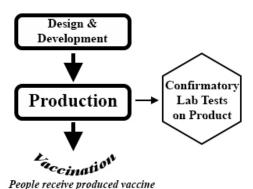


## **Update: COVID-19 Vaccine Candidates and Abortion-Derived Cell Lines**

Accurate information about the development and production of COVID-19 vaccines is essential, especially because many proposed candidates use newer molecular technologies for production of a viral vaccine. One concern regarding the ethical assessment of viral vaccine candidates is the potential use of abortion-derived cell lines in the development, production or testing of a vaccine. This analysis utilizes data from the primary scientific literature when available, along with data from clinical trial documents, reputable vaccine tracking websites, and published commercial information.<sup>1</sup> It is the hope that by providing accurate data, recipients can make well-informed decisions regarding vaccine choices.

## For additional background and guidance, please see:

- \* A Visual Aid to Viral Infection and Vaccine Production for a visual primer on the various strategies for viral vaccine production.
- \* An Ethics Assessment of COVID-19 Vaccine Programs for discussion of ethical considerations in viral vaccine production.
- \* COVID-19 Vaccines & Fetal Cell Lines for an infographic description of how fetal cell lines are sometimes used to produce vaccines.



## Flow Chart for Creation and Testing of Vaccines

<u>Design & Development</u>: conceptualization, preparatory experiments, and specification for how vaccine will be constructed and produced.

<u>Production</u>: process used to manufacture final vaccine to be given to people.

<u>Confirmatory Lab Tests on Product</u>: tests to analyze quality, nucleic acid or protein sequence, protein conformation, antibody reactivity, etc. of final vaccine product.

Vaccination: giving final produced vaccine to people.

Analysis of SAR		<b>2 (COVID-19) V</b> dated 10 November 2020	accine Ca	<u>ndidates</u>	DOES USE :	USE abortion-derived ce abortion-derived cell line	9
Sponsor(s) <sup>1</sup>	Country	Strategy <sup>2</sup>	Clinical Trial Status <sup>3</sup>	Public Funding <sup>4</sup>	SOME DO.  [BLANK] Currently  Design &  Development	undetermined Production	Confirm- atory Lab Tests
WHOLE VIRUS VACCING Beijing Institute of Biological Products/ Sinopharm	E – LIVE A	TENUATED or INACT Inactivated virus "BBIBP-CorV" Given: Intramuscular	Phase 3 Phase 1/2		Vero monkey cells	Vero monkey cells	



Wuhan Institute of Biological Products/ Sinopharm	China	Inactivated virus Unnamed Given: Intramuscular	Phase 3 Phase 1/2		Vero monkey cells	Vero monkey cells <u>Xia et al., JAMA</u> 324, 951,  13Aug2020	
John Paul II Medical Research Institute	USA	Live attenuated virus	Pre-clinical		Ethical cell lines as a matter of policy	Perinatal human cells (term umbilical cord and placental)	
Sinovac Biotech Co., Ltd.	China	Inactivated virus "PiCoVacc" Given: Intramuscular	Phase 3 Phase 3 Phase 1/2 Phase 1/2 Phase 1/2		Vero monkey cells	Vero monkey cells Gao et al., Science 369, 77, 3July2020	protein test HEK293 cells Supplement Gao et al., Science 369, 77, 3July2020
VIRAL VECTOR-BASED	VACCINE						
Altimmune	USA	Replication-deficient Adenovirus vector "AdCOVID" Given: Intranasal	Pre-clinical		PER.C6 cells	PER.C6 cells Same platform as NasoVAX NasoVAX uses PER.C6 Licensed PER.C6 from Janssen	•
AstraZeneca University of Oxford	USA UK	Replication-deficient Adenovirus vector "AZD1222" "ChAdOX1nCoV-19" Given: Intramuscular	Phase 3 Phase 3 Phase 3 Phase 2/3 Phase 2/3 Phase 1/2 Phase 1/2	Operation Warp Speed HHS-BARDA \$1.2 Billion CEPI up to \$384 Million	HEK293 cells	HEK293 cells van Doremalen et al., Nature preprint, 30July2020	•
CanSino Biologics, Inc. Beijing Institute of Biotechnology, Academy of Military Medical Sciences, PLA of China	China	Replication-deficient Adenovirus vector "Ad5-nCoV" Given: Intramuscular	Phase 3 Phase 3 Phase 2 Phase 2 Phase 2 Phase 1 Phase 1		HEK293 cells	HEK293 cells Biospace, 12May2020	•
Gamaleya Research Institute	Russia	Replication-deficient Adenovirus vectors	Phase 3		HEK293 cells	HEK293 cells	<b>•</b>



		(rAd26-S+rAd5-S) "Sputnik V" Given: Intramuscular	Early approval in Russia August 2020 Phase 1/2 Phase 1/2				
Institut Pasteur and Themis and Merck	USA France	Replication-competent recombinant measles virus "TMV-083" Given: Intramuscular	Phase 1	CEPI up to \$4.9 Million		Vero monkey cells	
Janssen Research & Development, Inc. Johnson & Johnson	USA	Replication-deficient Adenovirus vector "Ad26" Given: Intramuscular	Phase 3 Phase 1/2	Operation Warp Speed HHS-BARDA \$1,457,887,081 total	PER.C6 cells	PER.C6 cells Tostanoski et al., Nature Medicine, 3Sept2020; J&J, 30March2020; Janssen Vaccine Technologies	•
Merck and IAVI	USA	Replication-competent recombinant vesicular stomatitis virus (VSVΔG) "V590" Given: Intramuscular	Pre-clinical	Operation Warp Speed HHS-BARDA \$38,033,570	Vero monkey cells	Vero monkey cells <u>Use rVSV Ervebo</u> <u>platform</u> <u>Ervebo uses Vero</u> <u>cell culture-11</u> <u>Description</u>	
Shenzhen Geno-immune Medical Institute	China	Lentivirus minigenes + Adult human APC (antigen-presenting cells)	Phase 1				
Shenzhen Geno-immune Medical Institute	China	Lentivirus minigenes + Adult human CD/T cells (dendritic cells and T cells) "LV-SMENP-DC"	Phase 1/2				
Vaxart	USA	Replication-deficient Adenovirus vector "VXA-CoV2-1" plus dsRNA adjuvant Given: Oral	Phase 1		HEK293 cells	HEK293 cells Moore et al., bioRxiv 6Sept2020	•



Anhui Zhifei Longcom Biopharmaceutical/Institute of Microbiology, Chinese Academy of Sciences	China	Protein vaccine Recombinant RBD dimer plus adjuvant Given: Intramuscular	Phase 2 Phase 1/2 Phase 1		HEK293T cells Dai et al., Cell 6Aug2020	CHO hamster cells Dai et al., Cell 6Aug2020	Pseudovirus HEK293T cells Dai et al., Cell 6Aug2020
Clover Biopharmaceuticals, Inc.	China	Protein vaccine "SCB-2019" plus adjuvant CpG 1018 Given: Intramuscular	Phase 1	CEPI up to \$69.5 Million		CHO hamster cells Trimer-Tag system; Liu et al., Scientific Reports 2017	
John Paul II Medical Research Institute	USA	Recombinant Protein Perinatal human cells (term umbilical cord and placental)	Pre-clinical		Ethical cell lines as a matter of policy	Perinatal human cells (term umbilical cord and placental)	
Medicago	Canada	Protein on Virus-Like Particle "CoVLP" Plant-expressed spike protein particle with adjuvant, CpG1018 or AS03 Given: Intramuscular	Phase 1		Recombinant DNA sequence in Agrobacterium, transformation of plant cells	Plant expression of protein and VLP Ward et al., medRxiv 6Nov2020	Pseudovirus HEK293 cells Ward et al., medRxiv 6Nov2020
Novavax	USA	Protein vaccine "NVX-CoV2373" Baculovirus expression plus Matrix M adjuvant Given: Intramuscular	Phase 3 Phase 2 Phase 1	Operation Warp Speed HHS-BARDA \$1,600,434,523 CEPI up to \$388 Million		Sf9 insect cells Bangaru et al., bioRxiv preprint, 6Aug2020; Graphical view	Pseudovirus HEK293 cells Bangaru et al., bioRxiv preprint, 6Aug2020
Sanofi and GSK Protein Sciences	USA France	Protein vaccine Baculovirus expression plus AS03 adjuvant Given: Intramuscular	Phase 1/2	Operation Warp Speed HHS-BARDA \$2,072,775,336 total		Sf9 insect cells  Baculovirus  expressed  recombinant protein  ;	
Sorrento	USA	Protein vaccine "T-VIVA-19" SARS-Cov-2 spike protein S1 domain fused with human IgG-Fc Given: Intramuscular	Pre-clinical			CHO cells Herrmann et al., bioRxiv preprint, 30June2020	



Sorrento	USA	Protein vaccine "STI-6991" SARS-Cov-2 spike protein expressed on K562 cells	Pre-clinical			K562 cells Concept: Ji et al., Medicine in Drug Discovery March2020	
University of Pittsburgh	USA	Protein vaccine Adenovirus-expressed recombinant proteins "PittCoVacc" Given: Microneedle arrays	Pre-clinical		HEK293 cells	HEK293 cells Kim et al., EBioMedicine , 2April2020	•
University of Queensland and CSL Ltd.	Australia	Protein vaccine "V451" Recombinant protein with proprietary molecular clamp Given: Intramuscular	Phase 1 Phase 1 Phase 1	CEPI up to \$4.5 Million		expiCHO hamster cells	
RNA VACCINE							
Arcturus Therapeutics	USA	mRNA vaccine self-transcribing, replicating "LUNAR-CoV19" ("ARCT-021") in vitro transcription reaction with T7 RNA polymerase from STARR plasmid template LUNAR proprietary lipid nanoparticle encapsulated Given: Intramuscular	Phase 1/2		Sequence designed on computer	No cells used de Alwis et al., bioRxiv 3Sept2020	protein test de Alwis et al., bioRxiv 3Sept2020
CureVac	Germany	mRNA vaccine non-replicating "CVnCoV" in vitro transcription lipid nanoparticle encapsulated Given: Intramuscular	Phase 2 Phase 1	CEPI up to \$15.3 Million	Sequence designed on computer	No cells used Rauch et al., bioRxiv 23Oct2020	Protein test Reticulocyte lysate, HeLa cells Rauch et al., bioRxiv 23Oct2020



Moderna, Inc. with National Institutes of Health	USA	mRNA vaccine non-replicating "mRNA-1273" T7 RNA polymerase- mediated transcription from DNA plasmid template LNP (lipid nanoparticle) encapsulated Given: Intramuscular	Phase 3 Phase 2 Phase 1	Operation Warp Speed HHS-BARDA \$2,479,894,979 total CEPI up to \$1 Million	Sequence designed on computer	No cells used Corbett et al., Nature, 5Aug2020	protein test & pseudovirus HEK293 cells Corbett et al., Nature, 5Aug2020
Pfizer and BioNTech	USA Germany	mRNA vaccine non-replicating "BNT- 162a1,b1,b2,b3,c2" nucleoside-modified mRNA <i>in vitro</i> transcribed by T7 polymerase from a plasmid DNA template LNP (lipid nanoparticle) encapsulated Given: Intramuscular	Phase 2/3 Phase 1/2 Phase 1/2 Phase 1 Phase 1	Operation Warp Speed HHS-BARDA \$1.95 Billion	Sequence designed on computer	No cells used Vogel et al., bioRxiv 8Sept2020	protein test & pseudovirus HEK293 cells Vogel et al., bioRxiv 8Sept2020
Sanofi Pasteur and Translate Bio	USA France	mRNA vaccine non-replicating "MRT5500" synthesized by in vitro transcription employing RNA polymerase with a plasmid DNA template LNP (lipid nanoparticle) encapsulated Given: Intramuscular	Pre-clinical		Sequence designed on computer	No cells used Kalnin et al., bioRxiv 14Oct2020 mRNA production in the lab; Translate Bio scientific platform	protein test & pseudovirus HEK293 cells Kalnin et al., bioRxiv 14Oct2020
DNA VACCINE							
Genexine	Korea	DNA vaccine "GX-19" DNA synthesized in vitro, placed in plasmid vector	<u>Phase 1/2</u>		Sequence designed on computer	No cells used Seo et al., bioRxiv 100ct2020	



		Given: Intramuscular and Electroporation					
Inovio Pharmaceuticals	USA	DNA vaccine "INO-4800" DNA synthesized in vitro, placed in plasmid vector Given: Intradermal Electroporation	Phase 1/2 Phase 1	Operation Warp Speed CEPI up to \$22.5 Million	Sequence designed on computer	No cells used Smith et al., Nature 20May2020	protein test & pseudovirus HEK293 cells Smith et al., Nature 20May2020
Symvivo Corporation	Canada	DNA vaccine Genetically engineered Bifidobacterium longum "bacTRL-spike" Given: Oral, bacteria bind to gut lining	Phase 1			No cells used	

<sup>1.</sup> Data accumulated from primary literature as referenced in the Chart; <u>AND</u> "COVID-19 Treatment and Vaccine Tracker," Milken Institute, <a href="https://covid-19tracker.milkeninstitute.org/">https://covid-19tracker.milkeninstitute.org/</a>; <u>AND</u> "Draft landscape of COVID-19 candidate vaccines," World Health Organization (WHO), <a href="https://www.who.int/publications/m/item/draft-landscape-of-covid-19-candidate-vaccines">https://www.who.int/publications/m/item/draft-landscape-of-covid-19-candidate-vaccines</a>

NOTE that patents are <u>not</u> considered because they are unreliable sources; even the most relevant patents are prospective documents that provide examples of potential use, but do not provide information about actual, current application of an invention or technology.

- 2. Prentice, DA and Sander Lee, T. June 15, 2020. A Visual Aid to Viral Infection and Vaccine Production. *On Science Series 1. Accessed 19 June 2020* at: <a href="https://lozierinstitute.org/a-visual-aid-to-viral-infection-and-vaccine-production/">https://lozierinstitute.org/a-visual-aid-to-viral-infection-and-vaccine-production/</a>
- 3. Phases of Clinical Trials: Pre-clinical- laboratory and animal studies; Phase I- 10-100 people, study safety and dosage; Phase II- tens to hundreds of people, study efficacy, dosage, side effects; Phase III- hundreds to thousands of people, study efficacy and adverse reactions.
- 4. HHS-BARDA = U.S. Health and Human Services-Biomedical Advanced Research and Development Authority; CEPI = Coalition of Epidemic Preparedness Innovations; BARDA's rapidly-expanding COVID-19 medical countermeasure portfolio. *Accessed 29 Sept 2020* at

https://www.medicalcountermeasures.gov/app/barda/coronavirus/COVID19.aspx; CEPI's COVID-19 Vaccine Portfolio, Accessed 29 Sept 2020 at https://cepi.net/COVAX/