

International Neuropsychiatric Disease Journal 5(4): 1-4, 2016; Article no.INDJ.22181 ISSN: 2321-7235, NLM ID: 101632319



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Resveratrol Reduces C-reactive Protein and Cholesterol in Schizophrenia

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Authors' contributions

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

Article Information

DOI: 10.9734/INDJ/2016/22181 <u>Editor(s):</u> (1) Weike Mo, Scientific Director for Pharmacogenetics, Molecular Testing Labs, USA. <u>Reviewers:</u> (1) Ds sheriff, Benghazi University, Libya. (2) Pietro Scicchitano, University of Bari, Italy. Complete Peer review History: <u>http://sciencedomain.org/review-history/11890</u>

Case Study

Received 21st September 2015 Accepted 3rd October 2015 Published 19th October 2015

ABSTRACT

Patients with schizophrenia (SZ) are generally found with overweight and obesity and several metabolic disorders. Additionally these patients have less life expectancy, with cardiovascular disease (CVD) the main cause of increased mortality in these patients. This is a case-report study of a male, 46 years old, with Diagnosis of SZ following a 1-month resveratrol supplementation program. In summary, we observed association of resveratrol supplementation (200 mg/day) with reduced C-reactive protein (CRP), and reduced cholesterol and triglycerides. The benefits included the improvement in Framingham score and mean reduction of 3% of risk of a heart attack over the next 10 years.

Keywords: Schizophrenia; Cardiovascular Diseases; Inflammation; Resveratrol.

1. BACKGROUND

Patients with schizophrenia (SZ) are generally found with overweight and obesity and several

metabolic disorders. Additionally these patients have less life expectancy, with cardiovascular disease (CVD) the main cause of increased mortality in these patients [1].

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Several studies support a link between inflammation and atherogenesis. Low-grade inflammatory status in overweight persons has been proposed as one of the mediating processes in metabolic disease development, such as CVD and diabetes [2]. Elevated Creactive protein (CRP) is predictor of CVD events in the general population [3]. To prevent these factors, many food compounds have been to have anti-inflammatory reported and antioxidant effects in human metabolism [2] and there are some evidences that resveratrol supplementation have a significant potential benefits to treat disorders like CVD [4].

Resveratrol is a natural polyphenolic compound with cardioprotective, anticancer and antiinflammatory properties [4], involved in antiatherogenic activities and vasculoprotection [5], identified in more than 70 species of plants, including grape-vines (*Vitis vinifera*), mulberries, and peanuts. Additionally, a recent meta-analysis of 10 studies suggested that the consumption of this polyphenol lowered CVD risk [6]. According to this line of evidences, the objective of this study was to determine the efficacy of resveratrol supplementation on inflammatory status and CVD risk prevention in patient with SZ.

2. CASE REPORT

This is a case-report study of a male, 46 years old, with Diagnosis of SZ by Structured Clinical Interview for DSM-IV-Axis I Disorders (SCID-I), following a 1-month resveratrol supplementation program, covered by Public Health Assistance at the Schizophrenia Program of Hospital de Clinicas de Porto Alegre (HCPA), Brazil. The research participant provided signed, informed consent and was on stable dose of atypical antipsychotics for 6 months. This research study was approved by the Research Ethics Committee/HCPA (register number: 110553).

Clinical evaluation included anthropometric measures (weight, height, waist circumference, body mass index – BMI), smoking status, physical activity, biomarker of inflammation (CRP), fasting glucose and lipid profile (cholesterol, LDL, HDL, triglycerides). All measures were taken on the first and the last day (day 1 and day 30) in a 1-month follow-up.

The subject took 2 dietary supplements each day (200mg of resveratrol). Resveratrol was obtained from a manipulation pharm (*trans*-resveratrol, 98% purified) in Porto Alegre, RS, Brazil. He was

instructed to take the first supplement after the baseline measurements (day 1) and the last supplement in the end of 4 weeks (day 30). The subject was instructed to abstain from foods containing substantial amounts of resveratrol (e.g., wine, red grapes, peanuts, and berries) and was advised not to take any other food supplements and maintain their usual diet and physical activity during the study period. To ensure that the subjects adhered to the study protocol the team used two practices: a) telephone calls every week over the study; b) pill count in the last day (day 30).

The characteristics of the subject are described in Table 1. After 30 days resveratrol supplementation, the subject displayed reduced waist circumference and measures of cholesterol, LDL, HDL, triglycerides and CRP, with a slight increase in weight, BMI, body fat percentage and number of smoked cigarettes. There were no changes in medication, exercise and glucose, without significant adverse events.

3. DISCUSSION

In summary, we observed association of resveratrol supplementation (200 mg/day) with reduced CRP, and reduced cholesterol and triglycerides. The benefits included the improvement in Framingham score [7,8] and mean reduction of 3% of risk of a heart attack over the next 10 years.

In a similar research, resveratrol inhibited astrocyte production of CRP, which plays a role in a variety of chronic inflammatory disorders. This suggests that resveratrol may be an effective therapeutic agent in neurodegenerative diseases initiated or maintained by inflammatory processes [9].

A recent study showed that nutraceuticals, including resveratrol, are effectively able to reduce the burden of the atherosclerosis process. The main mechanisms that explain such a positive action on the cardiovascular system are not well understood but the result supports the hypothesis about the use of primary nutraceuticals cardiovascular in prevention protocols to reduce the overall burden of cardiovascular disease morbidity and mortality [10]. According to this line of evidences, the case provides additional support for the protective effect of resveratrol in different pathological conditions, associated with reduced risks of CVD, cancer and diabetes, maybe through its anti-inflammatory effect [4].

| Characteristics | Day 1 (Baseline) | Day 30 | Results |
|--------------------------|------------------|--------|--------------|
| Age (years) | 46 | 46 | = |
| Clozapine dose (mg) | 600 | 600 | = |
| Smoking (yes/not) | Yes | Yes | = |
| Number of cigarette/day | 6 | 10 | ^ |
| Exercise (yes/not) | Yes | Yes | = |
| Weight (kg) | 83,2 | 83,7 | ^ |
| BMI (kg/m²) | 26,11 | 26,26 | ^ |
| Waist circumference (cm) | 111 | 110 | $\mathbf{+}$ |
| Body fat percentage (%) | 29,7 | 30,1 | ^ |
| Glucose (mg/dL) | 103 | 103 | = |
| Cholesterol (mg/dL) | 187 | 161 | $\mathbf{+}$ |
| LDL (mg/dL) | 110,4 | 94,8 | $\mathbf{+}$ |
| HDL (mg/dL) | 36 | 34 | $\mathbf{+}$ |
| Triglycerides (mg/dL) | 203 | 161 | $\mathbf{+}$ |
| CRP (mg/dL) | 8 | 4 | $\mathbf{+}$ |
| Framingham score (%) | 10 | 7 | $\mathbf{+}$ |

Table 1. Characteristics of subject with schizophrenia in resveratrol supplementation

BMI: Body Mass Index; CRP: C-reactive Protein

This case-report is a pilot study and there are some limitations. The case was selected by convenience, and could represent a special case, with additional unidentified protective factors. Because of this, it is important to confirm these results with larger samples. Second, it may be necessary assessment of additional blood biomarkers (e.g. cytokines and oxidative stress) to increase generalizability of effects of resveratrol add-on.

Despite these limitations, the case provides additional effect of resveratrol supplementation with a decrease in lipid profile, in inflammation, and may help prevent CVD risk. We think this can improve therapeutic and clinical outcome, and individual quality of life.

4. CONCLUSION

In conclusion, we have shown that 4 weeks of a resveratrol supplementation (200 mg/day) may reduce inflammation and improve lipid profile. These findings indicate that resveratrol deserves additional attention for the clinical care of SZ, by its role in comorbidity prevention and protective effect over CVD in SZ. Further studies in larger cases are needed to implement the actual findings associated with this hypothesis.

COMPETING INTERESTS

Authors have declared that no competing interests exist.

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Zortea et al.; INDJ, 5(4): 1-4, 2016; Article no.INDJ.22181

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Peer-review history: The peer review history for this paper can be accessed here: http://sciencedomain.org/review-history/11890