Research Letters

Sterols and sterolins in Hypoxis hemerocallidea (African potato)

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Commercially available health supplements and herbal remedies containing sterols and sterolins, either from African potato (Hypoxis hemerocallidea) alone, or whether enriched with sterols and sterolins, are claimed to be efficacious in the treatment of a variety of ailments. Sterols and sterolins in African potato are purported to be the relevant constituents that are required for the therapeutic claims of such products. A patent describing the extraction of sterolins from African potato plant material has claimed that approximately 9 mg sterolins can be isolated from 100 g of an enriched aqueous African potato extract. Our analysis of African potato plant material and its sterol and sterolin content, when similarly prepared, shows that the measureable content of sterols and sterolins in African potato is far less than the amounts of these compounds that have been claimed to be necessary for therapeutic benefit. We conclude that therapeutic claims relating to sterol and sterolin content in African potato are unsubstantiated, in view of the extremely low content of such compounds that we have isolated from our plant material, and in products containing African potato, or extracts thereof.

Introduction

African potato (AP), Hypoxis hemerocallidea, apart from its perceived nutritional value, is of active medical interest, and is purported by many South Africans as possibly being the best-known medicinal plant. Although African potato is the common name, its underground growth is not a tuber, but a corm. Furthermore, the term African potato is also, and confusingly, attributed to another plant, Plectrantus esculentus. The name Hypoxis will therefore be used throughout this text when referring to the AP. Extracts of the corms have been ingested by man as a dietary supplement and for a diversity of ailments. In recent years, the increasing interest in functional foods and the use of phytosterols, their glycosides (sterolins) and steryl esters (>2 g/day) for reducing serum cholesterol and increasing immunity has resulted in these compounds regaining considerable attention. There are many herbal formulations containing sterols for this reason, most of which have been fortified with additional amounts of free sterols, stanols and sterolins. Commercially available health supplements and herbal remedies containing sterols and sterolins as well as Hypoxis, alone or enriched with sterols and sterolins, are claimed to be efficacious for a variety of ailments. In particular, purported sterol and sterolin content in products containing Hypoxis is identified to be the important feature for the therapeutic claims of these products.

Sterols and stanols are claimed to play an important role in the realm of health supplements, with extensive scientific argument for their prophylactic and therapeutic use for various medical conditions and ailments, such as atherosclerosis, benign prostatic cancer and colon cancer. However, many reports on the medicinal properties of sterols are largely based on in vitro analyses or unrealistic high in vivo dosages, making the therapeutic claims for these compounds disputable.

The average daily human dietary consumption of phytosterols is approximately 250 mg, mostly derived from vegetable oils, cereals, fruits and vegetables. Dietary phytosterols have been estimated to be almost twice this intake in the case of vegetarians. Phytostanols are much less abundant in nature than phytosterols and are consequently consumed in relatively lower amounts (c. 25 mg/day) in human diets. Phytosterols have also been used since the mid-1990s in strategies for lowering cholesterol, and for reducing the risk of cardiovascular diseases (CVD). The advent of the ‘statin’ compounds has resulted in a rapidly diminished use of phytosterol products. Lipophilic extracts of Hypoxis corms have been used in Europe for the treatment of prostate problems. These extracts are claimed to have anti-inflammatory activity, and to relieve the symptoms of prostate adenoma. The components claimed to be responsible for this effect are mixtures of phytosterols, and it is claimed that...

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\(\beta\)-sitosterol (Fig. 1) is the main active compound. A commercial product, Harzol\textsuperscript{9}, was launched in 1974, and consists of a combination of \(\beta\)-sitosterol and its glucoside, BSSG (Fig. 2) and has gained wide acceptance in Germany. The glucosides (including BSSG) were purported to be obtained from Hypoxis, and their action in the treatment of benign prostate hypertrophy was attributed to inhibition of 5\(\alpha\)-reductase, or to diminished binding of dihydrotestosterone within the prostate gland.\textsuperscript{9} Various other commercial preparations were subsequently launched, and were claimed to contain Hypoxis extracts, or other plant material, specifically for their sterol/sterolin content. One such formulation, initially purported to contain Hypoxis, was stated to contain sterols-to-sterolins in the mass ratio of 100:1. This specific ratio has been claimed to enhance the in vitro proliferative response of human T cells, more than individual sterols at the same concentration, and this claim was used to market and promote sales of this particular product.\textsuperscript{5–10} The Hypoxis component, however, was subsequently removed from this product, and the included phytosterols have been subsequently claimed to be sourced from elsewhere.

With the absence of convincing ethnomedical information relating to the efficacy of the 100:1 ratio of sterols-to-sterolin, coupled with the non-availability of data from other sterol and sterolin ratios, the situation raises doubt regarding the validity of the efficacy and therapeutic claims for such products. Claims of phytophsterol content in Hypoxis nevertheless continue to be made, such as a recent advertisement for Santjie Marx Products, which includes the statement: 'The plant sterols and sterolins that are essential to proper immune system function, are highly concentrated in the Africa Potato plant' (http://www.smarx-produkte.com/).

Early claims relating to the medicinal properties of \(\beta\)-sitosterol and its glucoside in Hypoxis have been reported in the literature.\textsuperscript{11–12} Although not substantiated by definitive pharmacological data, several patents have been filed on the usage and medicinal claims of sterolin-enriched extracts of Hypoxis for various therapeutic purposes.\textsuperscript{12–17} In particular, a patent filed by Pegel and Liebenberg in 1973\textsuperscript{18} claimed that the potency of an extract from Hypoxis species in the treatment of prostate hypertrophy was correlated with chemically non-defined phytophsterol glucosides (sterolins such as BSSG).

It should be noted that due to poor aqueous solubility and limited bioavailability of free phytosterols, their purported serum cholesterol-lowering effects were not always found to be consistent, and very high oral doses (up to 25–50 g/day) appear to be required for efficacy. The problems of solubility and bioavailability have led to many contradictory results in early clinical studies.\textsuperscript{3}

The highest yield recorded in the US patent entitled ‘Extraction of sterolins from plant material’, was about 9 mg sterolin from 100 g of an enriched aqueous extract of Hypoxis.\textsuperscript{17–20} The extractive value of the above aqueous isolate, defined as the ratio of BSSG (molecular weight, 552.89) with BSS (molecular weight, 414.65), 9 mg will stoichiometrically convert to about 6.8 mg \(\beta\)-sitosterol, which is far lower than the reported therapeutic dose required for efficacy.\textsuperscript{7} To date, except for the claims made in the aforementioned patent, to our knowledge there have been no research publications describing the isolation and analysis of sterols or sterolins from Hypoxis. Nor are there any scientific data to substantiate the amount of these phytoconstituents that is claimed to be required for therapeutic efficacy. Their presence in adequate therapeutic concentration in any species of the genus Hypoxis deserves scrutiny. The issue has been raised by Nicoletti \textit{et al.}\textsuperscript{24} and Hostettmann \textit{et al.}\textsuperscript{8}

Based on our argument, we propose that any nutritional or therapeutic value of Hypoxis is unlikely to be due to the presence of sterolins in Hypoxis or extracts therefrom, as final extractive recovery of these compounds is <0.01% w/w of enriched extract.

**Content of sterols and sterolins in Hypoxis**

A Hypoxis extract was prepared from Hypoxis plant material, in accordance with the example procedure recorded in the patent.\textsuperscript{16} We measured the content of sterols (\(\beta\)-sitosterol and stigmasterol) and BSSG recovered. Extractive procedure is similar to the Hypoxis decoctions prepared according to the common recipe used by traditional healers (\textit{sangomas}) in the Eastern Cape province of South Africa. Analysis involved a validated HPLC procedure, coupled with evaporative light-scattering detection for the qualitative and quantitative determination of sterols and BSSG.\textsuperscript{5} The aqueous extract did not show any detectable quantities of BSSG or BSS and this was confirmed by spiking the sample solutions with known concentrations of BSS and BSSG standards.

The content of sterols (stigmasterol, \(\beta\)-sitosterol and stigmastanol) and sterolins in the preparation outlined above is less than our analytical detection limit of 0.002% w/w. This equates to an amount of less than about 6 mg BSSG/100 g of aqueous extract, in contrast to the content of 9.01 mg/100 g aqueous extract, as given in the patent information. Notwithstanding the patent claims, we confirm a lower sterol content of Hypoxis, and we show that this so-called enriched extract of Hypoxis contains less than 0.002% sterolins.

We conclude that because the daily consumption by consumers...
Table 1. Content of sterols and sterolins in African potato.

<table>
<thead>
<tr>
<th>Sample no.</th>
<th>Sample profile</th>
<th>Amount</th>
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</thead>
<tbody>
<tr>
<td>1</td>
<td>Reported analytical profile of AP&lt;sup&gt;23&lt;/sup&gt;</td>
<td>Amount of sterolin (BSSG)/100 g of enriched aqueous extract of AP</td>
</tr>
<tr>
<td>2</td>
<td>Reported consumption of AP&lt;sup&gt;23&lt;/sup&gt;</td>
<td>Average daily consumption of traditional extract of AP</td>
</tr>
<tr>
<td>3</td>
<td>Current analytical method</td>
<td>Limit of detection for BSSG</td>
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<tr>
<td></td>
<td></td>
<td>Amount of BSSG in 100 mg fresh AP/ml</td>
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of *Hypoxis* is about 200–500 ml of such an aqueous decoction (typically containing about 37 mg fresh *Hypoxis* per ml, or c. 18.5 g of *Hypoxis* plant material),<sup>26</sup> the daily intake of BSS and BSSG, if present in the decoction, will be negligible (about a milligram) and will be inadequate for any therapeutic efficacy (Table 1).

Health-product preparations of *Hypoxis hemerocallidea* are readily available from retail outlets, and more recently via the internet, and the suppliers and manufacturers of these products continue to make a variety of claims regarding the efficacy of such products. It is the purpose of this communication to present a scientific appraisal of these claims, and to afford evidence that they are suspect.

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