Diabetes and OSA

Diabetes and sleep apnoea: a hidden epidemic?

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A diagnosis of OSA should be considered in patients with type 2 diabetes

It is now well established that obstructive sleep apnoea (OSA) is associated with an increased risk for cardiovascular disease, and that this is probably related to the fact that patients with sleep apnoea are likely to have a high prevalence of the risk factors that comprise the metabolic syndrome—namely, central adiposity, dyslipidaemia (low HDL cholesterol, raised triglycerides), high blood pressure, insulin resistance, and hyperglycaemia—even after correction for adiposity.\(^1\)\(^-\)\(^3\) Type 2 diabetes is also typically associated with a similar metabolic profile, so it is not surprising that the prevalence of diabetes in obese hypertensive patients with sleep apnoea may be as high as 40%.\(^4\) Given that both type 2 diabetes and OSA are highly prevalent in the general population, it is important to establish the prevalence of OSA in people with type 2 diabetes as this may represent a significant population with undiagnosed sleep apnoea that may be amenable to treatment.

In this issue of *Thorax* West and colleagues\(^5\) have attempted to establish the prevalence of OSA in a community and hospital based population of men with established type 2 diabetes. Initial screening was by questionnaire to over 1600 men with diabetes, which achieved an overall response rate of 56%; this identified populations of “high risk” and “low risk” subjects based on self-reported height, weight, and sleep related symptoms. A subset of each group was invited for a formal sleep study. OSA was found in 31% of high risk patients and 13% of low risk patients; the overall prevalence of OSA in the diabetes population was estimated at 23% compared with 6% in a community based prevalence study conducted a few years earlier.

This study does have limitations related to possible selection bias—for example, the prevalence of OSA in non-responders to the questionnaire is unknown, and most of the patients with diabetes were cared for in secondary care and may have had more severe disease than would be expected in a purely community based sample. Furthermore, the use of historical controls at a time when the prevalence of obesity in the population was rising rapidly may have underestimated the current prevalence of OSA in the non-diabetic population. Nevertheless, it seems likely that the overall conclusions of this study are valid, and that OSA is highly prevalent in men with type 2 diabetes.

As expected, OSA was more common in the “high risk” group who had more symptoms and were more likely to be obese, but there was only a weak correlation with glycaemic control as assessed by HbA1c, which is perhaps unsurprising given the confounding effects of oral hypoglycaemic drugs and insulin that are used to treat diabetes. It is disappointing that the questionnaires used to identify high and low risk groups for likelihood of OSA were relatively insensitive; however, the high risk group had a greater proportion of patients who reported sleepiness, and this is the group who would be offered treatment under current guidelines.\(^6\) West and colleagues\(^5\) did not include women with diabetes in their study and, although sleep apnoea is generally less common in women than in men, it is important that future research also addresses this issue in women.

The results of this study have important implications for those treating people with type 2 diabetes in both primary and secondary care. Clinicians should consider the possible diagnosis of OSA in patients with type 2 diabetes and, in patients who are symptomatic, determine whether treatment should be offered. This may not just improve sleep related symptoms but also has the potential to improve some features of the metabolic syndrome, notably blood pressure and possibly insulin resistance.\(^7\)\(^-\)\(^9\)

The mechanisms that underlie the association between type 2 diabetes and OSA are likely to be complex; both develop in the context of the “common soil” of obesity, but there is evidence to support the hypothesis that sleep apnoea may predispose to diabetes, and that diabetes increases the risk of sleep apnoea. There is a growing body of research that supports the concept that reduced sleep duration may itself increase the risk of diabetes, possibly via hormonal mechanisms related to stress such as catecholamines and cortisol.\(^10\) In addition, arousals in the context of sleep apnoea will activate the sympathetic nervous system and this could contribute both to insulin resistance and sympathetic mediated impairment of β cell function.\(^11\) Both insulin resistance and sleep apnoea have been associated with increased circulating concentrations of pro-inflammatory cytokines,\(^12\) some of which may derive from adipose tissue and may be secreted in greater amounts in response to tissue hypoxia.\(^13\) Adiponectin, a recently discovered adipose tissue derived hormone, has been shown to protect against the development of insulin resistance; one small study has shown lower adiponectin concentrations in patients with OSA than in those with simple snoring, implying that low levels of this hormone might contribute to the development of insulin resistance in OSA.\(^14\) Moreover, adiponectin secretion may also be reduced by hypoxia.\(^15\) Physical activity is strongly protective against the development of insulin resistance and type 2 diabetes, but the possibility that individuals with OSA are less physically active as a result of daytime sleepiness has not been explored. Diabetes can result in autonomic dysfunction, and this has been shown to result in abnormalities in respiratory function, particularly in the response to hypoxia and hypercapnia that may have implications for the development of symptomatic OSA which appears to be common even in non-obese diabetic patients with autonomic neuropathy.\(^16\)\(^-\)\(^17\)

There seems little doubt that patients with diabetes are more likely to have sleep apnoea than the general population. This has important implications for public health, especially given the prevalence rates of OSA reported by West et al of 23% in the diabetes population. In the UK alone, with an estimated 3 million people with type 2 diabetes, this represents nearly three quarters of a million individuals. Preliminary evidence suggests that treatment of OSA in people with type 2 diabetes improves glycaemic control, as well as reducing blood pressure and improving sleep related symptoms.\(^14\) In a largely non-diabetic population with severe OSA, long term use (over 10 years) of continuous positive airways pressure (CPAP) ventilation was associated with a more than 50% lower incidence of both fatal and non-fatal cardiovascular disease.\(^18\) Given that cardiovascular disease...
Obstructive sleep apnoea is independently associated with a higher prevalence of metabolic syndrome. Eur Heart J 2004;25:735-41.