some
501 migratory nematodes (including *Pratylenchus* and *Radopholus* spp.) [3].
502

503 The species with the greatest agricultural importance are those that infect the roots, this
504 because many of the control strategies, such as chemical control, are inefficient [104]. Root-
505 knot nematodes (*Meloidogyne* spp.) are one of the most polyphagous and damaging genera
506 between plant parasitic nematodes. The species of this genus are biotrophic endoparasites
507 capable of infecting a large number of higher plant species and have an almost
508 cosmopolitan distribution [103]. Several genera of nematodes exploit all parts of plants, but
509 the economically most important species are those that infect the roots. In part because

510 many of the control strategies, such as chemical control, are inefficient [104]. Root-knot
511 nematodes (*Meloidogyne* spp.) are one of the most polyphagous and damaging genera of
512 plant parasitic nematodes. The species of the genus *Meloidogyne* are biotrophic
513 endoparasites capable of infecting a large number of higher plant species and have an
514 almost cosmopolitan distribution [103].

515
516 The soil texture strongly influences the distribution of *Meloidogyne* spp., since they occur
517 predominantly in sandy soils, besides the texture, the presence and development of *M.*
518 *incognita* is associated to neutral pH soils, where they may persist in dormancy for a certain
519 period of time [105]. [106] state that in the absence of host plants the eggs of this nematode
520 are resistant to environmental and nematicidal stresses and hatch only under the stimulation
521 of plant root emanations. During this period of dormancy the eggs can withstand soil water
522 deficiency and only hatch under favorable conditions to the development of the pathogen,
523 making it difficult to control.

524
525 The species of the genus *Meloidogyne* are highly diversified, showing diversity in terms of
526 cytogenetics, mode of reproduction, specialization in parasitism, and range of hosts [107]. In
527 general, this high level of diversity contributes to a complex pathogen-host interaction
528 leading to highly successful parasitism. For example, the three main species of the genus
529 *Meloidogyne* (*M. incognita*, *M. javanica* and *M. arenaria*) are highly polyphagous, infecting
530 more than 3,000 plant species [107].

531
532 *Meloidogyne* spp. has a sedentary endoparasitic lifestyle with second-stage infectious-
533 mobile juveniles (J2) that invade the plant through the roots and migrate intercellularly
534 through tissues to developing vascular cells [106], reproduced by mitotic parthenogenesis
535 [108]. These nematodes establish an intimate relationship with their hosts, inducing the
536 formation of gall, the main visible symptom of root-knot nematode infection [106].

537
538 Infection occurs when the second stage mobile juvenile (J2) is attracted to the host plant root
539 system. J2 migrates intercellularly toward the vascular cylinder, where it injects secretions
540 through its stylet into approximately six cells of the parenchyma. The affected plant cells
541 undergo a transformation, becoming giant and multinucleate in which the nematode
542 nourishes non-destructively during the rest of its life cycle. During their biological cycle, the
543 nematodes feed on nutrients and water from the host plant. This impairs the plants growth,
544 causing wilt and increasing the susceptibility of the plant to other pathogens and, in some
545 cases leading to death plant [110].

546
547 According to [111] the secretions originating from the esophageal glands of the nematode
548 play important roles in the penetration of the host and in the formation of giant cells. The
549 root-knot nematodes have three esophageal glandular cells, one dorsal and two subventral
550 gland cells. Each gland is a single enlarged cell that is specialized to export secretions to the
551 esophagus of the nematode [112]. Once in the lumen, the secretions of the glands are
552 exported out of the nematode through the stylet. Some studies report that the subventral
553 glands produce pectinases and cellulases that help the nematode penetrate the host root
554 [104]. This hypothesis was confirmed by the identification of several genes encoding
555 cellulases and pectate lyases that are expressed in the subventral glands of the root-knot
556 nematodes [104].

557
558 After the feeding cycle, the *Meloidogyne* spp. will cause reduction in the translocation of
559 water and nutrients from the roots to the leaves causing symptoms, in the aerial part,
560 identical to those of nutritional deficiency. The increase in the metabolic activity of giant cells
561 induced by nematode parasitism will stimulate the mobilization of photoassimilates from
562 shoot to roots and, in particular, to the giant cells themselves, which are used to feed the

563 nematode [111]. The intensity in reducing the size and production of the plant varies
564 according to the population of nematodes, the association with other pathogens (*Fusarium*
565 spp., *Rhizoctonia solani*, etc.) and with other factors, such as the degree of susceptibility of
566 the cultivated variety and soil fertility.

567

568 Another way to consider the impact of nematodes on crops is through the management
569 strategies used in their control. In 2010, about US \$ 7.3 billion in chemical products were
570 sold in Brazil for the treatment of pests and diseases of plants, corresponding to about 7% of
571 total sales of agricultural products in the country. [113] However, issues such as
572 groundwater contamination, mammalian and avian toxicity, and residues in food have
573 caused much more severe restrictions on the use of chemicals, and in many countries the
574 use of effective but highly toxic nematicides, were and are still prohibited [6]. The literature is
575 also replete with studies on organic media, such as green manures, to control nematodes,
576 but these strategies are ineffective [114].

577

578 Even if environmentally safe nematicides are developed, or if biocontrol agents are
579 identified, host resistance will continue to be the safest and most efficient form of nematode
580 control [104]. Resistance to nematodes is yet to be identified for many cultivated plants,
581 although several resistance genes have been identified [115]. [116] reports that in peppers
582 (*Capsicum annuum*) there are ten genes for dominant resistance to *Meloidogyne* spp. (*Me1*,
583 *Me2*, *Me3*, *Me4*, *Me5*, *Me6*, *Me7*, *Mech1*, *Mech2* and *N*). According to the aforementioned
584 author, four of the ten genes (*Me1*, *Me3*, *Me7* and *N*) confer resistance to a wide range of
585 species of the genus *Meloidogyne*, serving as important resources for the development of
586 peptide cultivars resistant to meloidogynosis.

587

588 Although about 100 species of *Meloidogyne* are known so far [117], historically, species of
589 the genus *Meloidogyne* were divided into main and secondary species. According to [108]
590 *M. arenaria*, *M. incognita*, *M. javanica* (occurring in tropical regions) and *M. hapla* (occurring
591 in temperate regions) are considered the four main species of *Meloidogyne*. However, these
592 authors consider five more species as secondary species. They are: *Meloidogyne chitwoodi*,
593 *M. fallax*, *M. enterolobii*, *M. minor* and *M. paranaensis*.

594

595 The vast majority of research focused on only four species: *M. arenaria*, *M. hapla*, *M.*
596 *incognita* and *M. javanica*. The reason for the high status of these four species is in part due
597 to the fact that they are extremely widespread and infect a wide range of hosts, but to a
598 certain extent it is also historical and can be attributed to a study of [108]. In this study, the
599 authors reported that *M. arenaria*, *M. hapla*, *M. incognita* and *M. javanica* made up 99% of
600 all species identified in more than 660 isolates from 65 countries.

601

602 [119] formulated a study to list the most harmful nematodes for agriculture, for which the
603 authors interviewed 371 nematologists worldwide. More recently, another survey was carried
604 out with the same purpose and was attended by 225 nematologists [120]. In both cases, the
605 votes were grouped, resulting in a list of the 10 most important nematodes for agriculture
606 (Table 01). In general, the lists are similar, mainly in the first five positions, and in both
607 stands out importance of the genus *Meloydogine* classified as the most important in both
608 studies.

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Table 1. Ranking of the main phytonematoids according to economic importance in a survey carried out by Sasser and Freckman (1987) and Jones et al. (2013)

Classification of Sasser and Freckman (1987)		Classification of Jones et al. (2013)	
1°	<i>Meloidogyne</i>	1°	<i>Meloidogyne</i> spp.
2°	<i>Pratylenchus</i>	2°	<i>Heterodera</i> spp. e <i>Globodera</i> spp.
3°	<i>Heterodera</i>	3°	<i>Pratylenchus</i> spp.
4°	<i>Ditylenchus</i>	4°	<i>Radopholus similis</i>
5°	<i>Globodera</i>	5°	<i>Ditylenchus dipsaci</i>
6°	<i>Tylenchulus</i>	6°	<i>Bursaphelenchus xylophilus</i>
7°	<i>Xiphinema</i>	7°	<i>Rotylenchulus reniformis</i>
8°	<i>Radopholus</i>	8°	<i>Xiphinema index</i>
9°	<i>Rotylenchulus</i>	9°	<i>Nacobbus aberrans</i>
10°	<i>Helicotylenchus</i>	10°	<i>Aphelenchoides besseyi</i>

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3.1 *Meloidogyne enterolobii*

Meloidogyne enterolobii was initially reported causing severe damage to a pacara-earpod tree (*Enterolobium contortisiliquum*) population on the island of Hainan in China [121]. Since this report, *M. enterolobii* has been the topic of few published studies, which seemed to be of little importance [121], which is in opposition to the fact that the species is considered a polyphase with a range of hosts similar to *M. incognita* [122]. Most registered hosts often include vegetables such as tomato, pepper and watermelon [122]), fruit trees such as guava [123], ornamental plants [124], weeds and medicinal plants [125]. Despite being less frequently reported in the literature, *M. enterolobii* is a highly aggressive species with a high rate of root infestation that induces more severe symptoms than other species of gall nematodes [126].

In heavily infested areas, cultivation may become unfeasible, as for guava in Brazil [127]. Another interesting fact is the ability of *M. enterolobii* to develop in genotypes of crops resistant to the main species of *Meloidogyne*, including cotton, sweet potato, tomato (*Mi-1* gene), potato (*Mh* gene), soybean (*Mir1* gene), chili (*N* gene), sweet pepper (*Tabasco* gene) and cowpea (*Rk* gene) [128, 129, 130]. Few cultures are recorded as resistant to *M. enterolobii*, including grapefruit, sour orange, garlic and peanut [131].

Until recently, it was assumed that the distribution of *M. enterolobii* was restricted to regions with typically tropical climatic conditions, such as Africa, South and Central America, the Caribbean and Asia. In 2004, during the regulatory sampling in ornamental nurseries, this nematode was detected in Florida [131]. In addition, the presence of *M. enterolobii* was reported in greenhouses in France and Switzerland [132]. There is the possibility that its distribution has been underestimated due to misidentification in different regions of the world, including Europe. As *M. enterolobii* is also likely to survive in the hottest parts of Europe and in greenhouses throughout the region, the risk of its establishment and dissemination in this area is very likely [133]. In 2010, *M. enterolobii* was added to the alert list of the European Organization for the Protection of Mediterranean Plants (EPPO) [134], with a recommendation by that organization that its member countries define this nematode as a quarantine pest.

652 In Brazil, the first report of this nematode causing damages in cultivated plants was carried
653 out by [135] in the municipalities of Petrolina in Pernambuco, and of Curaçá and Juazeiro in
654 Bahia in commercial plantations of guava (*Psidium guajava* L.). The main symptoms
655 described were the presence of large galls associated with necrosis of the root system,
656 resulting in drastic reduction of radicles, important in plant nutrition. Subsequently, *M.*
657 *enterolobii* was recorded in several regions of the country causing severe damage in
658 commercial guava orchards, such as: Rio de Janeiro [136] Ceará [137], Piauí [138], Paraná
659 [139], Mato Grosso do Sul [140], Espírito Santo [141] and Maranhão [142].
660

661 **3.2 *Meloidogyne incognita***

662
663 *Meloidogyne incognita* is a pathogen that causes deformation in the root system of
664 parasitized hosts, causing gall formation and occurrence of cracks in the cortex. The most
665 frequent symptoms as a consequence of the presence of the pathogen are wilt, chlorosis
666 and nutritional deficiency symptoms [143]. It is a highly polyphagous species, causing
667 significant economic losses [144], being harmful to cotton (*Gossypium hirsutum*), sugar beet
668 (*Beta vulgaris*), tomato (*Solanum lycopersicum*), pepper (*Capsicum annuum*), watermelon
669 (*Citrullus lanatus*), melon (*Cucumis melo*), okra (*Abelmoschus esculentus*), soybean
670 (*Glycine max*) among other cultivated species [23]. *Meloidogyne incognita* is a control
671 species more difficult than any other species of the genus *Meloidogyne* due to its high
672 variability within the species, with four species known for this species [143].
673

674 It is a parasite that has the ability to develop rapidly under appropriate conditions. This rapid
675 population growth is mainly due to the rapidity of its biological cycle (about 28 days),
676 combined with the high female fecundity. The exact number of eggs produced varies
677 depending on environmental conditions. Under favorable conditions, a single female can
678 produce 500 to 2000 eggs [143]. The eggs have shells containing chitin as protection and
679 are deposited by the female in a gelatinous matrix resistant to desiccation secreted by the
680 female. Although males occur, reproduction occurs exclusively via mitotic parthenogenesis
681 (apomixis) [143].
682

683 **3.3 *Meloidogyne javanica***

684
685 *Meloidogyne javanica* is one of the most important species among the group of gnats
686 nematodes, has a wide range of hosts among crop cultivars, pastures and grasses,
687 horticultural and ornamental crops. This species is highly adapted to the most diverse
688 environmental conditions. [124] *M. javanica* is distributed throughout the world, developing at
689 temperatures ranging from 3 ° C up to a maximum of 36 ° C, and can tolerate up to 4 or 5
690 months without rainfall, and has a preference for clay-poor soils [108].
691

692 The embryogenesis of *M. javanica* has been studied for over 40 years, since it has been
693 reported that the stages of development of the embryo are easily identifiable, with eight
694 stages in egg formation [144]. There are several cell divisions leading to the additional
695 elongation stage, resulting in the juvenile first stage, the J1 stage of *M. javanica* has
696 approximately 500 cells, the eggshell has three layers, the yolk layer being very peripheral,
697 on a chitinous layer and a more internal layer of lipids, the *M. javanica* egg can withstand
698 long periods of drought and only hatch under conditions favorable to the development of the
699 pathogen [144].
700

701 **4. REACTION OF MEDICINAL PLANTS TO *Meloidogyne* spp.**

702
703 Among the diseases can compromise the qualitative and quantitative characteristics of the
704 pharmacological and productive properties of medicinal plants, nematode diseases can be

705 highlighted [8], especially the species of the genus *Meloidogyne* [8]. Some studies have
706 been developed with the purpose of evaluating the reproduction of *Meloidogyne* spp. in
707 medicinal species. From these studies, resistant species can be identified and used as
708 resistance sources genes, as well as their use in crop rotation systems aiming to decrease
709 the nematode population in infested areas.

710
711 [145] evaluated the reproductive rates of *Meloidogyne incognita* race 2 and *Meloidogyne*
712 *javanica* in eight species of medicinal plants. Evaluations were performed based on egg
713 mass indexes and nematode reproduction factors. In relation to the results, *Achillea*
714 *millefolium* (millefolium), *Arctium lappa* (burdock), *Bryophyllum calycinum* (brasilian name:
715 “folha-da-fortuna”) and *Crassula portulaca* (balm) were non-efficient hosts or unfavorable to
716 both species. *Plectranthus barbatus* (forskohlii) and *Polygonum hydropiperoides* (swamp
717 smartweed) were efficient to reproduce the two species. *Achyrocline satureoides* (marcela)
718 and *Tropaeolum majus* (nasturtium) were efficient for *M. javanica* and not for *M. incognita*.

719
720 [146] tested the pathogenicity of *Meloidogyne javanica* in sweet basil (*Ocimum basilicum*),
721 holy basil (*Ocimum sanctum*), common balm (*Melissa officinalis*), and tropical White weed
722 (*Ageratum conyzoides*). The level of infection by the nematode was quantified by the
723 number of galls and eggs per root system, expressed on a scale of 0 to 5. The four species
724 evaluated were highly susceptible, with all the inoculated plants receiving a grade 5 in
725 relation to the gall index and to the number of eggs. However, only sweet basil was resistant
726 to infection by *M. javanica*, presenting a significant reduction in fresh weight and shoot dry
727 weight.

728
729 [147] evaluated four species of medicinal plants, in protected cultivation, regarding
730 hostability to *Meloidogyne incognita*. The reaction of the medicinal plants was done by
731 counting the eggs and juveniles of the roots, estimating the Host Susceptibility Index (ISH).
732 *Achillea millefolium* (bloodwort), *Arctium lappa* (greater burdock) and *Bryophyllum calycinum*
733 (brasilian name: “folha-da-fortuna”), were resistant, while *Ageratum conyzoides* were
734 susceptible.

735
736 In a study published in 2004 [148], 22 species of medicinal plants were evaluated for
737 hostability to *M. hapla* in greenhouse. The evaluations were performed based on the gall
738 index and the reproduction factor. Twelve of these species [*Angelica dahurica* (angelica),
739 *Arctium lappa* (greater burdock), *Astragalus membranaceus* (astragalus), *Carthamus*
740 *tinctorius* (safflower), *Codonopsis lanceolata* (deodeok), *C. pilosula* (dang shen), *Coriandrum*
741 *sativum* (coriander), *Glycyrrhiza uralensis* (chinese liquorice), *Leonurus sibiricus* (honey
742 weed), *Ligusticum tenuissimum* (vinum), *Ostericum koreanum* (tyosen-nodake) and
743 *Peucedanum japonicum* (coastal hog fennel)] were registered as susceptible to *M. hapla*.
744 *Cassia tora* (foetid cassia), *Coix lacryma-jobi* (adlay) and *Perilla frutescens* (beefsteak plant)
745 were immune, neither galls nor nematodes were found in these plants. *Achyranthes japonica*
746 (japanese chaff flower), *Atractylodes japonica* (japanese Atractylodes), *Hibiscus manihot*
747 (aibika), *Ricinus communis* (castor bean) and *Sophora flavescens* (shrubby sophora) were
748 considered resistant. *Doellingeria scabra* (korean aster) and *Agastache rugosa* (korean mint)
749 were susceptible and hypersensitive, respectively.

750
751 In the study of the reaction of 14 species of medicinal plants to *Meloidogyne paranaensis*, [8]
752 evaluated the multiplication of the nematodes in the roots by counting eggs and J2 in the
753 root systems, calculating the reproductive factors (RF). Plants with $FR < 1$, susceptible with
754 $FR \geq 1$ and immune to $FR = 0$ were considered resistant. The species that showed
755 resistance to *M. paranaensis* were *Taraxacum officinale* (common dandelion), *Matricaria*
756 *recutita* (German chamomile), *Melissa officinalis* (common balm), *Hyssopus officinalis*
757 (hyssop), *Lippia alba* (bushy lippia), *Kalanchoe pinnata* (cathedral bells) and *Sedum*

758 *praealtum* (green cockscomb); and the susceptible ones were: *Ocimum basilicum* (sweet
759 basil), *Plectranthus barbatus* (forskohli).

760

761 When evaluating 15 species of medicinal plants for hostability to *M. javanica* and *M.*
762 *incognita*, [149] they counted the number of galls and eggs, as well as the determination of
763 the reproduction factor. In this study with inoculation of *M. incognita*, chamomile was
764 susceptible, with FR = 1.64, being considered a good host, the other plants were considered
765 resistant, with FR <1. For *M. javanica*, all plants behaved as resistant, with FR <1, with
766 immunity to myrrh, rue and balm, which presented FR = 0.

767

768 The reaction of seven species of medicinal plants to *M. paranaenses* was studied by [150]
769 with evaluations through determinations of gall index and reproduction factor. The
770 accessions of *Pfaffia glomerata* (brazilian ginseng), *Hypericum perforatum* (eola-weed) and
771 *Melissa officinalis* were highly susceptible to *M. paranaensis*. *Pogostemon cablin* (patchouly)
772 had an intermediate reaction and was classified as susceptible. *Artemisia annua* (sweet
773 sagewort) and *Catharanthus roseus* (madagascar periwinkle) were highly resistant, and
774 *Cordia verbenacea* (brasilian name: "erva-baleeira"). *Catharanthus roseus* was distinguished
775 because it had a high gall index but did not allow the multiplication of the nematoid.

776

777 [6] evaluated the susceptibility of 30 species, being 20 ornamental and 10 medicinal
778 [*Peumus boldus* (boldo), *Ocimum gratissimum* (african basil), *Mentha arvensis* (wild mint),
779 *Mentha villosa* (mojito mint), *Plectranthus amboinicus* (mexican mint), *O. basilicum* (sweet
780 basil), *Rosmarinus officinalis* (rosemary), *Cymbopogon citratus* (lemon grass), *Lippia alba*
781 (bushy lippia), *C. winterianus* (cymbopogon grass)) to *M. incognita*. The evaluation of the
782 plants was performed by measuring the number of galls and eggs, egg mass index,
783 reproduction factor and reproduction factor reduction. From these variables, the reaction of
784 the plants to the nematode was classified. With regard to medicinal plants, the species *M.*
785 *villosa*, *L. alba* and *C. citratus*, *C. winterianus* and *P. boldus* did not present galls in their root
786 systems, the other species behaved as moderately susceptible, with few galls and, or
787 females isolated at their roots.

788

789 **5. FINAL CONSIDERATIONS**

790

791 The identification of resistance sources among medicinal species is an important alternative
792 in the management of cultivated areas infested with nematodes. In cultivation systems these
793 species can be used in consortium or rotation with other cultivated species aiming at
794 reducing the inoculum concentration in the field, reducing agricultural losses. In addition, the
795 knowledge of these possible resistance sources, in the medium and long term, can be used
796 as a subsidy for breeding programs in order to select materials for nematode resistance of
797 the genus *Meloidogyne*. The literature indicates that resistance attributed to medicinal plants
798 comes from substances derived from the secondary metabolism of these species, such as
799 phenolic compounds, steroids, triterpenes, anthraquinones, flavonoid glycosides, saponin
800 glycosides, condensed tannins, hydrolysable tannins and sugars. Knowing which of these
801 substances are involved in resistance to root-knot nematodes, it is possible to isolate these
802 compounds for later use in the formulation of products capable of controlling this pathogen.

803

804 **COMPETING INTERESTS**

805

806 Authors have declared that no competing interests exist.

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