Women who smoke
A review of the evidence

Background
Women who smoke are at significantly greater risk of developing a smoking related disease than men. They are also at risk of pregnancy related complications due to smoking and have more difficulty quitting. There are important gender differences in smoking behaviour that have implications for the quitting process. Advice to female smokers should take these factors into account and support should be tailored to their needs.

Objective
This article presents a summary of the evidence and a range of gender specific strategies that general practitioners can use to optimise the support they give to female smokers.

Discussion
Women may need more intensive behavioural and pharmacological support when quitting. Particular emphasis should be placed on addressing stress, social support, smoking cues and concerns about weight gain. Nicotine replacement therapy is less effective in women. However, it is still useful if used in adequate doses and can be used in pregnancy if the woman is unlikely to quit without it. Varenicline and bupropion are not approved in pregnancy or in lactation.

Keywords: smoking; women; health promotion

There is an increasing awareness of the special risks and needs of women who smoke. Women are at significantly greater risk of developing a smoking related disease than men, as well as being susceptible to gender specific health issues and pregnancy complications. Women also have more difficulty quitting smoking than men and may need extra support and advice to help them quit. In spite of this, women are less likely than men to be asked by their general practitioner about smoking or to be advised to quit. Even in pregnancy, GP intervention rates are well below recommended levels.

Smoking rates are lower in Australian women than men: 17.7% of women and 21.1% of men smoke cigarettes daily. In the indigenous population, 49% of women smoke daily compared to 51% of men. Women also smoke fewer cigarettes; 13 cigarettes per day on average compared to 14.5 per day for men.

However, women are less likely than men to quit successfully, whether quitting on their own, with counselling or by using nicotine replacement therapy (NRT). In one large study of 6600 smokers, women were 16% less likely to have quit than men at 13 years follow up.

Nicotine in women
Premenopausal women metabolise nicotine faster than men and postmenopausal women, and have significantly lower blood nicotine levels for the same cigarette intake. Women on the combined oral contraceptive pill (COCP) metabolise nicotine faster still, and the rate accelerates even further in pregnancy. Nicotine withdrawal symptoms are equally severe in men and women, and, in women, greater during the luteal phase.

Smoking in men is strongly reinforced by the psychoactive effects of nicotine. Women are more influenced by non-nicotine factors such as a desire to reduce negative mood, to facilitate social interaction and to prevent weight gain. Women also smoke more in response to cues or triggers, such as being with friends who smoke or when under stress.

Health effects
Women are at significantly greater overall risk from the health effects of smoking than men. A meta-analysis of 19 studies found that women had a 35% increased overall risk compared to men for a range of serious smoking related diseases such as stroke, acute myocardial infarction and cancers. In heavier smokers the additional risk for women was substantially greater. Female smokers also have a 60% increased risk of coronary heart disease compared to men who smoke, and in women taking COCP, smoking...
is associated with a 10-fold increased risk of coronary heart disease. This risk increases with age and cigarettes smoked per day and is regarded as a contraindication for women over 35 years of age who smoke 15 or more cigarettes per day. Gender specific health issues experienced by women are outlined in Table 1.

**Smoking in pregnancy and lactation**

Although most Australian women are aware of the substantial health risks to themselves and their fetus, 14.8% of nonindigenous women and 51.8% of indigenous women reported smoking during pregnancy. Only 7% of nonindigenous women and 3% of indigenous women are able to quit smoking in the first half of pregnancy. Nicotine and carbon monoxide cross the placenta freely and reach high concentrations in the fetus, reducing placental blood flow and creating fetal hypoxia and growth restriction. Smoking in pregnancy approximately doubles the risk of stillbirth, spontaneous miscarriage and preterm delivery. Smoking during pregnancy is associated with a 10-fold increased risk of coronary heart disease. This risk increases with age and cigarettes smoked per day and is regarded as a contraindication for women over 35 years of age who smoke 15 or more cigarettes per day. Gender specific health issues experienced by women are outlined in Table 1.

**Table 1. Gender specific health issues**

<table>
<thead>
<tr>
<th>Condition</th>
<th>Effect of smoking</th>
</tr>
</thead>
<tbody>
<tr>
<td>Cervical cancer</td>
<td>RR: 1.8343</td>
</tr>
<tr>
<td>Mucinous ovarian cancer (16% of total ovarian cancers)</td>
<td>RR: 2.1044</td>
</tr>
<tr>
<td>Breast cancer</td>
<td>44% increased risk in slow ‘NAT 2 gene acetylators’ (50–60% of Caucasians)</td>
</tr>
<tr>
<td>Endometrial cancer</td>
<td>Reduced by 25% in postmenopausal women who smoke</td>
</tr>
<tr>
<td>Infertility in women</td>
<td>OR: 1.6 2 month delay in conception</td>
</tr>
<tr>
<td>Assisted reproduction</td>
<td>OR: 0.5448</td>
</tr>
<tr>
<td>Dysmenorrhoea</td>
<td>OR: 1.3749</td>
</tr>
<tr>
<td>Menopause</td>
<td>1–4 years earlier</td>
</tr>
<tr>
<td>RR = relative risk; OR = odds ratio</td>
<td></td>
</tr>
</tbody>
</table>

Weight gain is a major concern for many women who smoke. Fifty percent of female smokers say that concerns about weight gain discourage them from trying to quit. On average, women gain more weight than men after quitting smoking, with 60% of the extra weight gained in the first year. Women are also more prone to extreme weight gain; 19.1% of women gain 20% or more of their baseline weight compared to 7.6% of men. Weight gain during or after smoking cessation increases the risk of relapse.

**Management**

**Optimising quit attempts**

Smoking cessation support in general practice should follow Australian guidelines, which are based on assessing the smoker’s readiness to change and the 5As: Ask, Assess, Advise, Assist and Arrange. However, guidelines need to be adapted to the special needs of women. In view of their lower success rates, women may need more intensive interventions to improve their chance of quitting successfully. In premenopausal women, quitting in the follicular phase may minimise withdrawal symptoms. Quit rates may be higher at this time with NRT, although a recent trial found better results from quitting in the luteal phase with bupropion.

**Managing weight gain**

Two recent reviews of weight gain prevention strategies for patients who were quitting smoking concluded that interventions are of borderline clinical benefit and are not justified on current evidence. There was mixed evidence for exercise in reducing weight gain. Giving general advice about reducing kilojoules was not shown to be effective in controlling weight or in achieving abstinence and is not recommended.

Nicotine replacement therapy, bupropion and probably varenicline reduce weight gain while being used, but this effect is not maintained at 12 months after quitting.

A pragmatic solution is to advise women to eat sensibly, exercise regularly and to accept some weight gain. Explain that one in four quitters will lose weight or stay the same and that the health benefits of quitting are almost always greater than the health effects of the extra weight.

**The role of medication in quitting**

The Smoking cessation guidelines for Australian general practice advise pharmacotherapy for all motivated smokers who are nicotine dependent. NRT is less effective in women than men. A meta-analysis found that the increase in quitting due to the nicotine patch versus placebo was only about half as large in women as in men at 6 months (4.6% vs. 9.3%).
Another meta-analysis of all nicotine replacement therapies (gum, patch, inhaler and spray) found they were equally effective in women and men in the short term but were significantly less effective in women at long term follow up. However, they did reduce withdrawal symptoms effectively in women. This gender difference in response to NRT may be partly due to genetic differences.

Nicotine replacement therapy is still worth considering for smoking cessation in women due to its safety, accessibility over-the-counter, ease of use and low cost. Nicotine patches are now available on a Pharmaceutical Benefits Scheme (Authority required), reducing the cost even further. Because of the faster metabolism of nicotine in premenopausal women, larger doses of NRT may be required, especially in women taking COCP. All commercially available forms of NRT increase chances of successful smoking cessation. There is some evidence that precessation nicotine patches and combination therapy (nicotine patch with gum, lozenge or inhaler) are the most effective forms of NRT.

Varenicline and bupropion appear to be equally effective in women and men, although absolute quit rates are higher with varenicline at 1 year (RR: 1.52).

Quitting during pregnancy and lactation

In pregnancy, smoking cessation interventions are modestly effective. Interventions reduce smoking by only 6% compared to control groups but do improve health outcomes. Quitting without any medication is preferred. However, NRT should be considered if the woman is unable to quit without it, although NRT has not yet been demonstrated to be effective in pregnancy in clinical trials. Nevertheless, it is likely to be beneficial given the strength of data in nonpregnant smokers.

Nicotine replacement therapy product information advises using intermittent dosing products such as gum or lozenges in pregnancy as they are likely to deliver a smaller daily dose of nicotine. However, because of the accelerated metabolism of nicotine in pregnancy, higher than standard doses are likely to be needed to control withdrawal symptoms and cravings. In practice, the minimum effective dose that controls

---

Table 2. Smoking cessation guidelines for Australian general practice (adapted for women*)

<table>
<thead>
<tr>
<th>Ask</th>
<th>Ask all patients and document in medical record (women are less likely to be asked*)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>• Do you smoke? If yes, how many, previous quit attempts?</td>
</tr>
<tr>
<td></td>
<td>• Have you ever smoked?</td>
</tr>
<tr>
<td>Assess</td>
<td>Readiness to change</td>
</tr>
<tr>
<td></td>
<td>• How do you feel about your smoking?</td>
</tr>
<tr>
<td></td>
<td>• Are you ready to quit now?</td>
</tr>
<tr>
<td></td>
<td>Nicotine dependence</td>
</tr>
<tr>
<td></td>
<td>• How soon after you wake do you have your first cigarette?</td>
</tr>
<tr>
<td></td>
<td>• Did you have cravings or withdrawal symptoms in previous quit attempts?</td>
</tr>
<tr>
<td></td>
<td>• How many cigarettes do you smoke per day?</td>
</tr>
<tr>
<td></td>
<td>(Smoking within 30 minutes of waking, withdrawal symptoms in previous quit attempts and smoking &gt;15 cigarettes per day are indicators of nicotine dependence)</td>
</tr>
<tr>
<td></td>
<td>Barriers to quitting</td>
</tr>
<tr>
<td></td>
<td>• Weight gain,* social pressure,* stress,* withdrawal symptoms, fear of failure</td>
</tr>
<tr>
<td>Advise</td>
<td>Advise all smokers to quit in a clear but nonconfrontational way (eg. the best thing you can do for your health is to quit smoking)</td>
</tr>
<tr>
<td>Assist</td>
<td>Not ready smokers</td>
</tr>
<tr>
<td></td>
<td>• Discuss health issues; offer written information; invite to return when ready</td>
</tr>
<tr>
<td></td>
<td>Unsure smokers</td>
</tr>
<tr>
<td></td>
<td>• Discuss health issues; explore barriers to quitting; offer written information</td>
</tr>
<tr>
<td></td>
<td>• Do brief motivational interviewing</td>
</tr>
<tr>
<td></td>
<td>Ready to quit (women may need more intensive intervention*)</td>
</tr>
<tr>
<td></td>
<td>• Set a quit date (avoid the luteal phase*) or gradually cut down if preferred</td>
</tr>
<tr>
<td></td>
<td>• Discuss weight gain concerns and give simple dietary and exercise advice*</td>
</tr>
<tr>
<td></td>
<td>• Assess for anxiety and depression and offer assistance*</td>
</tr>
<tr>
<td></td>
<td>• Develop coping strategies for smoking cues such as social interaction, stress and alcohol*</td>
</tr>
<tr>
<td></td>
<td>• Advise social support*</td>
</tr>
<tr>
<td></td>
<td>• Encourage partner to quit*</td>
</tr>
<tr>
<td></td>
<td>• Pharmacotherapy if nicotine dependent:</td>
</tr>
<tr>
<td></td>
<td>• nicotine replacement therapy. Optimise with adequate dose, precessation nicotine patch, combination therapy*</td>
</tr>
<tr>
<td></td>
<td>• varenicline or bupropion if not pregnant*</td>
</tr>
<tr>
<td></td>
<td>• Consider referral to Quitline 13 78 48</td>
</tr>
<tr>
<td>Arrange</td>
<td>Follow up within 7 days</td>
</tr>
<tr>
<td></td>
<td>• Give encouragement; review progress, problems and medication; give relapse prevention advice</td>
</tr>
<tr>
<td></td>
<td>• Negotiate further follow up visits</td>
</tr>
</tbody>
</table>

*Factors of special importance to women
symptoms should be used.\textsuperscript{21} Expert consensus is that NRT is safer than smoking in pregnancy.\textsuperscript{41} Of women who do quit in pregnancy up to 80\% will relapse within 1 year of birth. If a pregnant woman is unable to quit, cigarette reduction has been correlated with increased birth weight. A reduction to less than eight cigarettes per day may minimise harm.\textsuperscript{42} 

Product information also advises intermittent dosing NRT products during lactation to minimise risk to the infant. However, nicotine concentrations in the infant from NRT are low and unlikely to be harmful. NRT is considerably less hazardous to both mother and child than exposure to cigarette smoke.\textsuperscript{21} 

Bupropion and varenicline are not approved in pregnancy or lactation.

**Summary of important points**

- Women have more difficulty quitting and need more intensive support.
- Particular emphasis should be placed on addressing stress and mood disorders, social support and smoking cues in women.
- Weight gain prevention strategies are not supported by evidence.
- NRT is less effective in women but still has a useful role. Precessation nicotine patch and combination therapies are most effective.
- NRT should be considered in pregnancy if the woman is unable to quit without it.
- Varenicline and bupropion are equally effective in women and men but are not approved in pregnancy.

**Author**

Colin Mendelsohn MBBS(Hons), is a general practitioner, Sydney, New South Wales. mendel@bigpond.net.au.

Conflict of interest: Dr Mendelsohn has received honoraria for teaching, consulting and travel from Pfizer and GlaxoSmithKline. He is on Pfizer's Champix Advisory Board and has served on GlaxoSmithKline's Nicotine Replacement Therapy Expert Panel. Both companies have sponsored articles in Your Health newsletter, of which Dr Mendelsohn is editor.

**Acknowledgment**

Thanks to Professor Nick Zwar and Associate Professor Renee Bittoun for reviewing the manuscript.

**References**

8. Benowitz NL, Sessos-Schlagger CN, Swan GE, Jacob 3rd P. Female sex and oral contracepti
34. Schnoll RA. Sex heterogeneity in pharmacogenetic smoking cessation clinical trials. Drug and Alcohol Dependence 2009;S94–99.


