

## South African tobacco smoking cessation clinical practice guideline

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*This guideline has been compiled on behalf of the South African Thoracic Society to provide practising clinicians with a resource from which to base individual cessation interventions for patients within the South African environment.*

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Tobacco smoking (i.e. cigarettes, rolled tobacco, pipes, etc.) is associated with significant health risks, reduced life expectancy and negative personal and societal economic impact. Smokers have an increased risk of cancer (i.e. lung, throat, bladder), chronic obstructive pulmonary disease (COPD), tuberculosis and cardiovascular disease (i.e. stroke, heart attack). Smoking affects unborn babies, children and others exposed to second hand smoke. Stopping or 'quitting' is not easy. Nicotine is highly addictive and smoking is frequently associated with social activities (e.g. drinking, eating) or psychological factors (e.g. work pressure, concerns about body weight, anxiety or depressed mood). The benefits of quitting, however, are almost immediate, with a rapid lowering of blood pressure and heart rate, improved taste and smell, and a longer-term reduction in risk of cancer, heart attack and COPD. Successful quitting requires attention to both the factors surrounding why an individual smokes (e.g. stress, depression, habit, etc.) and the symptoms associated with nicotine withdrawal. Many smokers are not ready or willing to quit and require frequent motivational input outlining the benefits that would accrue. In addition to an evaluation of nicotine dependence, co-existent medical or psychiatric conditions and barriers to quitting should be identified. A tailored approach encompassing psychological and social support, in addition to appropriate medication to reduce nicotine withdrawal, is likely to provide the best chance of success. Relapse is not uncommon and reasons for failure should be addressed in a positive manner and further attempts initiated when the individual is ready.

Key steps in smoking cessation include: (i) identifying all smokers, alerting them to the harms of smoking and benefits of quitting; (ii) assessing readiness to initiate an attempt to quit; (iii) assessing the physical and psychological dependence to nicotine and smoking; (iv) determining the best combination of counselling/support and pharmacological therapy; (v) setting a quit date and provide suitable resources and support; (vi) frequent follow-up as often as possible via text/telephone or in person; (vii) monitoring for side-effects, relapse and on-going cessation; and (viii) if relapse occurs, providing the necessary support and encourage a further attempt when appropriate.

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### 1. Tobacco smoking in South Africa

There are an estimated 1.3 billion smokers worldwide and over 5 million deaths per year attributable to tobacco smoking.<sup>[1]</sup> Even though smoking rates are declining, there are an estimated 7 million smokers in South Africa (SA).<sup>[2]</sup> Tobacco smoking is undoubtedly the primary risk factor for chronic obstructive pulmonary disease (COPD), which is estimated to be the third highest cause of death globally by 2030. SA has a particularly high prevalence of smoking (20%).<sup>[3,4]</sup> SA also has one of the highest burdens of tuberculosis (TB) and HIV, which are both risk

factors for COPD and exacerbate the effects of smoking.<sup>[5,6]</sup> Tobacco smoking increases the risk for TB, cancer, pneumonia, ischaemic heart disease and stroke, which are all leading causes of death globally.<sup>[1]</sup> The mortality among current smokers in SA is nearly double that of non- or ex-smokers.<sup>[7,8]</sup> Up to a third of all male deaths in SA adults over the age of 35 years have recently been attributed to tobacco use.<sup>[7,9]</sup> Passive smoking also increases the risk of cardiovascular disease in adults and respiratory disease, particularly among children. The cost of smoking-related disease to the SA economy is estimated to be R1.2 billion.<sup>[10]</sup>

The benefits of stopping smoking are almost immediate, with a lowering of blood pressure within minutes, and longer-term benefits

such as improved lung function and reduced risk for lung cancer, stroke and heart disease. Smoking cessation is a critical component of the effective management of COPD.<sup>[11]</sup> Savings of disposable income, achieved by quitting smoking, would also be available for basic necessities, particularly for those living in poverty.

Many smokers have no desire to quit and will require repeated engagement by health practitioners to affect behaviour change. Interviewing techniques have changed from the more traditional approach, typified by scolding or lecturing, to one that involves support, encouragement and the provision of information.<sup>[12]</sup> For those who wish to stop there are many options available such as 'cold turkey', cognitive behavioural therapy, acupuncture, hypnosis, internet and cellular phone-based support programmes, and medication.

This guideline assesses published evidence and reviews international guidelines, applying them to the specific needs and circumstances in SA in developing a clinical practice guideline for SA clinicians.

Published international guidelines from the US Centers for Disease Control, the American College of Chest Physicians (ACCP), the UK National Institute for Clinical Excellence (NICE) and the Cochrane Collaboration *Database of Systematic Reviews* for all topics relating to tobacco smoking were reviewed. In addition, PubMed was searched for newer studies on specific topics such as e-cigarettes, smoking in pregnancy, and smoking in persons infected with HIV or TB. Smoking in this guideline refers to all forms of smoking tobacco products such as cigarettes, cigars and rolled tobacco. Smoking cessation – also referred to as quitting, is the process of stopping smoking. Where available, evidence grading is provided with the grading source.

## 2. Tobacco smoking cessation strategies

Smoking cessation can be broadly divided into two phases: (i) the identification of smokers and assisting them to quit; and (ii) initiating and sustaining the quit attempt (Fig. 1). The strength of evidence for each intervention varies. For each stage and intervention, key points and strength of supporting data are presented and expert opinion was relied on where data was lacking.

### 2.1 Identifying smokers and initiating quit attempts

- The strategy of identifying smokers during routine consultations with healthcare practitioners increases quit attempts (Grade A<sup>[13]</sup>).
- Brief smoking cessation interventions should be implemented, regardless of the availability of, and access to, specialised services (Grade A<sup>[13]</sup>).

'If you don't think about it, you won't do it'. Using stickers, tags or reminders in folders for the nurses or doctors increases the likelihood of discussions with patients around smoking, counselling and referral for cessation advice. The inclusion of smoking as one of the vital signs identifies 80% or more of smokers attending primary care facilities.<sup>[14]</sup>

### 2.2 Clinical interventions to initiate and sustain smoking cessation

- Repeated encouragement and assistance provided by more than one physician/healthcare worker, through a range of supportive options, increase the likelihood of abstinence (Grade A<sup>[13]</sup>).

Interventions as brief as 3 min can significantly increase quit rates, but intensive interventions are more effective (more comprehensive treatments being those that occur over multiple visits, for a longer period of time and/or are provided by more than one clinician).<sup>[14]</sup> The goal should be for every tobacco user to be *identified* and *offered at least a brief intervention at every clinical consultation*. A commonly

recommended approach is the **5As model**, which is an effective, structured approach to providing brief interventions (3 - 10 min) in the primary care setting. The 5 steps are: (i) asking about smoking; (ii) alerting the patient to the benefits of quitting; (iii) assessing readiness to make a quit attempt; (iv) assisting those willing to quit; and (v) arranging for follow-up contact and referral to further resources, such as quitlines. Problem solving, skills training and psychosocial support during treatment are all effective elements of counselling. They can be used as part of brief interventions, but also form the foundation of more intensive interventions.<sup>[14]</sup>

### 2.3 Motivational interviewing approach

Many patients lack the motivation to quit or express a lack of readiness to quit. This may be because they lack information, have concerns about quitting, face significant social/environmental barriers or lack confidence in their ability to quit. Such patients may benefit from counselling methods derived from motivational interviewing (MI).<sup>[12]</sup>

In this approach, the clinician encourages the smoker to explore their feelings of ambivalence about smoking/quitting by discussing, for example, the pros and cons of smoking from their perspective, and by exploring how personally important cessation may be and how confident they feel about it. In this process, the provider supports and strengthens any intentions to change, which come from the patients themselves. The idea is to elicit *self-motivational* statements, rather than tell the patient what they should do and how they should do it. MI posits that a good collaborative relationship, in which a client is viewed as the expert on his/her own life, serves to minimise resistance to change and thereby enhances motivation.<sup>[15]</sup> This varies significantly from the traditional, advice-giving approach, which casts the provider in a dominant, directing role and the patient merely as a passive recipient of their expert knowledge. Meta-analyses have shown that MI can significantly improve smoking cessation rates over brief advice.<sup>[16]</sup>

### 2.4 The individual who is not ready to quit

- Encourage the individual to think about quitting, identify the reasons for smoking and the barriers to quitting and provide an opportunity for follow-up.

For individuals who are not ready or willing to quit, on-going encouragement and offers of support are necessary. The practitioner can provide information about the risks of smoking and the benefits of quitting and encourage the patient to return for further discussions if and when they *are* ready to quit. Assisting the patient to identify the perceived benefits that they gain from smoking (i.e. stress reduction, weight control), their perceived barriers to quitting (i.e. social pressure, withdrawal symptoms) and the benefits of quitting (i.e. health improvement, cost savings) can lay the foundation to behavioural change at a later date and enhance the possibility of a quit attempt.

### 2.5 Assisting the motivated patient to quit

- Counselling plus medication to treat nicotine withdrawal is more effective than either intervention alone (Grade A<sup>[13]</sup>).
- Multiple options are available and can be tailored to suit the individual patient and circumstance.
- Smokers who are motivated, have supportive networks and are ready to change, have higher success rates.
- Smokers who are highly nicotine dependent, have social stressors and psychiatric comorbidities, are less likely to be successful.

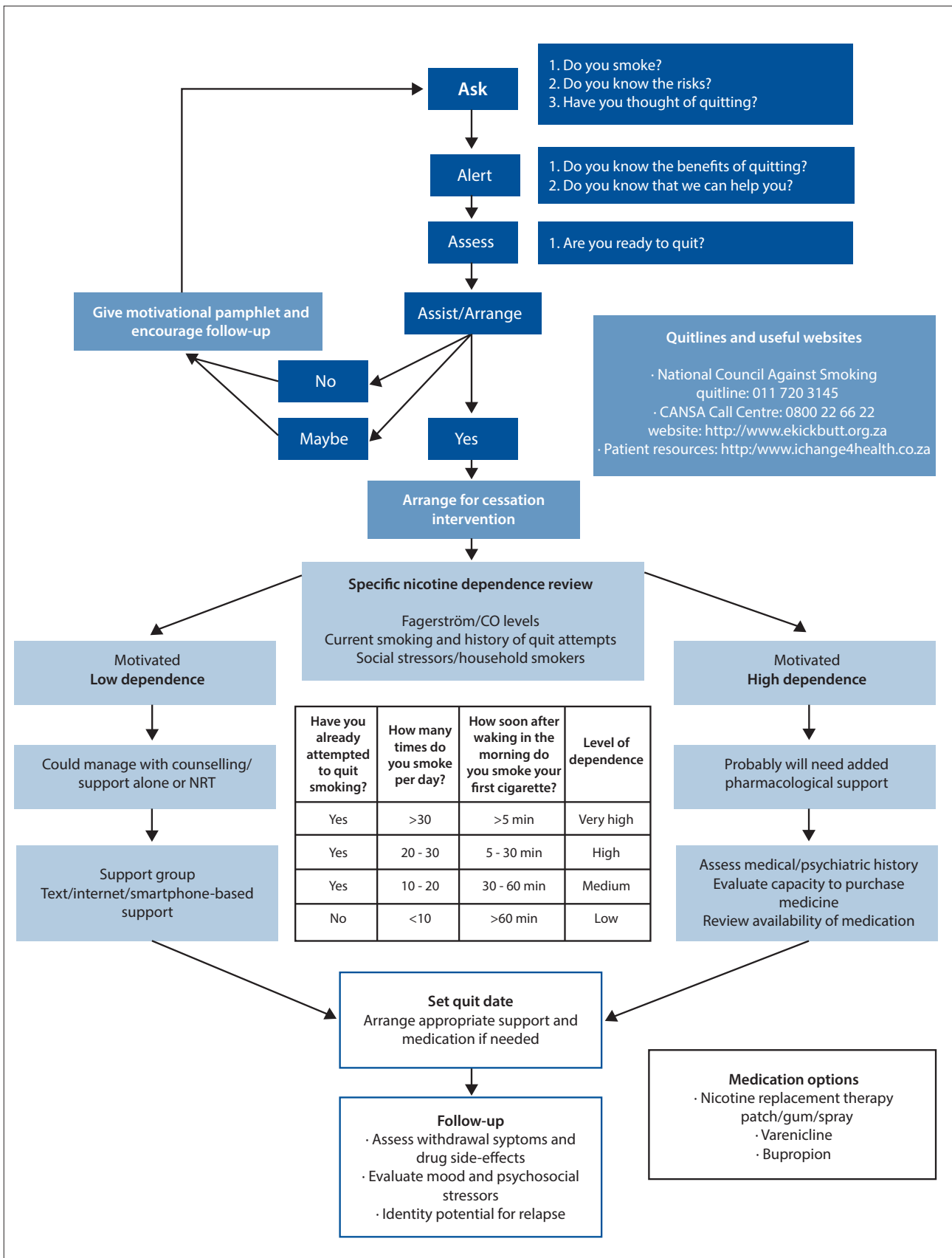


Fig. 1. Smoking cessation approach in clinical practice. All smokers should be identified and their readiness to quit evaluated. If resistance to quitting exists, information and suitable follow-up should be provided (top). If an individual is ready to make a quit attempt, evaluation of nicotine dependence, co-morbid disease and psychosocial factors should guide the clinician in the choice of effective interventions (bottom). CO = carbon monoxide; NRT = nicotine replacement therapy.

### 3. Pharmacological intervention (drug treatment to aid smoking cessation)

Several pharmacological strategies are available to assist in smoking cessation – predominantly to address acute nicotine withdrawal. These may be simply classified as nicotine replacement therapies (NRTs) or drugs that reduce addiction. Although both classes of drug are an aid to smoking cessation, they have little or no effect on the underlying addiction and do not address the psychosocial factors that cause a person to smoke. Most studies of pharmacological interventions have been conducted in subjects who smoke >10 cigarettes/day and frequently >20 cigarettes/day. There is little evidence to guide the use of pharmacological therapy in subjects who smoke only 3 - 5 cigarettes/day or in those with very low nicotine dependence. Table 1 outlines the comparative performance of the various drugs available to assist in smoking cessation.

#### 3.1 NRT

- NRT is effective and should be encouraged (Grade A<sup>[13]</sup>).
- A controller (patch) and reliever (gum, spray) approach is the most effective way to use NRT (Grade A<sup>[13]</sup>).

NRT has been the mainstay for addressing nicotine withdrawal. It is particularly effective when employing dual NRT – one as a ‘controller’ and one as a ‘reliever’ (odds ratio (OR) 3.6 (95% confidence interval (CI) 2.5 - 5.2) for success, and abstinence rates 36.5% (95% CI 28.6 - 45.3) at the end of therapy).<sup>[14]</sup> This approach, which is advocated by the ACCP, is more effective than using a single form of NRT (relative risk (RR) 1.34; 95% CI 1.18 - 1.51).<sup>[17]</sup> Titration of the NRT (via patch strength) to the level of symptom severity based on the Fagerström scale (Table 2), or of cotinine level is recommended to ensure that adequate control of withdrawal symptoms is achieved. If ‘breakthrough’ symptoms occur the ‘reliever’ NRT should be used (gum, spray, etc.) The use of nicotine spray may be preferred to gum in some individuals.<sup>[18]</sup> Combinations of NRT with bupropion are more effective than NRT alone (OR 2.57; 95% CI 1.05 - 6.32),<sup>[19]</sup> but based on the theory of their mechanism of action, there should be no value in using both NRT and varenicline together. Despite this, there are anecdotal reports of additional benefit, and this has recently been tested in the VARNIC study, the results of which are expected during 2014. Note: at the time of writing these practice guidelines, nicotine patches were not available in SA.

#### 3.2 Antidepressants

- Bupropion is an effective drug for smoking cessation particularly in combination with NRT (Grade A<sup>[13]</sup>).
- Nortriptyline is a moderately effective drug for smoking cessation (Grade A<sup>[13]</sup>).
- The use of bupropion or nortriptyline must be under medical supervision due to potential side-effects and interactions.

Bupropion (Zyban) used at doses of 150 mg twice daily for 7 - 12 weeks is effective (RR 1.69; 95% CI 1.53 - 1.85).<sup>[20]</sup> Additionally, bupropion appears to reduce long-term relapse and the weight gain associated with quitting.<sup>[21]</sup> Its most frequent side-effects are insomnia, dry mouth and nausea.<sup>[20]</sup> Bupropion may also reduce the seizure threshold and a seizure risk of 1/1 000 has been reported.<sup>[22]</sup> There is no evidence that selective serotonin re-uptake inhibitors (SSRIs) are effective for smoking cessation.<sup>[20]</sup>

Nortriptyline (not registered in SA) is generally considered a second-line therapy for those who have failed NRT and bupropion/varenicline.<sup>[23]</sup> Nortriptyline is a metabolite of amitriptyline, which is available in SA. There are, however, no published data on smoking cessation using amitriptyline. Given the increased associated risk of suicide in smokers,<sup>[24,25]</sup> the serious complications of amitriptyline overdose and its unknown efficacy as a smoking cessation aid, the drug cannot be recommended for smoking cessation until further data are available.

#### 3.3 Nicotine receptors agonists

- Varenicline is an effective smoking cessation therapy (Grade A<sup>[13]</sup>).
- Although not confirmed in large studies and meta-analyses, there is concern about incidents of suicide or suicidal behaviour.
- It is strongly advised that all patients who are prescribed these drugs are closely monitored for behavioural change and/or neuropsychiatric symptoms.

Two nicotine receptor partial agonists have been marketed: varenicline (Champix – registered in SA) and cytisine (Tabex – not registered in SA). By acting as partial agonists, they stimulate dopamine release and reduce nicotine withdrawal symptoms.<sup>[26-28]</sup> In controlled trials and meta-analyses, varenicline has been shown to be the most effective single drug for smoking cessation (RR 3.1; 95% CI 2.5 - 3.8)<sup>[29,30]</sup> whereas cytisine, although effective, appears to result in only a modest increase in quit rates.<sup>[29,31]</sup> The most common side-effects reported with varenicline include abnormal dreams,

**Table 1. Comparisons of abstinence rates achieved with different pharmacological interventions to support smoking cessation\***

Medication	Arm, n	OR (95%CI)	Estimated abstinence rate (95%CI)
Placebo	80	1	13.8
NRT			
Nicotine gum	15	1.5 (1.2 - 1.7)	19 (16.5 - 21.9)
Nicotine patch	32	1.9 (1.7 - 2.2)	23.4 (21.3 - 25.8)
Nicotine spray	4	2.3 (1.7 - 3)	26.7 (21.5 - 32.7)
Antidepressants/centrally acting agents			
Bupropion SR	26	2 (1.8 - 2.2)	24.2 (22.2 - 26.4)
Nortriptyline	5	1.8 (1.3 - 2.6)	22.5 (16.8 - 29.4)
Clonidine	3	2.1 (1.2 - 3.7)	25 (15.7 - 37.3)
Nicotine receptor agonist			
Varenicline (2 mg/day)	5	3.1 (2.5 - 3.8)	33.2 (28.9 - 37.8)

OR = odds ratio; CI = confidence interval; NRT = nicotine replacement therapy; SR = slow release.  
\*Data from Fiore *et al.*,<sup>[13]</sup> comparisons are with placebo treatment.

**Table 2. Fagerström test for nicotine dependence<sup>[85]</sup>**

Question	Score
1. How soon after you wake up do you smoke your first cigarette?	
Within 5 min	3
6 - 30 min	2
31 - 60 min	1
After 60 min	0
2. Do you find it difficult to refrain from smoking in places where it is forbidden?	
Yes	1
No	0
3. Which cigarette would you hate most to give up?	
The first in the morning	1
Any other	0
4. How many cigarettes per day do you smoke?	
≤10	0
11 - 20	1
21 - 30	2
≥31	3
5. Do you smoke more frequently during the first hours after waking than during the rest of the day?	
Yes	1
No	0
6. Do you smoke even if you are so ill that you are in bed most of the day?	
Yes	1
No	0
TOTAL SCORE _____	
<b>Interpretation of total score<sup>[86]</sup></b>	
0 - 3	Mild dependence
4 - 6	Moderate dependence
7 - 10	Severe dependence

nausea and headache.<sup>[32,33]</sup> While smokers as a group are at higher risk of suicide than non-smokers,<sup>[24,25]</sup> there are concerns about the neuropsychiatric side-effects of varenicline. While the US Food and Drug Administration (FDA) mandated a 'black box' warning in 2008, meta-analyses have not confirmed this risk.<sup>[29,34]</sup> Recent studies have also reported that varenicline appears to be safe in patients with schizophrenia<sup>[35]</sup> and in patients with cardiac disease.<sup>[29,36-38]</sup> Nevertheless, it should be used with caution and under supervision. Champix is a schedule 5 drug in SA and patients should be monitored regularly with particular attention to changes in their emotional state, unusual behaviour and suicidal ideation.

### 3.4 Electronic cigarettes

- There is no evidence that electronic cigarettes (e-cigarettes) are effective aids

to smoking cessation, although they may reduce the number of cigarettes smoked.

E-cigarettes are battery-powered devices, similar in appearance to conventional cigarettes that vaporise nicotine. A large variety of products are on offer. They are available in different flavours and may be used in an attempt to reduce the symptoms of nicotine withdrawal while allowing the smoker to participate in the 'ritual' of smoking. E-cigarettes are available over the counter and there is currently little legislative control on their use, availability and marketing. Recently however, both the FDA and the South African Medicines Control Council have begun to consider regulation of these products.<sup>[39]</sup>

The role of e-cigarettes in smoking cessation algorithms remains unclear. As they are likely to be less hazardous than tobacco smoking,<sup>[40]</sup> they may, in future, have a role in smoking

cessation, either to reduce nicotine intake or as a bridge to smoking cessation. In a small pilot study, they appeared to decrease cigarette consumption,<sup>[41]</sup> and they may be especially useful in reducing consumption in chronic psychiatric (schizophrenic) patients.<sup>[42]</sup> Side-effects include mouth and throat irritation, dry cough, nausea and headache, although these appear to decrease over time. Concerns have been raised about the long-term safety of e-cigarettes, particularly with regard to the flavouring used.<sup>[43]</sup> Additionally, although e-cigarettes do not produce classic smoke, they have been found to produce short-term adverse physiological effects on the airways.<sup>[44]</sup> Currently, given the lack of data on efficacy and limited long-term safety data, they are not recommended as part of smoking-cessation strategies.

### 3.5 Nicotine vaccine

- The theory behind nicotine vaccines is that they induce antibodies that bind to nicotine, reducing its availability to central receptors.
- Nicotine vaccines are still in development and their efficacy has not been confirmed.<sup>[45]</sup>

### 3.6 Complementary medicine

Several forms of complementary medical approaches are in common use and are widely advertised as aids to smoking cessation. These include hypnotherapy, acupuncture, acupressure and electro-stimulation. However, none of these methods are supported by convincing efficacy data when subjected to review using Cochrane Library methods. Although positive results have been reported in individual studies, there is a lack of data from large, randomised, controlled studies. When performed by experienced and qualified individuals, they are likely to be safe and may benefit some. However, they are not recommended as effective strategies.

#### 3.6.1 Hypnotherapy

- There is no evidence to support hypnotherapy as an effective aid to smoking cessation.

Hypnotherapy is a widely promoted aid to smoking cessation and multiple approaches, techniques and success rates have been reported. Two recently published meta-analyses, in which different study selection criteria were employed, concluded that hypnotherapy was not effective, as insufficient quality data existed comparing hypnotherapy with other methods for smoking cessation. Estimates in the two separate meta-analyses for success of hypnotherapy in achieving smoking cessation were: OR 4.55 (95% CI 0.98 - 21.01)<sup>[46]</sup> and RR 1.49 (95% CI 0.86 - 2.5)<sup>[47]</sup>



### 3.6.2 Acupuncture

- 'There is no consistent, evidence that either acupuncture or acupressure are effective in smoking cessation.'<sup>[48]</sup>

Several studies and meta-analyses have evaluated the effects of acupuncture on smoking cessation in a controlled, randomised manner.<sup>[48-50]</sup> Differing methods used in these studies (e.g. the nature of control procedures and the selection of control groups) do not provide sufficient consistency for a bias-free analysis. The most recent and comprehensive Cochrane review concluded that acupuncture is less effective than NRT (RR 1.05; 95% CI 0.82 - 1.35).<sup>[48]</sup>

## 4. Smoking cessation in specific situations

### 4.1 Pregnancy

- Smoking poses significant risks to mother and fetus.
- Brief counselling by a healthcare provider as part of routine antenatal care is effective and well received by pregnant women.
- There is, as yet, insufficient, high-quality evidence to determine whether the use of pharmacotherapy (i.e. NRT, bupropion or varenicline) is effective and safe during pregnancy.

The risks to mother and child from smoking during pregnancy, and in the post-natal period are well established. Maternal smoking is associated with obstetric risks (miscarriage and premature rupture of membranes, placental abruption, intra-uterine growth retardation and stillbirth).<sup>[51]</sup> Maternal exposure to second-hand smoke in pregnancy can also increase the risk of low-birthweight children.<sup>[52]</sup> Although quitting early in pregnancy will produce the greatest benefits, stopping at any stage during pregnancy yields benefits to the fetus and mother, and the child in the postnatal period.<sup>[53]</sup> Passive smoking increases the risk for asthma, middle ear and recurrent chest infections in children.<sup>[54]</sup> Counselling should be offered at the first antenatal visit and if possible, smokers in the household or workplace should be identified. A smoking cessation intervention involving brief counselling by lay counsellors, supported by midwives and by educational materials specifically tailored to pregnancy, was shown to be effective in increasing cotinine-validated quit rates among disadvantaged women attending public sector antenatal care clinics in SA.<sup>[55]</sup> On the basis of the substantial evidence of benefit from NRT in the general population and the limited evidence available among pregnant women,<sup>[53,56,57]</sup> some guidelines recommend the use of NRT in pregnancy under medical supervision, but only when behavioural therapy has failed and in heavily dependent smokers who remain motivated to quit.<sup>[58]</sup>

### 4.2 Adolescents/paediatrics

- Treatment strategies relying on behavioural change are effective.
- No smoking cessation medication is licensed for use in children aged less than 18 years.
- Paediatricians and healthcare workers providing care to children must address smoking by parents and advise on smoking-cessation strategies.

Tobacco smoking in adolescents frequently leads to long-term nicotine addiction and the consequent adverse health-effects. In SA in 2008, 21% of learners (grade 8 - 10) were found to be current smokers, with 6.8% having initiated smoking before the age of 10 years.<sup>[59]</sup> The smoking of *hookah* pipes ('hubbly-bubbly') should be addressed among adolescents who frequently believe it to be tobacco-free and, hence, safe. Preventing adolescents from starting to smoke is vital to reduce the numbers of adults who smoke.

Nearly half of adolescent smokers in SA attempt to quit each year but factors such as stress, depression, peer pressure and weight gain impact on their success.<sup>[59-62]</sup> Behavioural change can be encouraged by focussing on the health benefits and improved school and sport performance, and by recognising the effects of cultural differences and the social pressures that exist with regard to smoking.<sup>[60-62]</sup> A variety of methods (ranging from one-on-one sessions to group therapy, telephonic helplines and web-based programmes) have been tried and, if tailored to individual needs, may double success rates. There are insufficient studies comparing these models in order to recommend one particular strategy in adolescents.<sup>[13,63]</sup> It is imperative to speak to the adolescent alone (without the caregiver present) and to maintain confidentiality when encouraging behavioural change.

SA data show that adolescents who have been exposed to smokers are more likely to smoke than those who have not (74.5% v. 44%, respectively).<sup>[59]</sup> Counselling of parents on the harmful effects of smoking and on interventions to aid quitting may reduce secondary smoke exposure in children.<sup>[13]</sup> Parental smoking cessation is also associated with higher quit rates among adolescents.<sup>[62,64]</sup>

No smoking-cessation medications are FDA-approved in children or adolescents (under 18 years). The UK NICE guidelines, however, support the use of NRT if required, along with behavioural interventions in teenagers over 12 years of age.<sup>[58]</sup> If a decision is made to use NRT in a teenager, it should be used in conjunction with behavioural therapy and should be individualised. Although safety studies have been conducted with bupropion and varenicline in adolescents, both are only approved for adults in SA.<sup>[33]</sup>

### 4.3 Tuberculosis

- Smokers have approximately double the risk of developing TB and of dying from TB than non-smokers.
- Smoking cessation for people with active TB is a feasible and effective intervention.

There is substantial epidemiological evidence that smokers have a higher risk of developing TB (both latent and active) and dying from TB.<sup>[65-67]</sup> In SA, with the added risk of HIV, the importance of smoking cessation to reduce the risk of TB, HIV-related diseases and COPD is unquestionable.<sup>[66,68]</sup> There are some data to show that integrating smoking cessation efforts (support plus NRT) with TB treatment is feasible,<sup>[69]</sup> improves quit rates (77% v. 8.7%, respectively) and completion of TB treatment (97.5% v. 84.8%, respectively).<sup>[70]</sup> It is unclear, however, whether stopping smoking during TB therapy will reduce the excess mortality associated with smoking and TB,<sup>[71]</sup> nor are there any efficacy data available on whether using bupropion or varenicline with concomitant anti-TB therapy may be of benefit. Smoking cessation should, however, be encouraged in all TB patients and appropriate support provided.

### 4.4 HIV/antiretroviral therapy

- Smoking cessation is an important and effective intervention in individuals living with HIV.
- Drug-drug interactions may occur with bupropion.

The risks of smoking in HIV-infected individuals are well described and include *inter alia* pneumonia, TB and lung cancer.<sup>[72]</sup> There are several small trials on smoking cessation in the context of HIV that have concluded that it is an important and effective intervention.<sup>[72-74]</sup> Smoking and nicotine are known to induce hepatic enzymes,<sup>[75,76]</sup> but not those metabolising common antiretrovirals (ARVs). Enzyme inhibition/induction by ARVs such as lopinavir-ritonavir may

increase or reduce serum concentrations of bupropion and should be used with caution.<sup>[74]</sup> Varenicline is not metabolised by the liver, thus drug-drug interactions should not occur, but common side-effects such as nausea may interfere with ARV compliance.<sup>[73]</sup>

#### 4.5 Mental illness

- Smoking is common in individuals with mental illness.
- Adequate management of the underlying psychiatric illness is key to successful smoking cessation.

Anxiety disorders, depression, and schizophrenia are strongly associated with tobacco smoking.<sup>[35,77,78]</sup> There is evidence to suggest that psychiatric disorders may lead to self-medication with nicotine and that smoking may predispose individuals to mental illness.<sup>[78,79]</sup> Smoking cessation is important even in those with complex psychiatric disorders. Where necessary, consultation with an experienced psychiatrist may be beneficial in choosing the best options for management of the mental illness and drugs for smoking cessation if required. A recent Cochrane review concluded that bupropion and varenicline appear to be safe and effective in schizophrenia, but NRT and psychological interventions show no benefit.<sup>[35]</sup> In patients with current depression, management of the depression should be addressed first, as there is limited success of smoking cessation interventions (even bupropion).<sup>[77]</sup>

#### 4.6 In-hospital cessation

- In-hospital cessation counselling with post-discharge follow-up is an effective intervention to assist in smoking cessation.
- The addition of NRT substantially improves quit rates, but insufficient data exist as to the additional benefits of bupropion or varenicline.

As smoking is banned in most public places and particularly in hospitals, acute admissions in which smoking cessation is imposed are convenient opportunities to promote smoking cessation. Many trials have demonstrated that appropriate in-hospital counselling and post-discharge follow-up are effective.<sup>[80]</sup> The additional use of NRT appears to be highly effective (54% increase in cessation rates).<sup>[80]</sup> There are insufficient data concerning the use of bupropion or varenicline during hospitalisation. Acute pre-operative cessation has mixed reports of long-term success, although lower complication rates (in relation to wound healing and lung function) occur in those who quit at least 6 - 8 weeks prior to surgery.<sup>[81-84]</sup>

### 5. Relapse

Relapse is common; frequently, because the underlying psychosocial factors or nicotine addiction have not been adequately addressed. Relapses can be viewed as an opportunity to learn how to approach a subsequent quit attempt, rather than as outright failure. Many attempts may be required before long-term success is achieved. If relapse does occur, appropriate support should be provided and the reasons for relapse reviewed before another quit attempt.

### 6. Implementation and further research

Smoking cessation at the primary level will require adequate training of staff to provide the necessary counselling to motivate patients to attempt to quit. Lay counsellors providing HIV adherence support could be trained to offer smoking cessation counselling and support. The prescription of medication to support quit attempts may need to be restricted to clinicians/clinics who are able to fully assess nicotine dependence and monitor side-effects of the medication. Research into the optimal use of medications in HIV/TB patients as well as the possible use of amitriptyline is needed.

### 7. Summary and recommendations

Nicotine addiction and psychosocial stressors make smoking cessation difficult, even in those who are motivated to quit. Motivating those with no apparent interest in quitting nor confidence to quit, requires support and encouragement from healthcare practitioners at every contact and should involve the entire health team. Although time is limited in clinical practice, brief motivational counselling with appropriate referral of the individual who is motivated to quit, is effective.

These practice guidelines serve as an aid to the clinician in deciding on the best strategies to use for smoking cessation. Even the best programmes internationally have only modest success rates, with frequent relapses if the underlying reasons predisposing to smoking or the barriers to cessation have not been addressed. Informing the smoker about the immediate and long-term benefits of quitting, anticipating the difficulties that could be expected and problem solving with the patient, as well as prescribing appropriate medication where needed, all increase the possibility of success. If such interventions are widely applied to a large proportion of smokers, they have the potential to achieve important reductions in disease and associated excess healthcare costs.

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