Executive Summary

Overview: Evidence has been mounting over a short period and now shows a significant association between smoking and increased severity of COVID-19. While it is biologically plausible, evidence of the association between smoking and susceptibility to becoming infected with SARS-COV-2 is limited and inconclusive. There remains sound evidence to continue to support quitting during this time, and COVID-19 may increase motivation to quit.

Severity: Current evidence shows that smoking status is significantly associated with severity of COVID-19. A large study (n=8,910) of COVID-19 patients found that being a current smoker was independently associated with an increased risk of in-hospital death (OR 1.79, 95% CI 1.29-2.47, adjusted for age, sex and other known major risk factors such as heart disease).

Susceptibility: While it is biologically plausible that smoking may increase susceptibility to contracting COVID-19, there is currently no strong evidence that smoking increases susceptibility to COVID-19.

Shisha: The World Health Organization has recommended a ban of the public use of shisha (e.g. in cafes bars and restaurants) during the COVID-19 pandemic to reduce potential transmission from sharing mouthpieces and pipes. Shisha was previously implicated in an outbreak of tuberculosis in Queensland.

Vaping: No direct evidence is available regarding vaping and COVID-19. Vaping appears to be a risk factor for respiratory infections in general.

The benefits of quitting during the COVID-19 pandemic: No current evidence exists to demonstrate that quitting will reduce negative outcomes for COVID-19. However, there are many immediate benefits from quitting, including to the respiratory and cardiovascular systems, which may plausibly have ‘pre-habilitation’ benefits that flow on to better outcomes.

Current messaging: Many jurisdictions, including SA, are encouraging smokers to quit during this time for their respiratory health. The UK has been encouraging smokers to quit, with messaging saying that smokers are 14 times more likely to develop severe COVID-19 disease. Media reports indicate that this has prompted 300,000 smokers to quit (note: unpublished preliminary data).

The World Health Organization advised that smoking makes you more vulnerable to COVID-19 infection and recommended that people quit smoking (24 March 2020).

Context

- COVID-19 is predominantly a disease of the respiratory tract. Comorbidities associated with smoking including chronic obstructive pulmonary disease (COPD), cardiovascular disease (CVD), diabetes and hypertension are also associated with severe disease among patients with COVID-19. Smoking is a well-established cause of many of these medical comorbidities.
• Many agencies are currently encouraging smokers to quit, in light of COVID-19 and its impact on the respiratory system.

Summary of the evidence

Current evidence suggests that smoking status is significantly associated with severity of COVID-19.

• A large study (published in the NEJM), of 8,910 patients hospitalised with COVID-19 across Europe, Asia, and North America found that being a current smoker was independently associated with an increased risk of in-hospital death from COVID-19 (OR 1.79; 95% CI 1.29-2.47), adjusting for other known risk factors including age, sex, coronary artery disease, congestive heart failure, arrhythmia, chronic obstructive pulmonary disease, and receipt of ACE inhibitors, ARB, or statins (1). In univariate analyses, being a former smoker was not associated with death.

• A systematic review of 11 smaller studies reporting on a total of 2002 patients with COVID-19 from China similarly found that ongoing smoking history was significantly associated with the development of severe disease (leading to ICU admission, mechanical ventilation or death; unadjusted OR 1.98; 95% CI 1.29-3.05) (2).

There is currently no strong evidence that smoking increases susceptibility to COVID-19.

• Studies published to date using data from China (3-14) and the US (15) have reported levels of smoking prevalence among patients hospitalised with COVID-19 which have been noted as being much lower than smoking prevalence in the general population in those countries. These findings must be interpreted with caution. It is likely that smoking status has not been comprehensively recorded in the data analysed to date (completeness of smoking data may not have been a priority for medical professionals, especially in the face of overwhelmed medical systems). Complete data were only available for 5.8% of COVID-19 patients in the US study, for example (15). Stronger designs, e.g. cohort studies will be required to make a more accurate comparison of smoking prevalence in people with and without COVID-19.

However, it is biologically plausible that smoking increases susceptibility of COVID-19 for the following reasons:

• An extensive review of the literature by the US Surgeon General published in 2014 indicated that while the mechanisms are complex and not well understood, smoking is generally detrimental to the immune system and its responsiveness to infections (16). (Note: Evidence prior to COVID-19)

• Smoking is also known to be a major risk factor for acute respiratory tract infections in general (17). Evidence has found previously that cigarette smoking can increase the incidence, duration, and/or severity of respiratory viral infections (18). Smokers were also found to have higher mortality in the
MERS-CoV outbreak, a virus with some similarities to SARS-CoV-2 (19). (Note: Evidence prior to COVID-19)

- The SARS-CoV-2 virus enters cells via binding to the ACE2 receptors present on the surface of respiratory cells. Some studies have found that smoking increases expression of ACE2, the receptor for COVID-19 (20-22), which could theoretically increase susceptibility and severity. On the other hand, a group of French researchers (pre-print) have hypothesized that SARS-CoV-2 might be a nicotinic agent, meaning that nicotine might be used to treat acute infection (23). This study has received considerable media attention. However, critiques have been made of the study methodology, and one of the study co-authors (Jean-Pierre Changeux) has previously received funding from the tobacco industry. Awaiting the peer-reviewed version of this study is recommended.

- Smoking involves repetitive hand-to-face actions which may help to transmit the disease (24).

Electronic cigarettes / vaping

- No direct evidence is currently available regarding vaping and COVID-19. However, vaping also appears to be a risk factor for respiratory infections in general, with human and animal studies indicating that e-cigarettes impair immune systems, leading to an increased susceptibility and/or delayed recovery from infection (25).

Shisha and waterpipe tobacco smoking may pose additional risks of infection from shared use of the mouthpiece.

- Waterpipe use has been implicated in an outbreak of tuberculosis (15) and in the transmission of fungal infections (16). There is similarly a risk of transmission of viruses (17).
- Waterpipe tobacco smoking may pose additional risks of infection from shared use of the mouthpiece and other components (26).
- The Waterpipe Tobacco Smoking Knowledge Hub of the World Health Organization Framework Convention on Tobacco Control recommends a complete ban of the use of waterpipes in all public establishments to help limit the transmission of COVID-19 (18).
- News reports indicate that many middle eastern countries including Iran, Kuwait, Pakistan, Qatar and Saudi Arabia have banned the use of shisha in public places to avoid COVID-19 transmission.

Is this global pandemic a good time to recommend that people to quit smoking?

Evidence supports quitting at all times:

- While there isn’t direct evidence that quitting smoking now will help smokers avoid more severe consequences of COVID-19, being a former smoker was not associated with an increased risk of
death (univariate analysis) in the large multi-country study of 8,910 COVID-19 patients cited above (1).

- Quitting smoking reduces rates of respiratory infections in general (27). Airway function improves within four weeks of quitting, and the immune systems improves within a few months (28).

- Quitting smoking reduces risk of heart disease, which is associated with increased severity of COVID-19 (1, 29). After quitting smoking, the risk of coronary heart disease decreases rapidly. Within 20 minutes of quitting, blood pressure drops, and the risk of heart disease halves within one year (27). People who have already been diagnosed with heart disease can lower their risk of dying by quitting smoking (30).

- Encouraging people to quit now could help to reduce demands on the health system by reducing hospitalisations associated with conditions that are quickly reversible by quitting, such as pneumonia (28), compromised wound healing and respiratory complications after general surgery (31), and pregnancy complications including low birthweight (28), placental abruption (32) and ectopic pregnancy (16).

What are selected tobacco control experts and health organisations recommending?

- South Australia
  - SA Government now has a range of messages encouraging smokers to quit.
  - Tobacco smoking is monitored on a monthly basis through SA Health, and changes to tobacco use and quit attempts will be monitored during the epidemic.
  - Number of calls and contacts with the SA Quitline will also be monitored. Data shows that there was a 10% increase in the number of calls to the SA Quitline in the first quarter of 2020 compared to the first quarter of 2019, with the majority of the increase seen in February and March. Quitline counsellors have reported that while callers are not citing COVID-19 as their primary reason for wanting to quit, it is a topic that is regularly forming part of the call conversation, suggesting it is near the front of callers’ minds.

- Quit Victoria is about to commence a digital campaign including a tailored Quitline ad encouraging smokers to quit during COVID-19. The key message is that physical distancing gives smokers an opportunity to change their habits now they are removed from social triggers. Quit Victoria has also compiled some FAQs regarding COVID-19 and smoking.

- An article by Australian tobacco control experts in The Conversation on 14 April advised that there’s never been a better time to quit because smoking increases COVID-19 risk.

- World Health Organization:
  - The Waterpipe Tobacco Smoking Knowledge Hub of the World Health Organization Framework Convention on Tobacco Control has recommended a complete ban of the use of waterpipes in all public establishments to help limit the transmission of COVID-19 on 13 March.
The World Health Organization advised that smoking makes you more vulnerable to COVID-19 infection and recommended that people quit smoking, as well as posting a brief Q and A on smoking and COVID on 24 March.

- **Public Health England** has stated that smokers are 14 times more likely to develop severe disease (a figure from one of the studies (5) in the meta-analysis (2) described above) and issued specific advice against smoking on 3 April. An article in The Guardian on 4 May reported that 300,000 UK smokers may have quit due to COVID-19, based on unpublished survey data that found that 2% of smokers had quit because of concerns about coronavirus; 8% were trying to quit; 36% had cut down; and 27% were now more likely to quit.

- **The US Food and Drug Authority** currently advises people to quit smoking as smoking can leave you more vulnerable to respiratory illnesses such as COVID-19 (last updated 1 May 2020).

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References

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