Psychiatrists often fail to adequately address their patients’ smoking, and often underestimate the impact of ongoing tobacco use. Evidence suggests that heavy smoking is a risk factor for major depressive disorder; it also is associated with increased suicidal ideations and attempts.\textsuperscript{1,2} Tobacco use also has a mood-altering impact that can change the trajectory of mental illness, and alters the metabolism of most psychotropics.

Previously, psychiatrists may have been reluctant to prescribe the most effective interventions for smoking cessation—varenicline and bupropion—because these medications carried an FDA “black-box” warning of neuropsychiatric adverse effects, including increased aggression and suicidality. However, a large study called the EAGLES trial\textsuperscript{3} found that the neuropsychiatric risks associated with these medications were lower than previously thought. Consequently, in December 2016, the FDA removed the black-box warning related to serious mental health adverse effects from the labeling of varenicline and bupropion.\textsuperscript{4}

The EAGLES trial was a large, multi-site global trial that included patients with and without mental illness. Its primary objective was to assess the risk of “clinically significant” adverse effects for individuals receiving varenicline, bupropion, nicotine replacement therapy (NRT), or placebo, and whether having a history of psychiatric conditions increased the risk of developing adverse effects when taking these therapies. Overall, 2% of smokers without mental illness experienced adverse effects, compared with 5% to 7% in the psychiatric cohort, regardless of treatment arm. The rate of neuropsychiatric events and scores on suicide severity scales were similar across treatment arms in both cohorts.\textsuperscript{3}

We should take lessons from the EAGLES trial. We propose that clinicians ask themselves the following 6 questions when forming a treatment plan to address their patients’ tobacco use:

1. Does the patient meet DSM-5 criteria for nicotine use disorder and, if yes, what is the severity of his or her nicotine dependence? The Fagerstrom Test for Nicotine Dependence (FTND)\textsuperscript{5} is a 6-question instrument for evaluating the quantity of cigarette consumption, compulsion to use, and dependence. It provides clinicians with guidelines on preventing withdrawal by implementing NRTs, such as lozenges, an inhaler, patches, and/or gum. A score of 1 to 2 (low dependence) indicates that no NRT is needed; a score of 3 to 4 (low to moderate dependence) requires 1 NRT; and scores of 5 to 7 (moderate dependence) and ≥8 (high dependence) require a combination of NRTs.

In the EAGLES trial, all participants smoked at least 10 cigarettes per day, and...
had moderate dependence, with an average FTND score of 5 to 6.

2. What stage of change is the patient in, and how many times has he or she attempted to quit? Based on the answers, motivational interviewing may be appropriate.

In the EAGLES trial, the participants were motivated individuals who had an average of 3 past quit attempts. Research suggests that even patients who have a serious mental illness can be motivated to quit (Box).6-9

3. What is the patient’s mental health status? What is the patient’s psychiatric diagnosis and how clinically stable is he or she? What is his or her suicide risk? Consider using the Columbia Suicide Severity Rating Scale (C-SSRS).10

In the EAGLES trial, the psychiatric cohort included only patients who had been clinically stable for the past 6 months and had received the same medication regimen for at least the past 3 months, with no expected changes for 12 weeks. Patients with certain diagnoses were excluded (eg, delusional disorder, schizophreniform disorder, impulse control disorders), and only 1% had a personality disorder, which increases mood lability and likelihood of suicidality behavior.

4. Does the patient have another comorbid substance use disorder? In the EAGLES trial, those who had active substance use in the past year or were receiving methadone or buprenorphine/naloxone were excluded.

5. Does the patient have any medical conditions? Does he or she have a history of seizures or eating disorders? It is important to determine if a patient has a seizure disorder or another medical condition that is a contraindication for using varenicline or bupropion.

In the EAGLES trial, most adverse effects related to the medications administered involved sleep (insomnia) or the gastrointestinal system (nausea). The psychiatric cohort reported some anxiety with bupropion.

6. Have you discussed smoking cessation and a treatment plan with the patient at every visit? In the EAGLES trial, participants received 10-minute cessation counseling at every outpatient visit.

When it comes time to select a medication regimen, for bupropion, consider starting the patient with 150 mg/d, and increasing the dose to 150 mg twice a day 4 days later. The target quit date should be 7 days after starting the medication. Monitor the patient for symptoms of anxiety and insomnia.

For varenicline, start the patient at 0.5 mg/d, and increase the dose to 0.5 mg twice a day 4 days later. After another 4 days, increase the dose to 1 mg twice a day. Set a target quit date for 7 days after starting medication. Monitor the patient for nausea, insomnia, and abnormal dreams.

References

Research suggests that even patients who have a serious mental illness can be motivated to quit smoking.


