Authors: Andressa Lumi Akabane\textsuperscript{1} Isabella Parente Almeida\textsuperscript{1} João Carlos Lopes Simão\textsuperscript{1}

\textsuperscript{1} Division of Dermatology, Department of Internal Medicine, Faculdade de Medicina de Ribeirão Preto, Universidade de São Paulo - São Paulo(SP), Brazil.

Correspondence to: Hospital das Clínicas de Ribeirão Preto da Universidade de São Paulo Av. Bandeirantes, 3900 Monte Alegre – CEP: 14049-900 Ribeirão Preto-SP
E-mail: andressa.akabane@gmail.com

Received on: 10/11/2016
Approved on: 20/04/2017

Study conducted at the Division of Dermatology, Department of Internal Medicine, Faculdade de Medicina de Ribeirão Preto, Universidade de São Paulo - São Paulo(SP), Brasil.

Financial support: The product used in the study was provided by the company Melora.

Conflict of interests: The study was sponsored by Melora, but the methodology, execution and analysis of the results were performed by the institution researchers, with no participation of the company.

ABSTRACT

Introduction: Melasma is a pigmentation disorder that mainly affects women of childbearing age with high phototypes. Polypodium leucotomos has antioxidant, photoprotective and immuno-modulatory activity, and can be considered as an adjunctive treatment for melasma.

Objective: To evaluate the efficacy, in relation to quality of life and objective improvement, of the use of Polypodium leucotomos in the treatment of melasma.

Methods: Prospective and individualized study. Nine volunteers with melasma were submitted to treatment with Polypodium leucotomos for 45 days. MELASQoL, DLQI and MASI scores were calculated at the beginning (D0) and 45 days after (D45). Analysis of variance ANOVA with Tukey post-test for comparison between D0 and D45 (p <0.05).

Results: All patients were females, mean age of 37.18 ± 6.78 years. Family history of melasma in 55.6%; 88.9% with unprotected photoexposure and use of estrogen. After 45 days of treatment with Polypodium leucotomos there was a significant reduction of MELASQoL and DLQI (p <0.05) and improvement of MASI in 55.6% of the patients.

Conclusions: There was improvement of MASI in 55.6% of patients after 45 days of treatment. In spite of the slight improvement in MASI, there was a reflex in the improvement of quality of life scores (DLQI and MELASQoL).

Keywords: melanosis; Polypodium; quality of life; indicators of quality of life.

RESUMO

Introdução: Melasma é distúrbio de pigmentação que acomete principalmente mulheres em idade fértil com fototipos elevados. Polypodium leucotomos tem atividade antioxidante, fotoprotetora e imunomodulatória, sendo tratamento adjuvante do melasma.

Objetivo: Avaliar a eficácia, em relação à qualidade de vida e à melhora objetiva, do uso de Polypodium leucotomos no tratamento do melasma.

Métodos: Estudo prospectivo e individualizado. Nove voluntárias portadoras de melasma foram submetidas ao tratamento com Polypodium leucotomos durante 45 dias. Escores MELASQoL, DLQI e MASI foram calculados no D0 e no D45. Realizou-se a análise de variância Anova com pós-teste de Tukey para comparação entre D0 e D45 (p < 0,05).

Resultados: Todas as pacientes eram do sexo feminino, com média de idade de 37,18 ± 6,78 anos, história familiar de melasma em 55,6%, e fotoexposição desprotegida e uso de estrogênio em 88,9%. Após 45 dias de tratamento com Polypodium leucotomos houve redução significativa do MELASQoL e DLQI (p < 0,05) e melhora do MASI em 55,6% das pacientes.

Conclusões: Houve melhora do MASI em 55,6% das pacientes após 45 dias de tratamento. A despeito da discreta melhora no MASI, houve reflexo na melhora dos escores de qualidade de vida (DLQI e MELASQoL).

Palavras-chave: melanose; Polypodium; indicadores de qualidade de vida; qualidade de vida.
INTRODUCTION

Melasma is a pigmentation disorder that affects mainly women of childbearing age with higher phototypes, due to the induced hyperactivation dermal melanocytes, especially by the ultraviolet radiation. It is a frequent condition in the general population, with a high impact in the quality of life.

Polypodium leucotomos (PL), a Polypodiaceae plant extract has antioxidant, photoprotective and immunomodulatory activity, being adjuvant in photoinduced dermatoses, including melasma.

The objective of the study was to evaluate the efficacy in regards to the quality of life, and the objective improvement with the use of PL for the treatment of melasma.

MATERIALS AND METHODS

A comparative, prospective and individualized analysis of melasma patients, submitted to a 45-day treatment with PL, conducted according to the ethical procedures advised by the Declaration of Helsinki. Eleven volunteers took one 250mg tablet of PL every 12 hours and continued using sunscreen with an SPF equal to or higher than 50, three times a day. The volunteers did not use any topical products containing acids or bleaching agents in the previous three months. Medical (MELAS-QoL and DLQI and MASI) and photographic assessments (Visia®, Canfeld Imaging System- Fairfield, EUA) of the volunteers were performed in D0 and D45. The use of the device Visia® enabled a most accurate photographic record of melasma changes through ultraviolet lamps (UV spots), and the device’s digital analysis system enabled the photographic comparison in the different times of the study. The patients signed an informed consent. Two patients were excluded from the study because they were not present in the final assessment.

The photographs were taken in a frontal view, 45° to the right and 45° to the left on days D0 and D45. MELASQoL and DLQI were filled out by the volunteers. MASI was calculated by three collaborating physicians not involved in the study, previously trained, after seeing the photos of the patients, with no knowledge of the time of picture taking (before and after).

For the comparison of the DLQI, MASI and MELASQoL scores, the variance analysis Anova was performed with Tukey post-test with a program, GraphPad Prism 7.0.

RESULTS

All volunteers were female, with a mean age of 37.18 ± 6.78 years and a mean course of the disease of 6.36 ± 5.26 years. The majority (55.6%) had family history of melasma. In regard to the phototypes, (Fitzpatrick): 33.3% phototypes I-II; 55.6%, III-IV; 11.1%, V-VI (Table 1).

The most prevalent worsening factors for melasma were: unprotected sun exposure (88.9%) and the use of estrogen (88.9%). Worsening of melasma after pregnancy was observed in two of the nine patients (Table 1).

There was a significant reduction in the MELASQoL before and after 45 days using PL (p = 0.0164), as well as a significant decrease in the DLQI score (p = 0.0483). Despite MASI improvement in 55.6% of patients after 45 days of treatment (Figures 1 and 2), overall, there was slight reduction of the mean on D45 (11.01 ± 6.22) compared to D0 (11.82 ± 7.4). However, there was no statistically significant difference (p = 0.799) (Table 2).

DISCUSSION

Melasma is more common in females (9:1) and affects 8.8% of Latin women of childbearing age. The mean age seen (37.18 years) was similar to other melasma studies.

### TABLE 1: Social, demographic and clinical data of melasma patients

<table>
<thead>
<tr>
<th>Data</th>
<th>N (%)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Current age (Mean ± SDM)</td>
<td>37.18 ± 6.78</td>
</tr>
<tr>
<td>Course of the condition (Mean ± SDM)</td>
<td>6.36 ± 5.26</td>
</tr>
<tr>
<td>History of Melasma</td>
<td></td>
</tr>
<tr>
<td>Yes</td>
<td>5 (55.6)</td>
</tr>
<tr>
<td>No</td>
<td>4 (44.4)</td>
</tr>
<tr>
<td>Fitzpatrick Phototype</td>
<td></td>
</tr>
<tr>
<td>I-II</td>
<td>3 (33.3)</td>
</tr>
<tr>
<td>III-IV-V</td>
<td>5 (55.6)</td>
</tr>
<tr>
<td>VI</td>
<td>1 (11.1)</td>
</tr>
<tr>
<td>Worsening factors</td>
<td></td>
</tr>
<tr>
<td>Pregnancy</td>
<td>2 (22.2)</td>
</tr>
<tr>
<td>Estrogen use</td>
<td>8 (88.9)</td>
</tr>
<tr>
<td>Unprotected sun exposure</td>
<td>8 (88.9)</td>
</tr>
</tbody>
</table>

### TABLE 2: Comparison of the scores MELASQoL, DLQI and MASI before and after 45 days of treatment with Polypodium leucotomos, represented as Mean±SDM

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Mean</th>
<th>SDM</th>
<th>Minimum</th>
<th>Maximum</th>
<th>p</th>
</tr>
</thead>
<tbody>
<tr>
<td>MELASQoL D0</td>
<td>53.44</td>
<td>8.46</td>
<td>34</td>
<td>63</td>
<td>0.0164</td>
</tr>
<tr>
<td>MELASQoL D45</td>
<td>38.89</td>
<td>14.08</td>
<td>19</td>
<td>63</td>
<td>0.0483</td>
</tr>
<tr>
<td>DLQI D0</td>
<td>10.22</td>
<td>3.42</td>
<td>7</td>
<td>17</td>
<td>0.799</td>
</tr>
<tr>
<td>DLQI D45</td>
<td>6.11</td>
<td>4.73</td>
<td>1</td>
<td>16</td>
<td></td>
</tr>
<tr>
<td>MASI D0</td>
<td>11.82</td>
<td>7.40</td>
<td>4.2</td>
<td>28.7</td>
<td></td>
</tr>
<tr>
<td>MASI D45</td>
<td>11.01</td>
<td>6.22</td>
<td>4.2</td>
<td>24.4</td>
<td></td>
</tr>
</tbody>
</table>

SDM = standard deviation of the mean. p>0.05.
This condition is particularly prevalent in darker phototypes, especially in Fitzpatrick III to VI. In the present study, 66.7% had phototype III-VI and 55.6%, III-IV.

The exact pathogenesis of melasma is not completely clear, being genetics, sun exposure and hormonal therapy possible causal factors. The majority (55.6%) had first degree relatives affected; in 88.9% there was unprotected sun exposure and the use of estrogen at some point in life, with worsening of melasma. In 22.2%, pregnancy was a worsening factor.

PL use was associated to improvement of the quality of life, DLQI and MELASQoL scores (p < 0.05). There was improvement in MASI in five (55.6%) of the patients in the study after 45 days of treatment, however, with an overall p > 0.05, with two (22.2%) with higher values and two (22.2%) with the same values upon completion of the study.

Standardized photographic records, such as Visia®, enable better patient follow-up. The comparative analysis of the device’s parameters referring to UV spots and polarized light, was consistent with the evolution of the MASI scores obtained, in the majority of cases. Difficulties such as lack of uniformity of the automatic masking and other concomitant dyschromias can, however, contribute to conflicting scores.

The slight MASI improvement reflected in a significant improvement of the quality of life scores (DLQI and MELASQoL). The subjective evaluation of improvement by the patient and by the examiner’s perception reflected independent results, consistent with the low correlation coefficients found between MELASQoL and MASI (0.17 to 0.36) in the literature. Similarly, there was a strong correlation of psychometric measures of the perception of the severity of the condition, such as MELASQoL and DLQI.
CONCLUSION

A slight improvement in MASI improved the quality of life scores significantly (DLQi and MELASQoL) for the patients in the study. The positive results of the adjunctive treatment with PL for melasma favor its use, being a very promising therapeutic alternative. However, new studies with a higher number of patients are needed to best evaluate the efficacy of this drug quantitatively.

REFERENCES


