

# COVID-19 and Promising Expected Vaccines

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## ABSTRACT

**Background:** In spite of all precautionary measurements have been done to control the spread of SARS-COV2, the mortality and morbidity have increased, and the need for an effective vaccine is growing daily

**Aim:** In this review, the authors provide an overview of the current COVID-19 vaccines under the spotlight.

**Conclusion:** Despite of the urgent need for a COVID-19 vaccine, several practical and technical issues remain, but recent vaccine developments by many companies and organizations seem promising and could make us closer to reduce mortality and morbidity rates.

**KEYWORDS:** Coronavirus; SARS-CoV-2; COVID-19; Vaccine; Immune response; Pandemic

**ABBREVIATIONS:** ARDS: Acute Respiratory Distress Syndrome; COVID-19: Corona Virus Disease of 2019; H1N: Influenza A virus subtype; MERS: Middle East Respiratory Syndrome; SARS: Severe Acute Respiratory Syndrome; RT-PCR: Real Time Polymerase Chain Reaction; APC: Antigen-presenting cell; ChAdOx1 nCoV-19; CH: Chimpanzee; AD: Adenovirus-Vectored Vaccine; Oxford Novel Coronavirus 2019; AZD1222: AstraZeneca Vaccines; CV: Coronavirus Vaccines; AD26: Adenovirus type 26; AD5: Adenovirus type 5; NOVAVAX: Nova Vaccines

## INTRODUCTION AND BACKGROUND

Extreme acute respiratory distress syndrome (ARDS) caused by SARS-CoV-2 emerged as a local outbreak in Wuhan province at the end of December 2019, but later, when the disease spread to many countries worldwide, WHO declared SARS-CoV-2 as a pandemic

disease called COVID-19 [1]. The SARS-CoV-2 genomic sequence analysis revealed its close resemblance to the SARS-COV and the Middle East Respiratory Syndrome coronavirus (MERS-COV) [2]. For the SARS-CoV-2 structure in Figure 1.

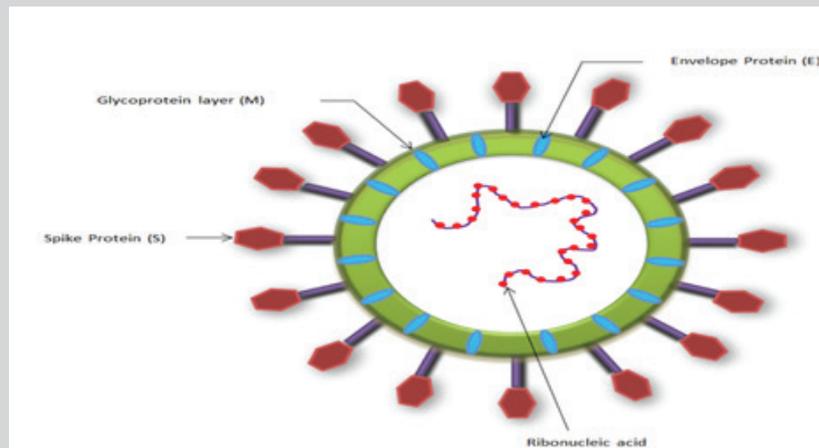
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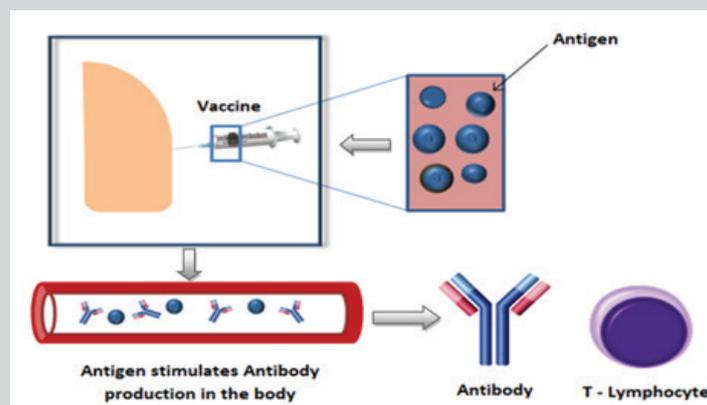
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**Figure 1:** Illustrates the main structural features of Coronavirus SARS-CoV-2.

After identifying the genus of COVID-19, the world started its journey towards finding an ideal diagnostic tool for this disease, until reached that PCR is the standard golden method, at least up to date [3,4]. Then the journey is continuing to find out an efficient COVID-19 treatment strategy. Even though, the treatment line is unclear enough and representing a debated issue among medical communities, but it seems that the promising strategy against

COVID-19 is the vaccination process. The process of COVID-19 vaccines manufacturing has been approved by health authorities and organizations in many countries. Moreover, some countries have started the COVID-19 vaccine campaign for their citizens, such as the UK, USA and Russia. Herein, the narrative review provides an overview of the current COVID-19 vaccines under the spotlight.



**Figure 2:** Schematic representing the administration of target antigen through a vaccine to stimulate the production of Antibodies and T-cell.

## CURRENT STATUS OF COVID-19 VACCINE

A vaccine is a substance that stimulates the immune system to produce antibodies, precisely as if it was exposed to the disease [5] in Figure 2. Generally, vaccine is divided into different types, including inactivated vaccines, live attenuated vaccines, vectored vaccines (DNA/ RNA vaccines), nucleic acid-based vaccines, and recombinant subunit vaccines [6-9].

Currently, there are more than 300 COVID-19 vaccine candidates [10]. Most of those vaccine candidates are developed relied on the previous experience with other coronaviruses to prompt humoral antibodies against the viral particles such as spike protein or receptor binding domains [11]. The larger parts of COVID-19 vaccines are at the pre-clinical phase (phase I and II).

Phase I to investigate the vaccine safety in small group, then phase II to test the immunogenicity in large subjects and phase

III to estimate efficacy and safety of a vaccine to take decision for approval [12]. Currently, some of vaccines are authorized and recommended to prevent COVID-19.

### Pfizer Inc. and BioNTech

Pfizer Company is a collaboration between Americans and Germans scientists through Pfizer Inc. and BioNTech SE. This collaboration led to innovate a vaccine candidate against COVID-19 [13]. Pfizer's vaccines based on using mRNA- candidate, which is considered as a new vaccines technique has never been approved in a vaccine before. According to Pfizer's announcement, the vaccines have demonstrated an efficacy rate of 95% after the first week of the second dose (3 weeks apart from the first dose) [14]. Data Monitoring Committee (DMC) has not raised any significant safety issues but has recommended that additional safety and efficacy data continue to be collected as planned and addressed consistently with regulatory authorities worldwide [15].

Regarding storage and distribution for vaccines, Pfizer's vaccines need to be stored in  $-70\text{ }^{\circ}\text{C}$  [16]. This frozen chain is going to be one of the most challenging aspects of the delivery of this vaccination because it will be difficult for rural areas and developing countries to access these types of ultra-cold freezers.

### Moderna's Vaccine (mRNA-1273)

This type of Vaccine is collaboration between the National Institute of Allergy and Infectious Diseases and Moderna, Inc in USA [17]. Moderna's vaccine against COVID-19 relies on a novel technology that uses messenger RNA (mRNA) similar to Pfizer's vaccine which also depends on mRNA to express a protein called spike [18]. According to the developers of this type of vaccines say, one of the mRNA advantages is faster and simpler to manufacture than traditional vaccines, which can take time to develop because researchers have to grow and inactivate whole pathogens or their particles [16]. Even though, this novel mRNA vaccine might be more durable against pathogens that tend to mutate, such as coronaviruses, but still the instability and breaking down by nuclease enzymes are main concerns [19,20].

The Moderna vaccine was 94.1% effective at blocking symptomatic Covid-19, measured starting from 14 days after the second dose (28 days from the first dose); [21]. It also has the same difficulty of storage in cold temperatures requires to be stored at  $-20\text{ }^{\circ}\text{C}$  [22].

### ChAdOx1 nCoV-19 Vaccine (AZD1222)

ChAdOx1 nCoV-19 vaccine popularly known as the Oxford vaccine was developed by Oxford University in collaboration with the biopharmaceutical company AstraZeneca [23]. The Oxford vaccine is composed of a replication-deficient chimpanzee adenoviral vector ChAdOx1 carrying the SARS-CoV-2 structural surface glycoprotein antigen (spike protein; nCoV-19) gene [23]. The mechanism of vaccine here based on inducing T cell lymphocytes within the first two weeks of vaccination and later after 28 days occurs humoral response (provoke of antibodies immunity) [24]. Although the Oxford vaccine is a type of a weakened or Live-attenuated vaccines structure of a live virus, which can induce long-term of immune responses, however, this type of vaccines tends to be risky for those people with weakened immune systems or have other health issues [25].

### Sinovac's COVID-19

CoronaVac is another type of COVID-19 vaccines that synthesized by a Chinese company called Sinovac Biotech. The CoronaVac used a traditional inactivated virus technique (an inactivated version of the SARS-CoV-2 virus). This technique is relied on a quick triggering the immune system to produce a humoral response within 4 weeks of immunization by giving 2 doses of the vaccine at a 14- day [26]. Although CoronaVac produced levels of antibodies lower than those reported in people who had recovered from Covid-19, but the CoronaVac innovators believe it could still provide protection from the unprecedented coronavirus [27]. This lower level of antibodies demonstrates what was published in previous studies that, inactivated virus technique is considered as not fully protective as live vaccines and might require adjuvant over time [25]. One of Sinovac's main advantages is that CoronaVac could be stored at normal refrigerator at  $2-8\text{ }^{\circ}\text{C}$  [27]. Sinovac's COVID-19 vaccine has been approved by Chinese health authorities to use in emergency case [28,29].

### Sputnik V

The name of Sputnik came from the first Soviet space satellite. Sputnik V is the first vaccine against COVID-19 approved by Russian authorities. It's based on two vectors, which include adenovirus vector 1 AD26 and adenovirus vector 2 AD5 [30]. This is a novel vectors technique developed by Gamaleya center of Epidemiology and Microbiology. This technique is thought to produce a longer-term immune response as compared to the vaccines using one and the same vector. The Sputnik V is available in dry form (lyophilized), which makes it easier to storage ( $2-8\text{ }^{\circ}\text{C}$ ) and distributes [31].

### NOVAVAX

NOVAVAX is an American company that has developed and is testing a candidate coronavirus vaccine called NVX-CoV2373. It's a type of nanoparticle vaccine consists of trimeric full-length SARS-CoV-2 spike glycoproteins and Matrix-M1 adjuvant [32]. The purpose of Matrix-M1 adjuvant is stimulating the entry of antigen-presenting cells (APC) into the injection site and thus enhancing the presentation of antigen to near lymph nodes, hence NVX-CoV2373 induced humoral immune response that block attaching of spike protein to cellular receptors and provided immunity from infection [33]. The storage condition of NOVAVAX is convenient, it is transferred and handling in an unfrozen atmosphere, liquid formulation allows to be kept at  $2\text{ }^{\circ}\text{C}$  to  $8\text{ }^{\circ}\text{C}$ , therefore for distribution could use standard vaccine channels [34].

### RE-INFECTION AND VACCINE

The question of the possibility of reinfection occurrence after taking the vaccine still is remaining unclear. Nevertheless, if SARS-CoV-2 follows the previous family of its coronavirus cousins, so, reinfection will soon become the rule, instead of the exception [35,36]. As it is seen in common-cold- to infect the same individual multiple times.

### VACCINE PRIORITY

Nearly, most of the Committees that supervise on vaccination campaigns are almost agreed to divide the vaccines takers into Priority groups. For instance, a high risk of severe illness in high-exposure occupations, Frontline health workers, and vulnerable people such as the eldest aged 70 years and over [37].

### TREATMENT

So far, there is no unified approach to agreed-upon COVID-19 treatment. However, there is one international trial called Solidarity. It launched by the WHO and partners, aims to find out an effective treatment for COVID-19 [38]. This trial evaluated four types of treatments, Remdesivir, Hydroxychloroquine, lopinavir/Ritonavir and Interferon. It found those treatments had insignificant effect on overall mortality and hospitalized patients [39]. Only corticosteroids have proven to some extent effective against critical COVID-19 patients cases [40].

### QUESTIONS TO BE ADDRESSED

There are some concerns among the public population about receiving COVID-19 vaccines. They are right toward these concerns because of the novelty and rapid development of it. Below are some questions that need convincing answers from the vaccine's developer.

1- How is the vaccine safety was evaluated? People who are vaccinated for trials and followed up for time period, is this period sufficient to detect most safety issues?

2- Will the Vaccine Be Effective for All Patients?

3- Should Testing for COVID-19 continues after vaccination, and if so, for what duration?

4- Is there is a transparency and equality in COVID-19 distribution for those vulnerable groups?

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Ebtehal M Fawzi, Sheima A Elbasheer, Alaa A Elhoussein, conceptualization and wrote the original draft; Ahmed O Abu Elhassan, Nagia S Ahmed, Mayada K Ali Acquisition of data and designed; Abdelmonem M Ali, Alfatih Aboalbasher Yousif, reviewed the final draft. All authors approved the final version of the article.

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