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of rapid waning of post-booster immunogenicity and immune escape of new variants of concern. Heterologous boosters with next-generation vaccines, such as multivalent vaccines (vaccines providing protection against different variants simultaneously), universal coronavirus vaccines, vaccines eliciting stronger T-cell responses, or mucosal vaccines (either intranasal or oral), are among the future options for COVID-19 vaccination. However, while awaiting these next-generation vaccines, booster immunisations are crucial to restore vaccine effectiveness against severe outcomes in clinically vulnerable populations. The results of this trial are important to help policy makers to determine who benefits most from booster dosing and when booster dosing should be implemented. The question of whether benefit can be gained with longer delays between boosters remains unanswered.

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Risk of arterial and venous thromboses after COVID-19



Infection can trigger thrombotic events. After respiratory and other infections, people have a 3–6-fold increased risk of arterial thrombosis, such as myocardial infarction and ischaemic stroke, and a 2–3-fold increased risk of venous thromboses, such as deep vein thrombosis of the legs and pulmonary embolism.^{1,2} The risk declines in the weeks after infection, although a higher risk can persist for a year or longer, particularly for venous thromboses.²

People with severe COVID-19 have a high risk of symptomatic and asymptomatic pulmonary emboli during their hospital stay.³ However, the longer-term risks of thrombotic events after mild COVID-19 are less clear, and a better understanding of the future risk of heart attack and stroke is a priority for people affected by COVID-19.⁴

In *The Lancet Infectious Diseases*, Edward Burn and colleagues⁵ report the 90-day cumulative incidences of venous or arterial thromboembolism and death after a COVID-19 diagnosis in primary care datasets from five countries: the Netherlands, Italy, Spain, the UK, and Germany. The study showed substantial

variation in the 90-day cumulative incidence following COVID-19 diagnosis between the different countries: for venous thromboses from two per 1000 in the Netherlands to eight per 1000 in Spain; and for arterial thromboembolism from one per 1000 in the UK to eight per 1000 in Spain. The incidence of venous and arterial events was higher in older people, and the risk of death after venous and arterial events was higher in people who had been diagnosed with or tested positive for COVID-19 than in people without COVID-19. In other studies, the cumulative excess risks up to 49 weeks after a COVID-19 diagnosis or positive test in linked primary and secondary care databases in England were 25 per 1000 for arterial and six per 1000 for venous thromboses,⁶ and the cumulative risk of venous events in Sweden up to 30 days after a COVID-19 diagnosis or positive test was about two per 1000.⁷

The wide variation in the incidence across countries highlights the challenges in combining estimates from different health-care systems and with different public health policies. Each primary care system has different coding practices, populations, and linked

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datasets to ascertain risk. Although the authors made considerable efforts to analyse their data to a common data model, linkage to hospital records (and hence the completeness of ascertainment of thrombotic events) varied between primary care systems, and different workloads and patterns of use of primary health care might affect coding. The different timing of vaccination programmes between countries could have led to different secular changes in the incidence and severity of COVID-19 and its consequences. However, multicountry studies such as this are crucial for helping to build an evidence basis for decisions about prioritising public health.

The prevention of arterial or venous thromboses through vaccination against common infections, or other population-level approaches, is appealing. Influenza vaccination reduces the relative risk of major cardiovascular events by about a third, from a meta-analysis of randomised controlled trials, and observational data suggest that COVID-19 vaccination has a similar protective effect in older people, although is subject to biases and residual confounding.⁸ Results of further studies in high-risk individuals are awaited.⁹ Although acute antithrombotic therapy might reduce the short-term risk of venous thromboses (with an increased risk of haemorrhage) after infection with SARS-CoV-2, aspirin does not seem to be of overall benefit.¹⁰ New trials of aspirin use could answer this question definitively for non-COVID-19 infections such as pneumonia.

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For more on trials of aspirin use to prevent cardiovascular events following pneumonia see <https://fundingawards.nihr.ac.uk/award/NIHR132968>



The state of tuberculosis in South Africa: what does the first national tuberculosis prevalence survey teach us?

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South Africa is among WHO's list of 30 high-burden tuberculosis countries and has one of the highest incidence rates of notified tuberculosis in the world.¹ In *The Lancet Infectious Diseases*, Sizulu Moyo and colleagues² report their findings from the first national tuberculosis prevalence survey in South Africa, which is a very important study that provides improved understanding of the true extent of tuberculosis and helps to identify groups who might be underserved by health services and where tuberculosis might be undiagnosed. Tuberculosis prevalence surveys are

massive endeavours; this survey included more than 35 000 participants across all nine provinces of South Africa, all of whom were screened using a symptom questionnaire and chest X-rays, and more than 9000 were eligible to provide sputum samples for Xpert MTB/RIF Ultra assay testing and mycobacterial culture. Although diagnosis of tuberculosis has improved in recent years it is still fraught with challenges. This survey, like many before it, has shown the inadequacy of using symptom screening to identify people living in the community who have culturable *Mycobacterium*