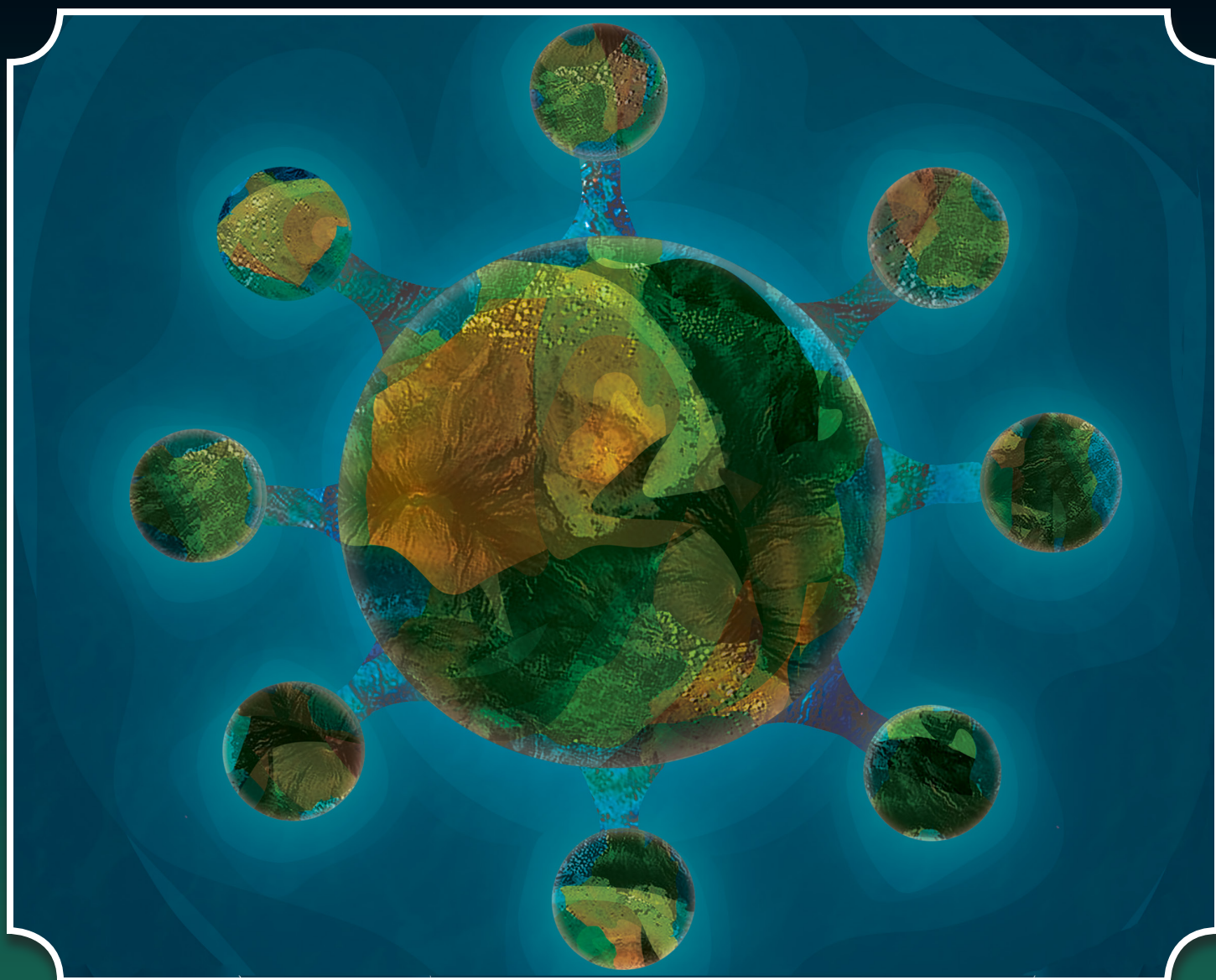


# VIRUS-BASED TOOLS

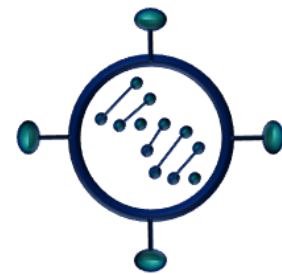
FOR DRUG DISCOVERY

Lentivirus | Vesicular Stomatitis Virus | Adeno-Associated Virus

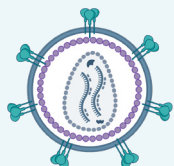


# Optimized Viral Options

Virus-based tools such as Lentivirus, Adeno-Associated Virus (AAV), and Vesicular Stomatitis Virus (VSV) are critical for cell engineering and the study of viral infection. We have designed a suite of ready-to-use viral reagents to address a wide span of research areas including virology (particularly Coronaviruses), immunotherapy, CAR-T therapy, CRISPR, cell signaling, and more.

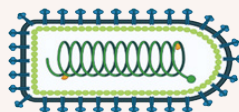


## Lentivirus



Ideal for pseudotyping or engineering stable cell lines, lentiviruses deliver relatively large genes that can integrate into the host genome.

## VSV



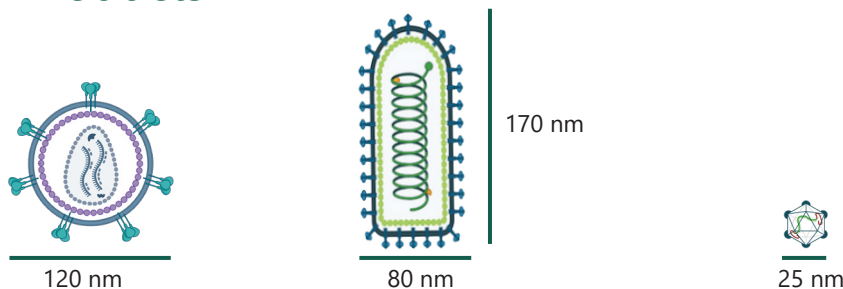
VSV is an excellent tool to model viral infection using pseudotyped viruses that replace the VSV-G protein with a desired viral protein of interest. Some cell infectivity models work best with VSV.

## AAV



AAV is an ideal viral vector for delivery into primary cells both *in vitro* and *in vivo*. Its low immunogenicity and pathogenicity enable safe gene therapy.

## Comparison of Viral Products



	HIV-based lentivirus	VSV Delta G	AAV
Genome size	9.7 kb	11 kb	4.7 kb
Suggested max insert size	10 kb	4.5 kb	2.5 kb
Genome type	ssRNA	ssRNA	ssDNA
Pseudotyping	Yes	Yes	No
Integration	Yes – stable (retrovirus)	No	No
Transduce exogenous gene of interest	Yes -stable	Yes - transient	Yes >6 months
Time to peak expression	72 hours	24-48 hours	7 days (2 weeks <i>in vivo</i> )
Biosafety level	2	2	1
<i>In vivo</i> use (animals)	Low efficiency	-	Most suitable
Immune response	Yes, medium	-	Ultra-low
Preferred applications	Gene transfer ( <i>in vitro</i> , stable)	Model viral infection	Gene transfer ( <i>in vitro</i> and <i>in vivo</i> )



# Our Advantages



## Produced In-House

- Made in the USA at our San Diego, CA laboratory
- Customized, personal support directly from our scientists



## Committed to Excellence

- ISO 9001:2015-certified Quality Management System
- Lot-specific quality control testing



## Expansive Portfolio

- Choose from over 140 ready-to-use lentivirus, AAV, and VSV vectors to study CAR-T, cell signaling pathways, coronavirus, CRISPR, and immunotherapy
- Long-term stable expression of a transgene with low immunogenicity, low toxicity, and high transduction efficiencies



## Custom Services

- Design a custom virus with reporters and selection markers of your choice
- Utilize our cell line development services to generate overexpression and reporter cell lines
- Generate knock-out/knock-in cell lines or integrating/non-integrating viruses

## Online Resources



### Lentivirus Tools Webinar

<https://bpsbioscience.com/videos?topic=lentiviruses>



### Lentivirus FAQs

<https://bpsbioscience.com/lentivirus-faq>



### Lentiviruses for SARS-CoV-2 Research Tech Note

<https://bpsbioscience.com/pseudoviruses-sars-cov-2-research>



### SARS-CoV-2 Pseudoviruses eBook

<https://bpsbioscience.com/ebooks?category=coronavirus>



# Lentivirus Products

Lentiviruses are enveloped retroviruses that fuse with the target cell membrane, delivering genetic material into the cytoplasm of the cell. Our replication-incompetent lentiviruses have been VSV-G pseudotyped, making these virus particles safe, stable and especially useful to target a wide range of cell types. For infectivity assays, we have developed lentiviral products pseudotyped with SARS-CoV-2 spike proteins, specific to variant mutations. Our suite of over 120 lentivirus products enables studies across a wide range of research areas.

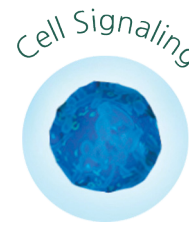
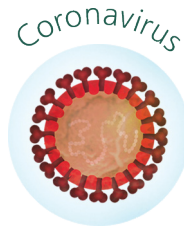
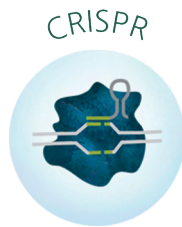
## Applications

- Stable cell line generation
- Protein expression
- CRISPR/Cas9 knockout
- Generating cellular reporter assays (GFP, luciferase)
- Screen for neutralizing antibodies

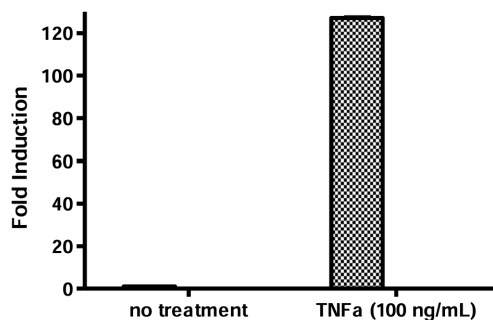
## Advantages

- Can infect actively dividing and non-dividing cells
- Can infect a wide range of cell stages
- Size of inserted DNA can be up to 10 kb
- Long term stable expression of a transgene
- Low cellular toxicity
- High transduction efficiency

## Research Areas

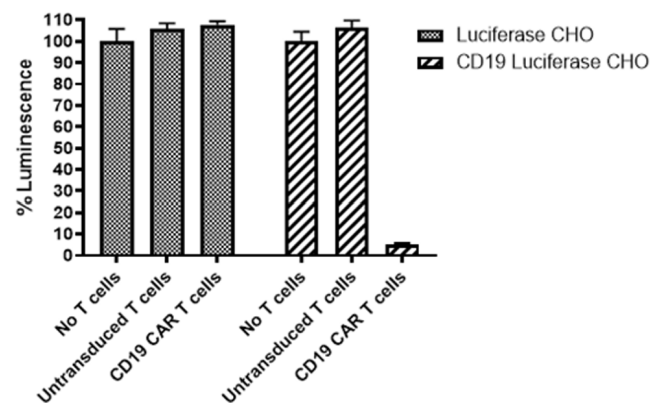


TNF $\alpha$  response of HEK293 cells transduced with NF- $\kappa$ B luciferase reporter lentivirus (#79564)



HEK293 cells transduced with NF- $\kappa$ B luciferase reporter lentivirus demonstrate induction of luciferase activity upon activation with TNF $\alpha$ . Fold induction was determined by comparing values against the control cells without TNF $\alpha$  treatment.

Activity of CD4/CD8 T cells transduced with anti-CD19 CAR Lentivirus (#78600)



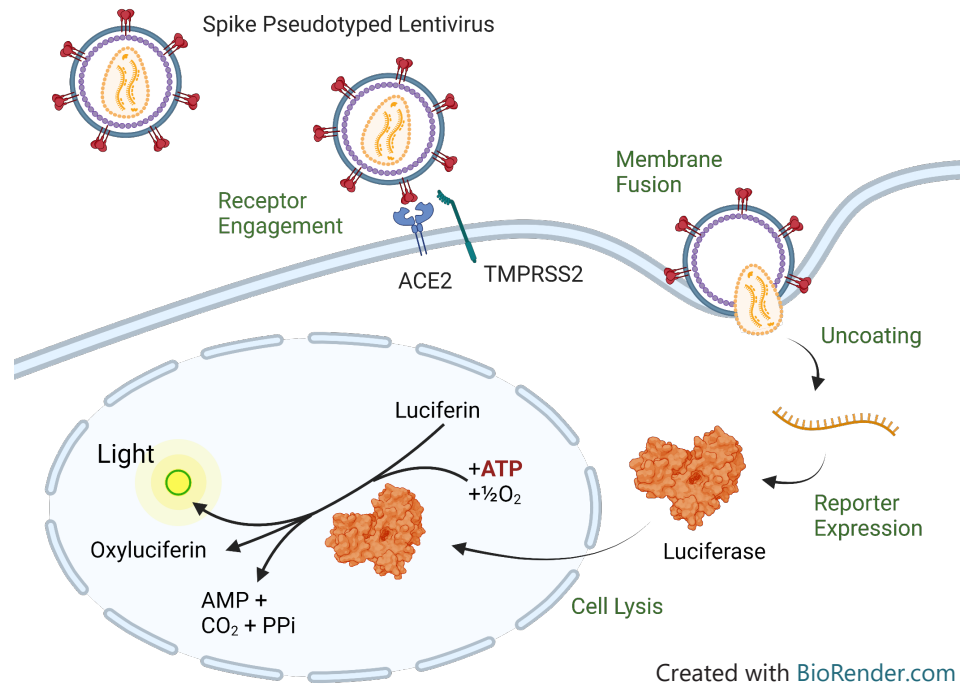
Anti-CD19 CAR Lentivirus-transduced T cells demonstrate specific killing of CD19/Luciferase CHO cells.



# Pseudoviruses for Modeling Infection

## Principle of the Assay

Lentivirus and VSV vectors can be pseudotyped, which involves replacing the native envelope protein with another viral protein of interest. For example, variant-specific SARS-CoV-2 Spike protein can be expressed on lentivirus or VSV delta G particles for infection of ACE2-expressing cells. The delivered genomes are engineered to express reporter genes such as luciferase or eGFP, enabling sensitive, quantitative readouts of infection. These systems serve as excellent models to screen for blocking antibodies or small molecule inhibitors of infection.



## Options for Optimal Experimentation

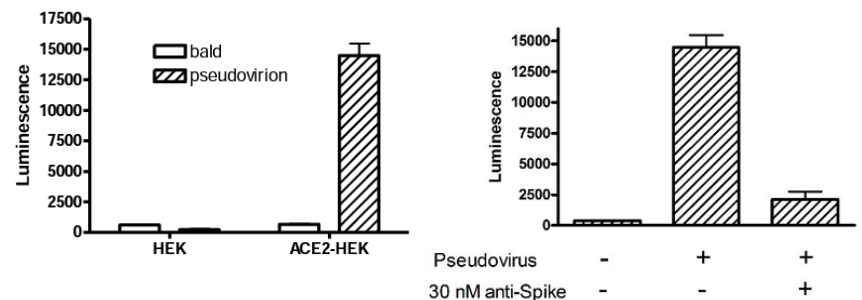
Virus Type	Reporters	Target Cell Types	Coronavirus Spike Variants
<ul style="list-style-type: none"> <li>Lentivirus</li> <li>VSV delta G (preferred for Vero E6 infection)</li> </ul>	<ul style="list-style-type: none"> <li>Luciferase</li> <li>eGFP</li> <li>Dual (Luc+eGFP)</li> </ul>	<ul style="list-style-type: none"> <li>HeLa (ACE2)</li> <li>CHO (ACE2)</li> <li>HEK293 (ACE2)</li> <li>Vero E6 (TMPRSS2)</li> </ul>	<ul style="list-style-type: none"> <li>SARS-CoV-2 emerging variants: BA.4/5, BA.2, BA.1</li> <li>Previous variants of interest: B.1.621, B.1.617.2, B.1.617.1, and many more.</li> </ul>

## Advantages

- High titer
- Simple protocols, suitable for high throughput assays
- Bald Lentivirus and VSV delta G controls
- Lentiviruses to express receptors: ACE2, TMPRSS2
- Quickly customizable to address emerging variant mutations or new viruses

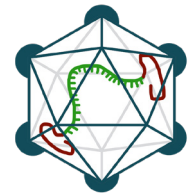
## Example Data

Spike (SARS-CoV-2) Pseudotyped Lentivirus (Luc-eGFP Dual Reporter) (#79982) transduction of ACE2-HEK293 cells monitored by luciferase activity



# AAV Gene Delivery and Reporter Vectors

Adeno-Associated Virus (AAV) is a small dependoparvovirus which was initially discovered as a contaminant in adenovirus preparations. AAVs are non-enveloped and consist of an icosahedral capsid containing a short, single-stranded DNA genome flanked by two Inverted Terminal Repeat sequences (ITRs).



Recombinant AAV used in gene therapy has been engineered to be integration-deficient and to deliver a gene of interest (up to  $\leq 5$  kb in length) in place of the viral genome. Inside the cell, the recombinant AAV vector exists as an episome and can result in sustained expression of the gene of interest for up to 6 months in non-dividing cells. Due to its low immunogenicity and lack of insertional mutagenesis, AAVs are safe for clinical use and are the vector of choice for many gene therapies currently in development.

## AAV Serotypes

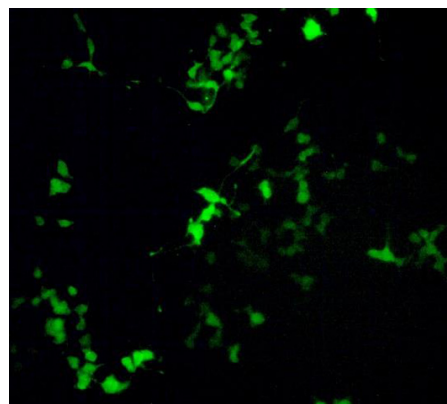
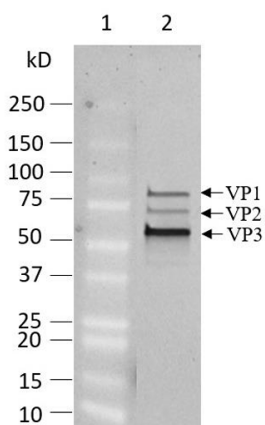
To date, 11 AAV serotypes have been characterized, each of these showing preferential binding for specific cell types and tissues. Thus, scientists can utilize this tropism to efficiently target specific cell types. In addition, several genetically engineered AAV serotypes have been developed to further increase tissue tropism and transduction efficiency for gene therapy purposes.

AAV1	CNS, Heart, Skeletal Muscle
AAV2	CNS, Kidney
AAV3	Liver
AAV4	CNS, Lung
AAV5	CNS, Lung
AAV6	Lung, Skeletal Muscle
AAV7	Liver, Skeletal Muscle
AAV8	CNS, Heart, Liver, Pancreas, Skeletal Muscle
AAV9	CNS, Heart, Liver, Lung, Skeletal Muscle

## AAV Reporter Particles

Reporter proteins, such as luciferase or fluorescent markers, are ideal to visualize and/or quantify protein expression following AAV transduction. Luciferase, eGFP, ZsGreen, and mCherry-containing AAVs can be used to optimize transduction and experimental conditions, track transgene expression over time, or be used as internal controls.

### Example data for AAV1 ZsGreen particles (#78443)



*Left:* Western blot of purified AAV1 ZsGreen particles display clear expression of AAV proteins: VP1, VP2, and VP3.

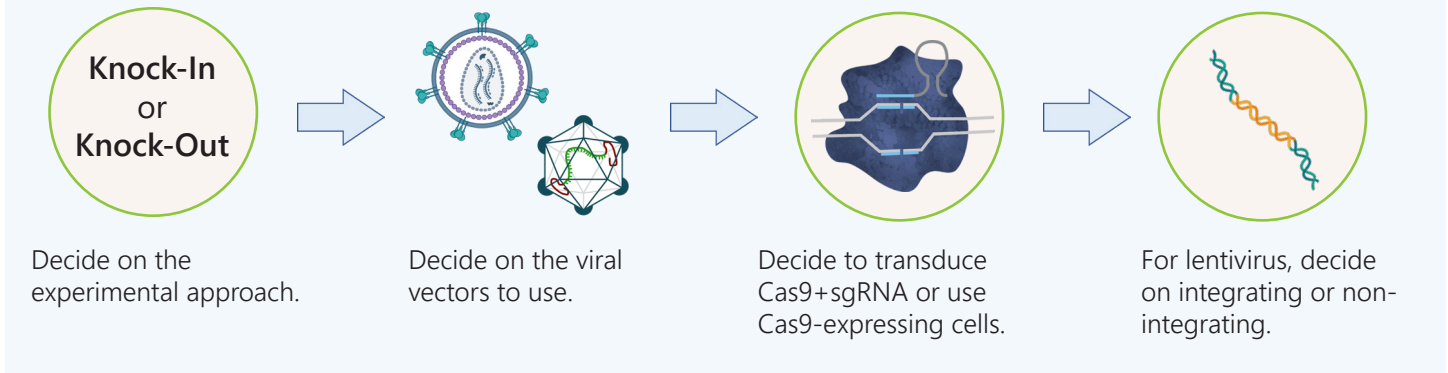
*Right:* Fluorescence microscopy of HEK293 cells 72 hours after transduction with AAV1 ZsGreen. ZsGreen expression was stable over time and still observed 30 days post transduction.



# CRISPR/Cas9 Cell Engineering

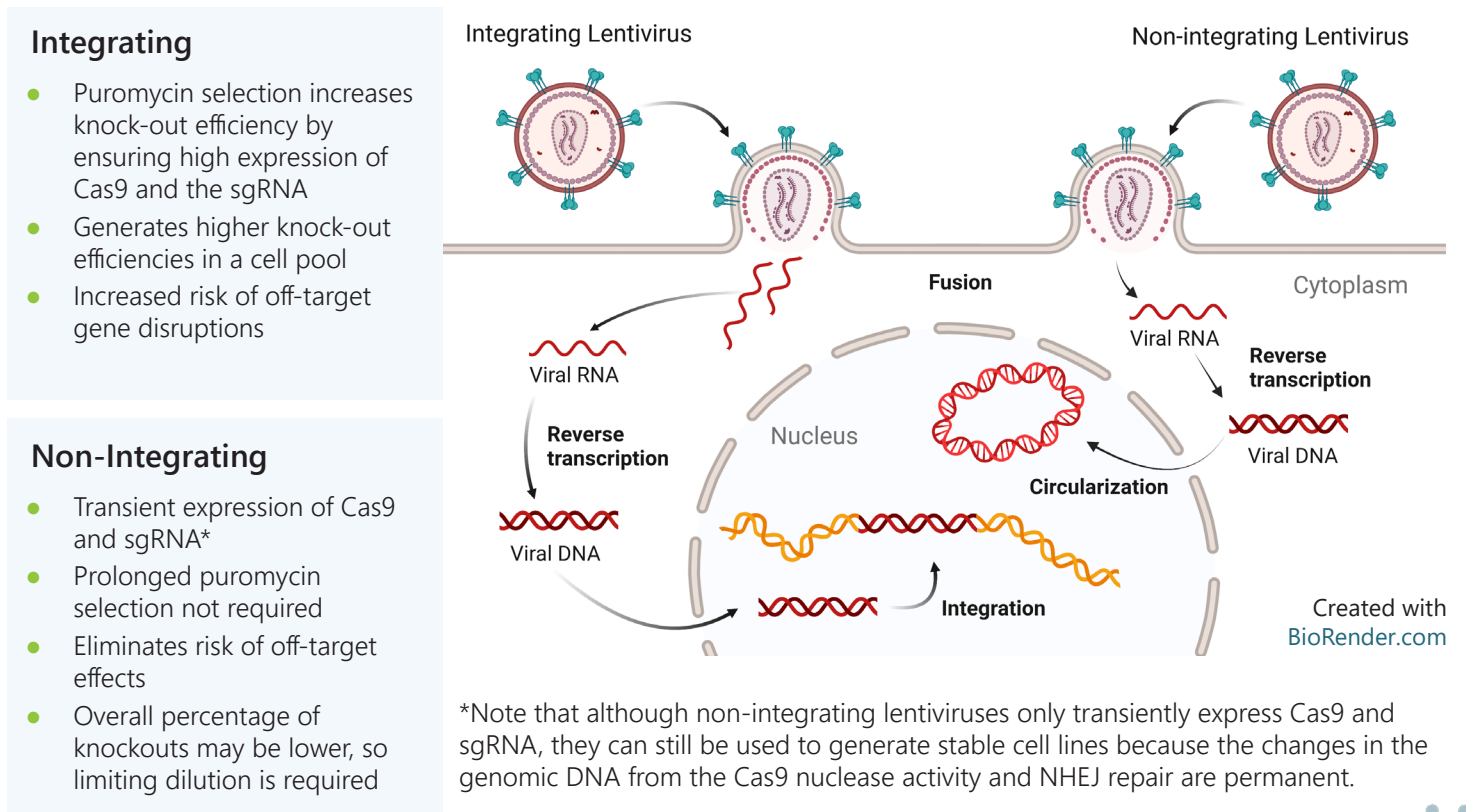
Lentivirus and AAV vectors can be used for CRISPR/Cas9-based cell engineering. Our off-the-shelf CRISPR lentiviruses are replication incompetent, HIV-based, VSV-G pseudotyped lentiviral particles that can transduce almost all types of mammalian cells, including primary and non-dividing cells. AAV can also be used to transduce primary cells, including *in vivo*, with SaCas9, derived from *Staphylococcus aureus*, which has high cutting efficiency in mammalian cells.

## The Logic Flow

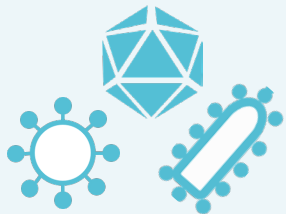


## Integrating vs Non-Integrating Lentiviruses

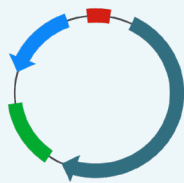
Lentiviruses are typically constructed to include the wild-type integrase enzyme that will integrate the Cas9 and sgRNA genes into the host genome. Alternatively, a non-active mutant integrase can be used, resulting in a non-integrating virus. Each has benefits and limitations which are compared below.



# Custom Virus Services



We can develop custom viruses for your research needs.



We can engineer your virus and cell lines with reporters, selection markers, variants, and specific mutations.



We can generate custom stable overexpression, knockout, or reporter cell lines using your virus.



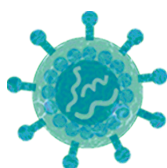
Choose integrating or non-integrating lentiviruses for cellular protein knock-out or knock-in.

## Our Milestone-Measured Process for Virus-Based Cell Engineering



**1**  
Molecular  
Biology

Viral vectors are generated using available clones, or through the use of synthetic DNA.



**2**  
Virus  
Production

The custom virus is manufactured for development of the stable cell line.



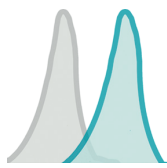
**3**  
Selection and  
Pool Generation

Parental cells are transduced with virus. The cell pool is selected for using antibiotics.



**4**  
Limiting Dilution  
and Clonal  
Selection

Based on the results of the initial pool testing, the cell pool is diluted and single cell-derived clones are selected.



**5**  
Confirmation of  
Expression

The expression level of the target protein is analyzed via Western blot or flow cytometry.



**6**  
Functional  
Validation

Cells are treated with a reference control compound to obtain dose-response titration data.



**7**  
Stability  
Testing

The desired number of clones are selected for passage stability testing. Mycoplasma testing and cell banking services are also available.

### Why choose BPS Bioscience for your custom projects?

- We have extensive expertise and experience in developing and manufacturing custom and off-the-shelf viral products.
- We have helped accelerate projects across large pharma, vaccine developers, biotech and basic research institutions.
- Our high quality custom products have returned excellent customer satisfaction scores.

*Give us a try today.*







Lentiviruses	Catalog#	Lentiviruses	Catalog#
BCMA Lentivirus	78714	EGR1 Promoter Luciferase Reporter Lentivirus	78664
Cas9 Lentivirus (Hygromycin Selection)	78067	Enhanced GFP Lentivirus (G418, Hygromycin and Puromycin)	78639
Cas9 Lentivirus (Neomycin Selection)	78432	EpCAM Lentivirus	78718
Cas9 Lentivirus (Puromycin Selection)	78066	Expression Negative Control Lentivirus (G418 or Hygromycin or Puromycin)	79902
CBL-B (Human) CRISPR/Cas9 Lentivirus (Integrating)	78343	FcER1G Lentivirus	79878
CBL-B (Human) CRISPR/Cas9 Lentivirus (Non-Integrating)	78344	FCGR2A CRISPR/Cas9 Lentivirus (Integrating)	78537
CD19 Lentivirus	78657	FCGR2A CRISPR/Cas9 Lentivirus (Non-Integrating)	78538
CD20 Lentivirus	78658	FcGRIIB (CD32B) Lentivirus	79877
CD22 Lentivirus	78659	FcGRIIIA (CD16a) Lentivirus	79876
CD47 CRISPR/Cas9 Lentivirus (Integrating)	78056	FcRL5 Lentivirus	78715
CD47 CRISPR/Cas9 Lentivirus (Non-Integrating)	78063	Firefly Luciferase Lentivirus (G418, Hygromycin and Puromycin)	79692
CD5 (Human) CRISPR/Cas9 Lentivirus (Integrating)	78119	Firefly Luciferase Lentivirus (UbC Promoter)	79880
CD5 (Human) CRISPR/Cas9 Lentivirus (Non-Integrating)	78198	Firefly Luciferase-eGFP Lentivirus (G418) or (Puromycin)	79980
CD8a Lentivirus	78648	GAL4 DBD-GR Lentivirus	78632
CD8a/CD8b Lentivirus	78650	GAS Luciferase Reporter Lentivirus (IFN- $\gamma$ /JAK/STAT1 Pathway)	78653
CEACAM5 Lentivirus	78719	GPC3 Lentivirus	78711
CEACAM6 Lentivirus	78720	GPRC5D Lentivirus	78716
CIITA (Human) CRISPR/Cas9 Lentivirus (Integrating)	78435	IL-2 Promoter Luciferase Reporter Lentivirus	79825
CIITA (Human) CRISPR/Cas9 Lentivirus (Non-integrating)	78434	IL-8 Promoter Luciferase Reporter Lentivirus	79827
Claudin-3 Lentivirus	78722	ISRE Luciferase Reporter Lentivirus (JAK/STAT Signaling Pathway)	79824
Claudin-4 Lentivirus	78723	Kinase (Human) CRISPR/Cas9 Lentivirus (Integrating)	78488
Claudin-9 Lentivirus	78721	LAG3 CRISPR/Cas9 Lentivirus (Integrating)	78053
CRBN CRISPR/Cas9 Lentivirus (Integrating)	78517	LAG3 CRISPR/Cas9 Lentivirus (Non-Integrating)	78060
CRBN CRISPR/Cas9 Lentivirus (Non-Integrating)	78518	LYPD1 Lentivirus	78724
CRE/CREB eGFP Reporter Lentivirus	78153	Myc Luciferase Reporter Lentivirus	78628
CRE/CREB Luciferase Reporter Lentivirus	79580	Nectin-4 Lentivirus	78712
CRISPR/Cas9 Kinase Knockout Lentivirus Library (Array Format)	78487	Negative Control eGFP Reporter Lentivirus	79927
CTLA4 CRISPR/Cas9 Lentivirus (Integrating)	78054	Negative Control Luciferase Lentivirus	79578
CTLA4 CRISPR/Cas9 Lentivirus (Non-Integrating)	78061	NF- $\kappa$ B eGFP Reporter Lentivirus	79926
eGFP Lentivirus (Inducible TET On)	78629	NF- $\kappa$ B Luciferase Reporter Lentivirus	79564

Lentiviruses	Catalog#	Lentiviruses	Catalog#
NFAT eGFP Reporter Lentivirus	79922	Spike (B.1.617.2; Delta Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78216
NFAT Luciferase Reporter Lentivirus	79579	Spike (B.1.617.2; Delta Variant) Pseudotyped Lentivirus (Luc Reporter)	78215
NFAT Luciferase-eGFP Reporter Lentivirus	78656	Spike (B.1.618 Variant) Pseudotyped Lentivirus (Luc Reporter)	78206
NFAT Luciferase-RFP Reporter Lentivirus	78617	Spike (B.1.621, Mu Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78618
NKp46 Lentivirus	78717	Spike (BA.1.1, Omicron Variant R346K) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78624
NLRP3 CRISPR/Cas9 Lentivirus (Integrating)	78545	Spike (BA.1.1, Omicron Variant R346K) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78623
NLRP3 CRISPR/Cas9 Lentivirus (Non-Integrating)	78546	Spike (BA.2, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78626
Non-secreted Gaussia Luciferase Lentivirus (CMV Promoter)	79893-C	Spike (BA.2, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78625
p53 Luciferase Reporter Lentivirus	78666	Spike (BA.2.12.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78646
PD-1 (Human) sgRNA-MS2 Lentivirus (Integrating)	78190	Spike (BA.2.12.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78645
PD-1 CRISPR/Cas9 Lentivirus (Integrating)	78052	Spike (BA.4/5, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78652
PD-1 CRISPR/Cas9 Lentivirus (Non-Integrating)	78059	Spike (BA.4/5, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78651
PD-L1 CRISPR/Cas9 Lentivirus (Integrating)	78057	Spike (BF.7, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78702
PD-L1 CRISPR/Cas9 Lentivirus (Non-Integrating)	78064	Spike (BF.7, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luciferase Reporter)	78699
PSMA Lentivirus	78726	Spike (BQ.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78700
Renilla Luciferase Lentivirus (G418 or Puromycin)	79565	Spike (BQ.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luciferase Reporter)	78697
RFP Lentivirus	78347-P	Spike (BQ.1.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78701
SBE Luciferase Reporter Lentivirus (TGFβ/SMAD Pathway)	79806	Spike (BQ.1.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luciferase Reporter)	78698
Secreted Gaussia Luciferase Lentivirus CMV Promoter or EF1A Promoter	79892	Spike (D614G) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78035
Spike (B.1.1.529 BA.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78349	Spike (D614G) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78028
Spike (B.1.1.529 BA.1, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78348	Spike (K417T, E484K, N501Y) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78143
Spike (B.1.1.7, Alpha Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78158	Spike (P.1, Gamma Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78159
Spike (B.1.1.7, Alpha Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78112	Spike (P.1, Gamma Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78144
Spike (B.1.351, Beta Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78160	Spike (SARS-CoV-1) Pseudotyped Lentivirus (eGFP Reporter)	78633
Spike (B.1.351, Beta Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luc Reporter)	78142	Spike (SARS-CoV-1) Pseudotyped Lentivirus (Luc Reporter)	78614
Spike (B.1.429, Epsilon Variant) Pseudotyped Lentivirus (Luc Reporter)	78172	Spike (SARS-CoV-2) Lentivirus	78010
Spike (B.1.617 Variant) Pseudotyped Lentivirus (Luc Reporter)	78204	Spike (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	79981
Spike (B.1.617.1, Kappa Variant) Pseudotyped Lentivirus (Luc Reporter)	78205	Spike (SARS-CoV-2) Pseudotyped Lentivirus (Luciferase Reporter)	79942
Spike (B.1.617.2.1; Delta Plus Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78219	Spike (XBB.1.5, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (eGFP Reporter)	78737
Spike (B.1.617.2.1; Delta Plus Variant) Pseudotyped Lentivirus (Luc Reporter)	78218	Spike (XBB.1.5, Omicron Variant) (SARS-CoV-2) Pseudotyped Lentivirus (Luciferase Reporter)	78736

Lentiviruses	Catalog#
Spike Variants (SARS-CoV-2) Pseudotyped Lentivirus Pack (Luciferase Reporter)	78616
Spike(SARS-CoV-2) Pseudotyped Lentivirus (Luc-eGFP Dual Reporter)	79982
SRE Luciferase Reporter Lentivirus	78627
STAT3 eGFP Reporter Lentivirus	78197
STAT3 Luciferase Reporter Lentivirus	79744
STAT5 Luciferase Reporter Lentivirus	79745
TCF/LEF Luciferase Reporter Lentivirus (Wnt/ $\beta$ -catenin Signaling Pathway)	79787
TCR Activator Lentivirus (CMV Promoter/Puromycin) or (EF1A Promoter/Puromycin) or (EF1A Promoter/Hygromycin)	79894
TCR CRISPR/Cas9 Lentivirus (Integrating)	78055
TCR CRISPR/Cas9 Lentivirus (Non-Integrating)	78062
TEAD Luciferase Reporter Lentivirus	79833
TGFBR2 CRISPR/Cas9 Lentivirus (Integrating)	78535
TGFBR2 CRISPR/Cas9 Lentivirus (Non-Integrating)	78536
TIGIT CRISPR/Cas9 Lentivirus (Integrating)	78058
TIGIT CRISPR/Cas9 Lentivirus (Non-Integrating)	78065
TMPRSS2 Lentivirus	78011
Trop2 Lentivirus	78710
UAS Luciferase Reporter Lentivirus	78631
YFP (Topaz) Lentivirus	79989

VSV	Catalog#
Bald VSV Delta G (Luciferase Reporter)	78636
Spike (B.1.617.2, Delta Variant) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78640
Spike (BA.1.1, Omicron Variant) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78641
Spike (BA.2, Omicron Variant) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78635
Spike (BA.2.12.1, Omicron Variant) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78643
Spike (BA.4/5, Omicron Variant) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78644
Spike (D614G) (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78642
Spike (SARS-CoV-2) Pseudotyped VSV Delta G (Luciferase Reporter)	78637
VSV-G Pseudotyped VSV Delta G (Luciferase Reporter)	78634



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