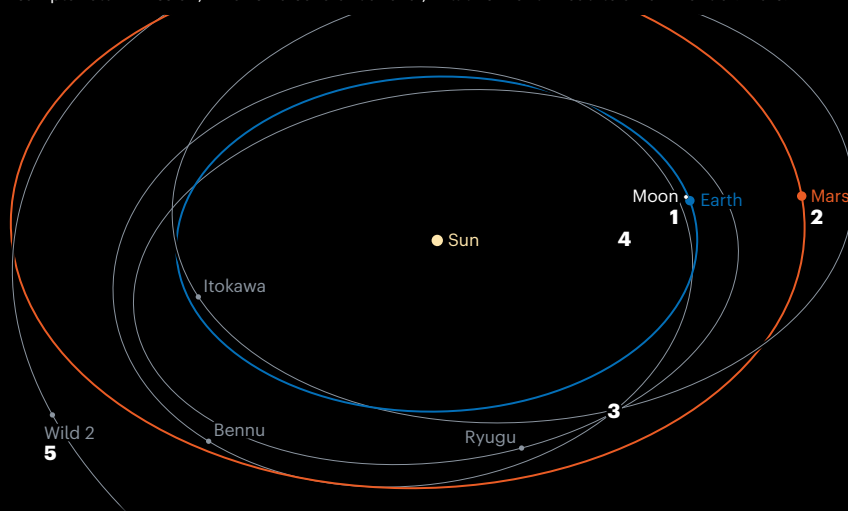


SAMPLING THE SOLAR SYSTEM

Agencies have sent spacecraft to the Moon, asteroids and comets, and into the middle of the solar wind, to grab rocks and particles and return them to Earth. These materials have transformed our understanding of the Solar System and its bodies. The next step towards a sample-return mission, NASA's Perseverance rover, will this month head to a new frontier: Mars.



1. Moon

1969–72: NASA's Apollo missions collected 382 kilograms of Moon rocks.
1970: Soviet Luna-16 mission, 101 grams.
1972: Luna-20, 55 grams.
1976: Luna-24, 170 grams.
Future: China's Chang'e-5 probe, NASA's Artemis missions.

2. Mars

Future: Japan's Martian Moons Exploration mission could return material from the satellite Phobos as early as 2029. NASA's Perseverance rover will fill 30 tubes with rock and soil on Mars, to be returned to Earth by 2031.

3. Asteroids

2005: Japan's Hayabusa probe collected more than 1,500 particles from asteroid Itokawa.
2019: Hayabusa2 left asteroid Ryugu to bring samples to Earth.
Future: NASA's OSIRIS-REx will collect samples from asteroid Benu.

4. Solar wind

2001–04: NASA's Genesis spacecraft collected more than 1,850 samples.

5. Comet Wild 2

2004: NASA's Stardust mission collected more than 10,000 particles of dust from Wild 2's tail.

should arrive on Earth soon. In December, JAXA's second asteroid mission, Hayabusa2, should return material from a carbon-rich asteroid called Ryugu, and NASA's OSIRIS-REx spacecraft is orbiting the diamond-shaped Benu asteroid, in the hope of grabbing a sample in October and returning to Earth in 2023.

NASA has also grabbed material from a comet, with its Stardust spacecraft. In 2004, the mission whizzed through the tail of Comet Wild 2. On Earth, the samples it gathered turned up huge surprises.

NASA named the mission Stardust because scientists thought the comet

contained ancient dust from other stars, frozen in ice for billions of years. "This idea was also spectacularly wrong," says Brownlee, the mission's principal investigator. Instead, they found the comet dust had formed close to the Sun at incandescently hot temperatures. That showed that hot materials had been transported throughout the early Solar System and somehow become incorporated into the icy body of the comet.

But returning samples from Mars is a bigger challenge than any other mission so far. The planet is farther away than the

Moon and has more gravity than a comet or an asteroid, making it harder to escape the surface and get back to Earth.

NASA wants Perseverance to drill out and store at least 30 tubes of Martian rock and soil at its landing site in Jezero Crater. Long-term plans call for NASA and the European Space Agency to collaborate to send a second rover to collect those tubes and launch them into Martian orbit, and a third spacecraft to fetch them from Martian orbit and fly them back to Earth. The aim is for the samples to reach Earth in 2031.

Alexandra Witze

CHINA'S CORONAVIRUS VACCINES ARE LEAPING AHEAD

But companies could struggle to run trials in enough people to satisfy regulators.

By David Cyranoski

Chinese companies are at the forefront of global efforts to create a vaccine for the coronavirus, with more than half a dozen candidates in clinical development. Last month, Tianjin-based CanSino Biologics published results¹ from an early clinical trial showing that its vaccine is

safe and can trigger an immune response.

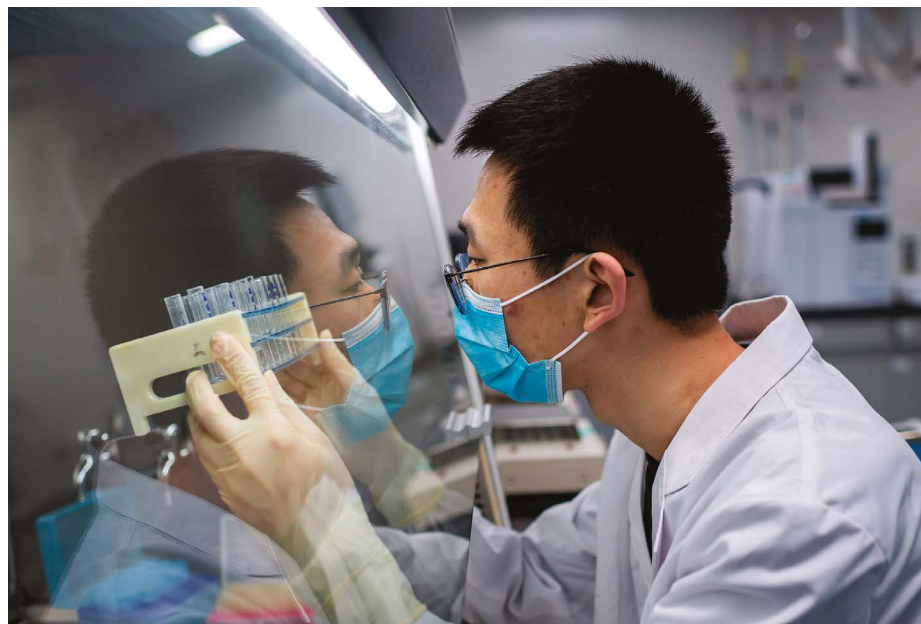
Yet the companies could face difficulty as they try to push vaccines through phase III trials, a crucial stage of testing that is needed to prove efficacy and secure approval from regulators. These trials usually require tens of thousands of participants, and with the outbreak in China largely under control, companies are having to test their vaccines

elsewhere. But researchers say they might still struggle to enrol so many participants and employ enough health-care professionals to collect data.

"The Chinese companies will need to step outside of China," says Jerome Kim, director-general of the International Vaccine Institute in Seoul. "The race is on," he says, "and it's really about who can set up in a high-risk area most quickly."

Chinese vaccine-makers will face other challenges, too. Vaccines will probably face extra scrutiny, given the country's opaque regulatory system and previous vaccine scandals, say scientists. In 2018, hundreds of thousands of children reportedly received defective vaccines against diphtheria, tetanus and whooping cough.

As the country where the first cases of the coronavirus were reported, China was fast out of the gate in developing vaccines. CanSino's offering is made from a common-cold virus,



Chinese companies have made several vaccines that are currently being trialled.

tweaked to mimic the coronavirus. Sinopharm, a state-owned pharmaceutical company in Beijing, is developing two vaccines made using particles of the coronavirus that have been inactivated so that they can no longer cause disease. The company said in press releases in June that both vaccines had produced antibodies in all participants in preliminary phase I and II trials. And Beijing-based company Sinovac has announced similarly promising results for its own inactivated-virus vaccine.

The next phase

This month, Sinovac launched a phase III trial of its vaccine in Brazil. Sinopharm will be testing its inactivated vaccines in the United Arab Emirates (UAE). Only three other coronavirus vaccines have entered phase III trials: one produced by biotechnology company Moderna in Cambridge, Massachusetts; one by the University of Oxford, UK, and drug maker AstraZeneca, based in Cambridge, UK; and one by biotech company BioNTech of Mainz, Germany, in collaboration with New York City-based drug firm Pfizer.

CanSino is also poised to launch a phase III trial. But the Chinese government has already said that its vaccine can be used by the military – making CanSino the first company to have a vaccine for COVID-19 approved for limited use in people. China has worked hard “to generate an efficient vaccine as soon as possible and to be transparent” when doing so, says Stéphane Paul, a vaccine researcher at the University of Lyon in France.

The speed with which Chinese vaccine-makers are moving has raised hopes around the world. Sinopharm has even promised to have a vaccine ready to distribute by the end of this year.

Inactivated vaccines are widely used, so it

makes sense for Chinese companies to focus on them, says Paul. “As a first line of vaccine, it is immunogenic, quick to develop and low-cost,” he says.

But some viruses become more potent when they infect organisms previously treated with inactivated vaccines, in a poorly understood phenomenon known as antibody-dependent enhancement (ADE). This was reported² last year in monkeys given a vaccine for the coronavirus that causes severe acute respiratory syndrome (SARS). Sinovac says its COVID-19 vaccine did not trigger ADE in monkeys, but the risk will be closely monitored in all the inactivated-vaccine phase III trials, says Paul.

Some observers also question whether Chinese companies will be able to work at the promised

“The race is on, and it’s really about who can set up in a high-risk area most quickly.”

speed, and with the precision required. And the fact that China was willing to approve CanSino’s vaccine for use in the military without phase III trials raised eyebrows. “The decision is political, and not scientific in nature. It doesn’t demonstrate anything on the potential efficacy of this vaccine,” says Marie-Paule Kiény, a vaccine researcher at INSERM, the French national health-research institute, in Paris.

Phase III trials present challenges for vaccine makers around the world, such as the need to recruit enough participants and qualified health staff. Demonstrating that vaccines provoke an immune response and protect people from the virus requires data on 20,000–40,000 people who have been split

into control and test groups and then followed closely for several months or even years, scientists say. To reach the numbers required, the trials might need to combine results from dozens of hospitals, each supplying data from hundreds of patients. “All of these things have to be done, and done correctly,” says Kim. “The number of sites that can do this and handle the volume is limited,” he adds. “Even the best sites will have difficulty.”

Many Chinese companies are at a disadvantage because they don’t have established networks of hospitals around the globe, says Kim. AstraZeneca published³ promising early trial results for its vaccine – based on a chimpanzee cold virus – on the same day as CanSino, and is carrying out phase III studies in the United Kingdom, Brazil and South Africa. Moderna has launched a trial of its vaccine, which elicits an immune response with synthesized RNA that mimics the RNA that the coronavirus uses to replicate, in 30,000 people across the United States. That country has a lot of experienced clinical researchers to carry out trials, and is tackling a large coronavirus outbreak.

International links

US President Donald Trump said last week that he was willing to work with any country that can deliver an effective vaccine, but Chinese companies had previously been ruled out of receiving funding from the US government’s Operation Warp Speed, which aims to accelerate vaccine development.

Still, Kiény points out that Sinopharm has partnered with the UAE’s government and Group 42 Healthcare, a local artificial-intelligence company, for its phase III trial, and that Sinovac has partnered with the Butantan Institute in São Paulo, Brazil. “So far, Chinese companies seem to have been successful in finding partners,” she says.

But some researchers question whether the trials in the UAE and Brazil will gather enough data to convince regulatory agencies that the vaccines work. In the UAE, where Sinopharm plans to enrol 15,000 participants to study its two vaccines, relatively few people are infected with COVID-19.

And although Brazil has a large coronavirus outbreak, the Butantan Institute plans to test Sinovac’s vaccine among health-care professionals because it is assumed they will face greater exposure to the virus than will other groups. Because of this, the trial will enrol only 9,000 people to test whether it works, says Ricardo Palacios, a clinical researcher at the institute who is leading the trial. “We designed a trial in order to obtain answers in a more efficient way,” says Palacios.

1. Zhu, F.-C. et al. *Lancet* [https://doi.org/10.1016/S0140-6736\(20\)31605-6](https://doi.org/10.1016/S0140-6736(20)31605-6) (2020).

2. Liu, L. et al. *JCI Insight* **4**, e123158 (2019).

3. Folegatti, P. M. et al. *Lancet* [https://doi.org/10.1016/S0140-6736\(20\)31604-4](https://doi.org/10.1016/S0140-6736(20)31604-4) (2020).