The association between smoking and the prevalence of metabolic syndrome and its components in patients with psoriasis aged 30 to 49 years

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Abstract

Introduction: Cigarette smoking may exacerbate and cause psoriasis. Moreover, smokers are more likely to develop insulin resistance and metabolic syndrome (MS).

Aim: To assess the prevalence of MS and its components in patients with psoriasis, who smoke, compared with the general Polish population of smokers.

Material and methods: We studied 29 patients with psoriasis (female = 9, male = 20), smokers, aged 30 to 49 years. Metabolic syndrome and its components were assessed using the IDF definition and compared to the results obtained in a representative sample of adult Poles in the NATPOL 2011 study in the same age group, including smokers.

Results: The results have shown that patients with psoriasis are more likely to be smokers ($p < 0.0034$) and the frequency of smoking in men is approximately 25% higher than in males of the control group ($p < 0.0017$). The prevalence of MS in patients with psoriasis who smoke was 27.58% and in the control group 25.2% ($p > 0.05$).

Mean body mass index was 26.07 kg/m$^2$ in psoriasis patients and 25.59 kg/m$^2$ in the control group ($p > 0.05$), and abdominal obesity was 88.82 cm and 90.02 cm ($p > 0.05$), respectively. There were no differences in hypertension (34.48% vs. 31.6%, $p < 0.05$) and mean HOMA-IR (1.80 vs. 1.77, $p > 0.05$). In lipid parameters, the differences were observed only in women with psoriasis – higher levels of HDL, triglycerides and ApoB/ApoA1 index compared with addicted women in the control group.

Conclusions: Men with psoriasis are more often addicted to smoking. Women with psoriasis who smoke have often disturbances of the lipid profile.

Key words: psoriasis, metabolic syndrome, insulin resistance, smoking.

Introduction

The presence of the same genetic predisposition is not sufficient for the development of psoriasis. Provocative factors that initiate a cascade of immune disorders play an important role in the formation of lesions.

Cigarette smoking is one of the factors that may exacerbate and cause psoriasis. The risk of psoriasis is higher in present and former smokers compared with those who never smoked [1–3]. Some studies confirm the relationship between smoking intensity and the severity of psoriasis [3, 4]. Smoking releases large amounts of free radicals and is a cause of “oxidative stress” [1, 3, 5].

The free radicals stimulate, regardless of cytokines, cell signalling pathways active in psoriasis including mitogen-activated protein kinase (MAPK), nuclear factor-$\kappa$B (NF-$\kappa$B), Janus kinase and JAK-STAT pathway [5]. Moreover, nicotine stimulates dendritic cells, macrophages and keratinocytes. They release cytokines, mainly tumor necrosis factor-\(\alpha\) (TNF-\(\alpha\)), IL-6, activating the T-cells and sustaining the course of chronic psoriasis [1, 5, 6]. Interleukin-6, in turn, helps to increase the production of C-reactive protein (CPR) in liver, which can lead to systemic inflammation [6] and contribute to the onset of metabolic disorders. Nicotine can also modulate the functional capacity of dendritic cells as antigen present-
ing cells (APC) by an increased expression of costimulatory molecules (CD86, CD40), adhesion molecules (LFA-1, CD54), and the expression of major histocompatibility complex class II (MHC-II) [1, 2, 4]. Furthermore, cigarette smoking increases the expression of genes predisposing to psoriasis, including HLA-Cw6, HLA-DQA1*0201 and CYP1A1 [5] and smokers with HLA-Cw6 have an 11 times greater risk of developing psoriasis compared to non-smokers without HLA-Cw6 [7].

Cigarette smoking, besides hypertension and hypercholesterolemia, is a major risk factor for cardiovascular disease [8]. It is the cause of severity of inflammation, endothelial dysfunction, atherogenic dyslipidaemia and coagulation disorders, resulting in the initiation and progression of atherosclerosis [9]. Smoking, inter alia, increases leukocyte count, and serum concentration of CRP, IL-6 and TNF-α [9].

According to the concept of the ‘psoriatic march’, which has been presented in recent years, psoriasis, obesity and smoking cause inflammation which influences the rapid development of atherosclerosis in patients [10]. Therefore, an attempt was made to assess the impact of psoriasis on prevalence of metabolic disorders in smokers.

Aim

The aim of this study was to assess the impact of cigarette smoking on metabolic syndrome (MS) and its components: insulin resistance, atherogenic dyslipidaemia, abdominal obesity and hypertension in patients with psoriasis aged 30 to 49 years, compared to Poles, who participated in the NATPOL 2011 study in the same age group.

Material and methods

The study included 62 patients with psoriasis aged 30 to 49 years, including 29 smokers (9 women and 20 men) treated at the out-patient clinic and the Department of Dermatology, Municipal Hospital in Olsztyn. Psoriatic patients with chronic and acute inflammatory diseases other than psoriasis, cancer, a history of cardiovascular complications and heart, kidney or liver failure (n = 1) were excluded from the study.

The control group consisted of 861 people of the Polish population aged between 30 and 49 years, including 250 smokers (101 women and 149 men) with a negative history of psoriasis. These people were included in the NATPOL research project in 2011 conducted by a team from the Hypertension and Diabetes Clinic, University Clinical Centre in Gdansk. The project involved the study of a representative group of 2400 Polish citizens in 2010–2011 and the prevalence of hypertension and other risk factors for cardiovascular disease, including lipid disorders and diabetes, among these people were estimated. A detailed description of the sampling will be published in “Polish Cardiology”.

It has been assumed that smoking ≥ 5 cigarettes per day is an addiction.

The following parameters were estimated in all subjects: body mass index (BMI), waist circumference measured in centimetres [11], blood pressure, according to the Polish Society of Hypertension guidelines 2003. The dermatological examination assessed the severity of psoriasis – PASI [12].

The patients had laboratory tests done under fasting conditions, 12 h after the last meal, evaluating the lipids (HDL, triglycerides, apolipoprotein A1 and B) and carbohydrate metabolism (glucose, insulin) and CRP. Insulin resistance index HOMA-IR (homeostasis model assessment-insulin resistance) was calculated [13] and to evaluate lipids, ApoB/ApoA1 indexes were estimated [14].

Metabolic syndrome was diagnosed using the International Diabetes Federation (IDF) criteria modified in 2009 [15], in which abdominal obesity (for Europeans: ≥ 94 cm for men, ≥ 80 cm for women) is not a prerequisite for the diagnosis, but it is one of the five criteria. The diagnosis of MS is about meeting three of five criteria (abdominal obesity, triglycerides level ≥ 150 mg/dl or treatment of hypertriglyceridaemia, HDL < 40 mg/dl in men and < 50 mg/dl in women, blood glucose ≥ 100 mg/dl or pharmacotherapy of hyperglycaemia and blood pressure: systolic ≥ 130 mm Hg and/or diastolic blood pressure ≥ 85 mm Hg or antihypertensive drug treatment) [16].

Statistical analysis

Statistical analysis of the results was performed using the statistical software package StatisticaPL v. 10. Analysis of variance and correlation was used to verify the hypotheses.

Results

Assessment of smoking in the study group compared to the NATPOL 2011 control group

The results showed that patients with psoriasis are more likely to smoke cigarettes compared to the NATPOL 2011 control group. These differences were also found considering patients’ gender: the frequency of smoking is 25% higher in males with psoriasis (Table 1).

Correlation between the severity of psoriasis (PASI) and cigarette smoking

Correlation analysis of PASI (with and without considering patients’ gender) and smoking showed no significant correlation between them. However, there was a significant correlation between smokers and alcohol drinkers.
Many studies have confirmed the association between smoking and psoriasis, especially pustular psoriasis. A study conducted by Naldi et al. has shown that one in five cases of psoriasis has been associated with smoking. Nicotine addiction may initiate psoriasis in genetically predisposed individuals and influences the disease severity. The risk increases with the number of cigarettes smoked daily and is higher in women than men: it is about 2.5 to 3.3 times higher among women who smoke more than 20 cigarettes per day and about 1.7 times higher among smoking men [1, 5]. Li et al. assessed the correlation between smoking and the prevalence of psoriasis in the U.S. Population (Nurses’ Health Study). Smokers, who smoked 1–14 cigarettes per day, had a relative risk of psoriasis of 1.81, those who smoked 15–24 cigarettes per day had 2.04, and those who smoked 25 or more – up to 2.29 [17]. The authors have also observed a gradual disease risk reduction after smoking cessation. The risk of psoriasis was particularly increased in heavy smokers and those with long-term addiction and it was similar for both genders [17]. However, there are also controversial reports. Chinese research showed that smoking is a risk factor for psoriasis only among men [18].

Our own research showed a higher percentage of people addicted to smoking among patients with psoriasis, especially in males (25% higher). This can confirm the aforesaid effect of smoking on psoriasis [1, 5, 19].

An interesting observation in our study is the frequent coexistence of smoking habit and alcohol consumption in patients with psoriasis. In the literature, alcohol consumption has been described as a factor responsible for triggering psoriasis, but it is said that smoking increases the risk of the onset of the disease [20]. Naldi says that smokers who drink are twice as likely to develop the disease as non-smokers and non-drinkers [2]. It should be noted that psoriasis can have an impact on self-esteem and suffering from the disease can cause stress and the patients are more likely to turn to alcohol and cigarettes [19].

### Discussion

#### Incidence of smoking in psoriatic patients

Many studies have shown that smoking increases the risk of developing MS. Nakanishi et al. found that the risk of developing MS was from 1.07 to 1.66 higher for smokers than non-smokers and it depended on the number of cigarettes smoked [18, 21, 22]. And smoking has an effect on metabolic disorders up to 20 years after quitting [21, 22]. However, there are also controversial reports. Polish studies by Chlebus et al. found a negative correlation between the prevalence of the MS and addictions [23].

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### Effect of cigarette smoking on metabolic syndrome and its components

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It was found that although people with psoriasis were more likely to smoke, the association between MS and psoriasis was not related to smoking [24].

Impact of smoking on obesity

It is believed that smoking can control weight. This mechanism is explained by the influence of nicotine on the increase in the resting metabolic rate and decreased appetite [22, 25]. Smoking 24 cigarettes per day increases energy expenditure by 10%. However, the comparison of waist circumference or abdominal obesity shows different results, and smokers were found to have the largest waist circumference [22, 25].

This phenomenon can be explained by the influence of nicotine on insulin resistance, predisposing to obesity and endocrine dysfunction among smokers. Long-lasting elevated catecholamines and growth hormone levels decrease the number of insulin receptors and lead to pancreatic cells toxicity [22, 25].

Moreover, visceral adipose tissue is sensitive to cortisol, which is elevated in the serum of smokers because of an increased sympathetic nervous system activity. Furthermore, the elevation in testosterone and low oestrogen levels lead to the increase in visceral obesity in menopausal women. Women who smoke have lower levels of bioavailable oestrogen [22]. The substances found in tobacco smoke also damage the granulosa cells of ovaries, causing a decrease in oestrogen and an increase in testosterone [26], resulting in abdominal obesity.

There is a lot of controversy about it. Mizuno et al. found no differences in waist circumference among non-obese patients, smokers or non-smokers. However, the waist circumference was significantly larger in obese patients who smoked compared to non-smokers [27].

Our own research showed no differences between the group of patients and the NATPOL group, considering patients’ gender.

Effect of smoking on blood glucose and insulin resistance

Numerous studies have shown that smoking reduces insulin sensitivity and causes type 2 diabetes and the longer the duration the higher the risk [22, 25, 27]. The study by Manson showed that the risk of diabetes increased with the number of cigarettes smoked per day: 20 cigarettes per day increased the risk by 50%, and more than 20 up to 70% [28]. Smoking cessation improves insulin sensitivity, despite weight gain.

From 5 to 10 years after quitting smoking it is the same as in people who have never smoked [22, 28]. The Physicians’ Health Study, which analysed over 22 thousand male physicians aged 40–84 who were addicted to smoking, showed that type 2 diabetes is more than twice as common in smokers as in non-smokers [29]. A similar correlation was found in the Nurse Health Study. The risk of developing diabetes was 1.4 for women who smoked more than 25 cigarettes per day [25].

There are reports which do not confirm such a correlation. The study by Godsland et al. showed that smokers had lower insulin levels after an oral glucose load than non-smokers [30]. Similar results were obtained by Szurkowska et al. identifying lower insulin levels in the fasting state and after a glucose load in smokers. Also Masuilli et al. found no effect of smoking on insulin resistance [23].

Our own research found no differences between the average glucose values and HOMA-IR among the psoriatic patients and the NATPOL group, considering patients’ gender.

Impact of tobacco smoking on lipid metabolism

Smoking modifies the lipid profile, leading to an increase in plasma levels of triglycerides (TG) and lower levels of HDL cholesterol [6, 21, 31–34]. Moreover, this correlation seems to be dependent on the number of smoked cigarettes [31]. Nicotine stimulates processes of lipolysis, leading to an increase in free fatty acids [21]. This reduces lipoprotein lipase activity and increases 3-hydroxy-3-methylglutaryl-CoA and glucose-6-phosphatase, which leads to lipid disorders [21]. In addition, there is an increased lipid oxidation and circulating lipid peroxidation products and antibodies against ox-LDL are significantly elevated in smokers. It is common knowledge that ox-LDL are antigens initiating the process of plaque formation [6].

The results have shown higher levels of HDL cholesterol in smoking women with psoriasis than in the control group, but also higher levels of triglycerides and ApoB/ApoA1. No such differences were seen among non-smokers.

Impact of tobacco smoking on hypertension

The effect of smoking on blood pressure is also controversial. Frati et al. showed that blood pressure and heart rate increase during smoking, but this effect disappears within an hour [21]. Moreover, Weitzman et al. found that active smoking was associated with a lowering of blood pressure [6, 34]. Mild reduction in blood pressure in smokers may be related to decreased body weight [35]. This observation is supported by the weight gain and increased blood pressure among former smokers versus that observed among people who never smoked [35]. A vasodilator effect of cotinine, the major metabolite of nicotine, also may help to lower blood pressure [34].

According to other reports, the incidence of hypertension is increased among those who smoke 15 or more cigarettes per day [36]. Chronic smoking causes arterial stiffness, which may persist for several years after smoking cessation [36]. Hypertension among smokers may be related to chronic sympathetic nervous system activity of
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Conclusions

Cigarette smoking among male patients with psoriasis is 25% higher than in the general Polish population. Our study confirms that cigarette smoking may be a provoking factor for psoriasis, especially in men. Women with psoriasis who smoke had higher levels of HDL cholesterol, triglycerides and ApoA1/ApoB index than smoking women in the Polish population. There were no differences in the prevalence of the MS, obesity, insulin resistance and hypertension between patients with psoriasis who smoke and the Polish population.

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Conflict of interest

The authors declare no conflict of interest.

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