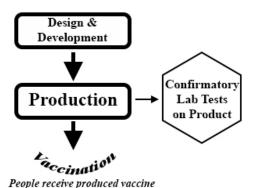


Update: COVID-19 Vaccine Candidates and Abortion-Derived Cell Lines

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.¹ 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 SARS-CoV-2 (COVID-19) Vaccine Candidates Last Updated 30 September 2020					DOES NOT USE abortion-derived cell line DOES USE abortion-derived cell line			
					SOME DO.	DO NOT use abortion-double undetermined	erived cells,	
Sponsor(s) ¹	Country	Strategy ²	Clinical Trial Status ³	Public Funding ⁴	Design & Development	Production	Confirm- atory Lab Tests	
WHOLE VIRUS VACCINI Beijing Institute of Biological	E – LIVE A'	Inactivated virus	Phase 3					
Products/ Sinopharm		"BBIBP-CorV" Given: Intramuscular	<u>Phase 1/2</u>		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	•



		(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	•



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)	
Novavax	USA	Protein vaccine "NVX-CoV2373" Baculovirus expression plus Matrix M adjuvant Given: Intramuscular	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 <u>Concept: Ji et al.,</u> <u>Medicine in Drug</u> <u>Discovery</u> 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"	Phase 1 Phase 1	CEPI up to \$4.5 Million			



		Recombinant protein with proprietary molecular clamp Given: Intramuscular	Phase 1			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 Given: Intramuscular	Phase 2 Phase 1	CEPI up to \$15.3 Million		No cells used	
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 <u>Corbett et al.</u> , <u>Nature</u> , <u>5Aug2020</u>	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	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



		polymerase from a plasmid DNA template LNP (lipid nanoparticle) encapsulated Given: Intramuscular					
Sanofi Pasteur and Translate Bio	USA France	mRNA vaccine non-replicating Given: Intramuscular	Pre-clinical		Sequence designed on computer	No cells used mRNA production in the lab; Translate Bio scientific platform	
DNA VACCINE							
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	

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

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: https://lozierinstitute.org/a-visual-aid-to-viral-infection-and-vaccine-production/
- 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/