

# CRISPR-Cas9

Screening | Knock-out | Knock-in | Cell Line Development

INNOVATIVE PRODUCTS & SERVICES  
TO ADVANCE GENE EDITING  
RESEARCH



# BPS Bioscience Advantages

## Scientist Founded, Scientist Driven



### Produced In-house



- All CRISPR products and services are produced and performed in the USA at our San Diego, California laboratory
- Get customized, personal support directly from our CRISPR experts

### Multiple CRISPR Editing Tools and Applications



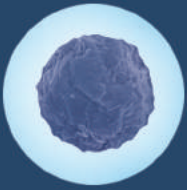
- CRISPR Knock-out
- CRISPR Knock-in
- CRISPR activation
- Cell lines, cell pools, lentiviruses, & plasmids
- CRISPR screens

### Customized For Your Research Needs



- Screening & Profiling: >200 optimized cell lines and cell-based assays
- Cell Line Development: Choose from >70 cell types and >20 reporter genes
- Ready to use Lentiviruses: Integrating and non-integrating options

# Knock-in Cell Lines



Introduce a specific point mutation or add a tag to your endogenous gene

## Project Milestones

1

### Molecular Biology



BPS will design and construct the gRNA and HDR template according to your experimental needs.

2

### Stable Cell Line Generation



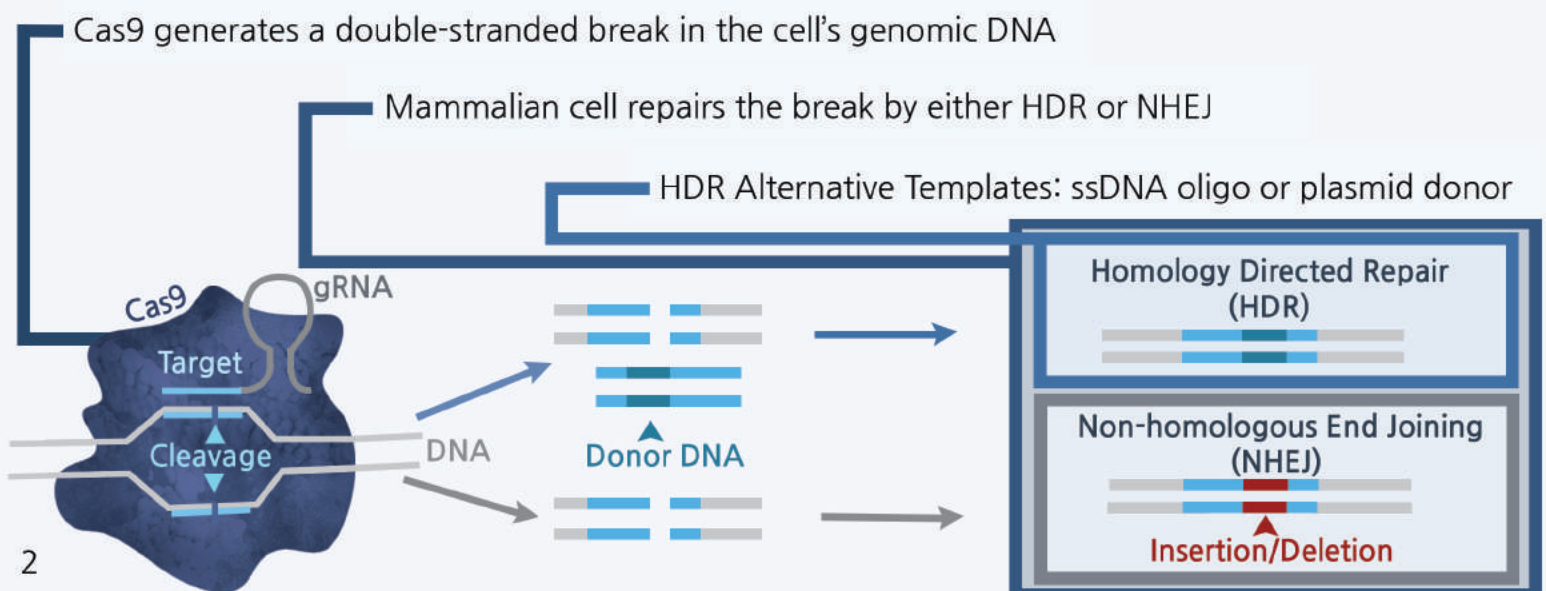
The cells will be transduced with the Cas9, gRNA, and HDR template, followed by genome editing evaluation. Single clones will be selected and expanded.

3

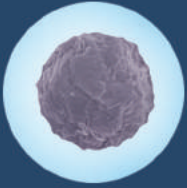
### Genotyping & Confirmation



The knock-in mutations will be confirmed by genomic sequencing, and positive clones will be expanded for further confirmation.



# Knock-out Cell Lines



- Knock-out your gene of interest for mechanistic or screening studies
- >20 available reporters
- >70 available cell types

## Project Milestones

1

### Molecular Biology



BPS will synthesize up to 5 sgRNA sequences and clone into a CRISPR expression vector for knock-out cell lines. BPS can also design the HDR template for knock-in cell lines.

2

### CRISPR Transfection



Depending on the cell type, cells can be transduced via electroporation, liposome-based transfection, or viral infection.

3

### Clonal Dilution



Based upon the results of the initial pool testing, the cell pool will be clonally diluted and the single cell-derived clones will be expanded.

4

### Confirmation of Expression



The expression level of the gene of interest will be analyzed via Western Blot or FACS.

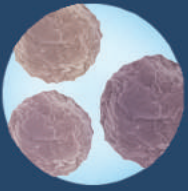
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### Confirmation & Delivery



Knock-out of the gene of interest will be confirmed by genomic sequencing. Confirmed clones will be expanded, frozen, and tested for Mycoplasma contamination.

# Knock-out Cell Pools



- Off the shelf products constitutively expressing Cas9 in variety of cell types: Jurkat, Neuro2A, A549, Raji, MDA-MB-231, HCT116, & more
- Cost-effective platform for setting up your own knock-out experiments or screens
- Custom services available for knock-out cell pool generation

## Project Milestones

1

### CRISPR Transfection



CRISPR transfection to express the Cas9

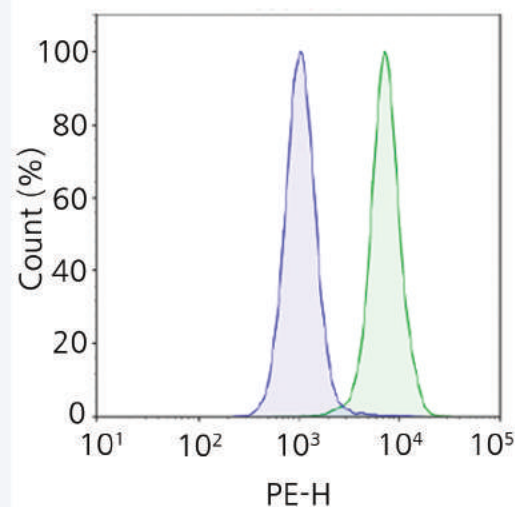
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### Cell Pool Testing

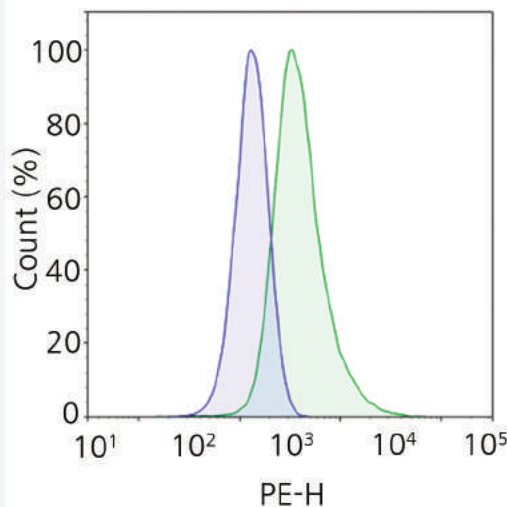


Cell pool testing to confirm presence of Cas9

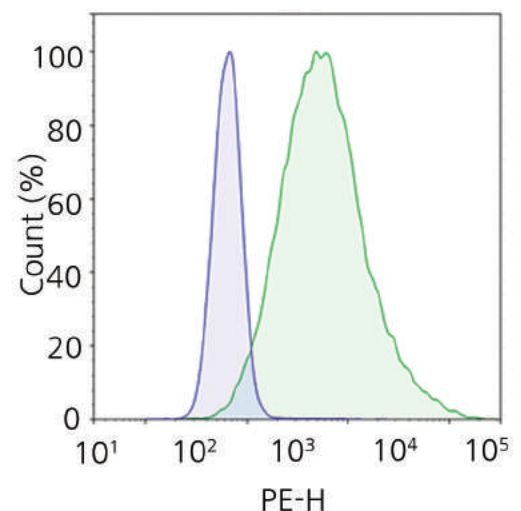
Expression of Cas9  
in Neuro2A Cell Pool



Expression of Cas9  
in Jurkat Cell Pool



Expression of Cas9  
in A549 Cell Pool



# CRISPR Activation



- Induce transcriptional activation and expression of any gene of interest
- Induction can be more than a hundred-fold, depending on the gene

## Synergistic Activation Mediator System (SAM) Components

sgRNA MS2

MS2-tagged sgRNA

dCas9 VP64

Mutated dCas9, lacking any endonuclease activity, fused to VP64, a transcriptional activator

MS2 p65 HSF1

P65 (Transcription Factor p65 or Nuclear Factor NF- $\kappa$ B p65), and HSF1 (Heat Shock Factor 1) fused with an MS2 tag

## SAM Activation Process

sgRNA MS2

Promoter Genomic DNA

1

sgRNA-MS2 targets the promoter region of the gene of interest

dCas9 VP64 MS2 p65 HSF1

sgRNA MS2

Promoter Genomic DNA

2

dCas9-VP64 and MS2-P65-HSF1 are recruited to the site in the genomic DNA

dCas9 VP64 MS2 p65 HSF1

sgRNA MS2

Promoter Genomic DNA

3

Transcription begins, inducing expression of the desired gene

## Gene Activation with CRISPRa

CRISPRa (SAM) Jurkat cells were electroporated with sgRNA-MS2 targeting PD-1 to induce PD-1 expression. Cells were stained with PE-labeled anti-PD-1 antibody and analyzed by FACS.

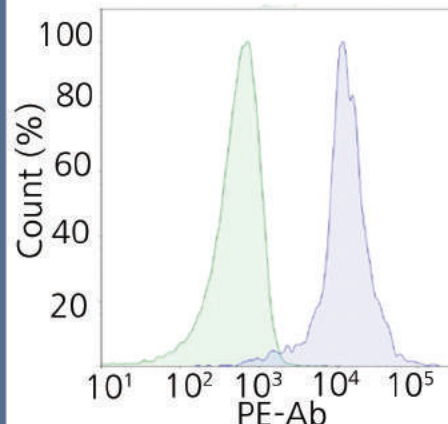


Parental CRISPRa Jurkat Cells



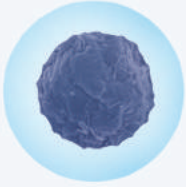
Transfected CRISPRa Jurkat Cells

Induction of CRISPRa Jurkat Cells



# CRISPRa Products & Services

## Cell Lines



- Stably express **dCas9 VP64** and **MS2 P65 HSF1**
- Transduce with **sgRNA MS2** via lentiviral transduction or electroporation
- Convenient and cost-effective platform for setting up your own activating experiments or screens without the need to first select for single cell clones expressing dCas9-VP64 and MS2-P65-HSF1

## Lentiviruses



### Integrating **dCas9 VP64** and **MS2 P65 HSF1** Lentiviruses

- Ready to be transduced into almost all types of mammalian cells, including primary and non-dividing cells
- These particles contain the genes for dCas9-VP64 (with blasticidin resistance) and MS2-P65-HSF1 (with hygromycin resistance)

### Integrating **sgRNA MS2** Activating Lentivirus

- Contain 4 validated sgRNA (single guide RNA) targeting the promoter region of your gene of interest, fused to MS2 and driven by a U6 promoter
- Ready-to-transduce into your dCas9-VP64 and P65-HSF1-MS2 expressing cell lines to stably activate expression of your gene of interest

## Plasmids



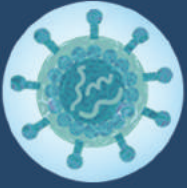
- **sgRNA MS2** plasmids can be transfected or electroporated into your cells for either transient or stable (following drug selection) activation
- Virus-free option, depending on your laboratory environment or preferences

## Custom Services



- With BPS Bioscience's cell line development services, our team of highly experienced scientists generate custom CRISPR activation cell lines
- Customizable project milestones and deliverables

# Lentiviruses



- Ready to transduce, no other components are required
- The surest way to quickly generate a knock-out cell line or cell pool
- Knock-out efficiencies as high as 90%

## Versatile & Ready to Use

The CRISPR Lentiviruses are replication incompetent, HIV-based VSV-G pseudotyped lentiviral particles that are ready to be transduced into almost all types of mammalian cells, including primary and non-dividing cells. These particles contain a CRISPR/Cas9 gene driven by an EF1a promoter, along with 4 validated sgRNA (single guide RNA) targeting your gene of interest, driven by a U6 promoter.

### Integrating Lentiviruses

Contains the wild-type integrase

Integrates randomly into the cell's genome to stably express both the Cas9 and sgRNA

Puromycin selection increases the knock-out efficiency by ensuring high expression levels of both Cas9 and the sgRNA

Generates higher knock-out efficiencies in a cell pool

Has a potentially higher risk of off-targeting due to random integrations into other genes

### Non-integrating Lentiviruses

Integrase-deficient

Generates only transient expression of the Cas9 and sgRNA

Prolonged puromycin selection is not required

Limited dilution is required because the overall percentage of knock-outs may be lower

Eliminates risk of off-targeting due to random viral integrations

## Cell Line Development Services

Conduct your research directly with a BPS lentivirus product, or choose to utilize our custom services to develop a cell line designed to meet your research needs. Our CRISPR experts will provide project guidance in addition to your custom lentivirus-generated knock-out cell line.

# Screening Services



- CRISPR kinase knock-out library available as an array or pool
- Customized specifically for your research project

## Knock-out Library

- Expansive CRISPR knockout library targeting human kinases

## Knock-out Array

- Integrating Lentiviruses are shipped ready-to-transduce into almost all types of mammalian cells, including primary and non-dividing cells
- Each format includes 4 sgRNA per gene, in addition to the proper controls
- Array format comes ready-to-use, delivered as one gene-per-well, without the need for any high-throughput screening platforms or bioinformatic analysis

## Knock-out Pool

- Our pooled format is delivered at high titer and customized for your screening capabilities
- Each format includes 4 sgRNA per gene, in addition to the proper controls

## Custom Services

- BPS can generate CRISPR libraries targeting your genes of interest
- Data is provided after completion of each milestone
- Our scientists work closely with you to provide customized tools to meet your research objectives

# CRISPR Cell Lines & Cell Pools, Proteins, Plasmids, Lentiviruses

## Product Listing

CRISPR Cell Lines & Cell Pools	Catalog#	Size	Plasmids	Catalog#	Size
Cas9 Expressing A549 Cell Pool	78072	2 vials	PD-1 sgRNA-MS2 for CRISPRa (Plasmid)	78091	5 µg
Cas9 Expressing Daudi Cell Pool	78089	2 vials			
Cas9 Expressing HCT116 Cell Pool	78073	2 vials			
Cas9 Expressing Jurkat Cell Pool	78070	2 vials			
Cas9 Expressing MDA-MB-231 Cell Pool	78069	2 vials			
Cas9 Expressing Raji Cell Pool	78071	2 vials			
Cas9-Expressing A549 Cell Line (High Expression or Low Expression)	78134	2 vials			
Cas9-Expressing Daudi Cell Line	78157	2 vials			
Cas9-Expressing HCT116 Cell Line (High or Low Expression)	78135	2 vials			
Cas9-Expressing HEK293 Cell Line	78166	2 vials			
Cas9-Expressing HeLa Cell Pool	78161	2 vials			
Cas9-Expressing Jurkat Cell Line (High or Low Expression)	78136	2 vials			
Cas9-Expressing MCF7 Cell Pool	78179	2 vials			
Cas9-Expressing MDA-MB-231 Cell Line (High or Low Expression)	78150	2 vials			
Cas9-Expressing Neuro2A Cell Line (High or Low Expression)	78137	2 vials			
Cas9-Expressing Neuro2A Cell Pool	78087	2 vials			
Cas9-Expressing Raji Cell Line	78156	2 vials			
CRISPRa (SAM) Jurkat Cell Line	78080	2 vials			
TCR Knockout NFAT-Luciferase Reporter Jurkat Recombinant Cell Line	79887	2 vials			
Proteins	Catalog#	Size	Lentiviruses	Catalog#	Size
Cas9, His-tag ( <i>S. pyogenes</i> )	100206	50 µg 100 µg 500 µg	Cas9 Lentivirus (Hygromycin Selection)	78067	500 µl x 2
			Cas9 Lentivirus (Puromycin Selection)	78066	500 µl x 2
			CD47 CRISPR/Cas9 Lentivirus (Integrating)	78056	500 µl x 2
			CD47 CRISPR/Cas9 Lentivirus (Non-Integrating)	78063	500 µl x 2
			CTLA4 CRISPR/Cas9 Lentivirus (Integrating)	78054	500 µl x 2
			CTLA4 CRISPR/Cas9 Lentivirus (Non-Integrating)	78061	500 µl x 2
			LAG3 CRISPR/Cas9 Lentivirus (Integrating)	78053	500 µl x 2
			LAG3 CRISPR/Cas9 Lentivirus (Non-Integrating)	78060	500 µl x 2
			PD-1 CRISPR/Cas9 Lentivirus (Integrating)	78052	500 µl x 2
			PD-1 CRISPR/Cas9 Lentivirus (Non-Integrating)	78059	500 µl x 2
			PD-L1 CRISPR/Cas9 Lentivirus (Integrating)	78057	500 µl x 2
			PD-L1 CRISPR/Cas9 Lentivirus (Non-Integrating)	78064	500 µl x 2
			TCR CRISPR/Cas9 Lentivirus (Integrating)	78055	500 µl x 2
			TCR CRISPR/Cas9 Lentivirus (Non-Integrating)	78062	500 µl x 2
			TIGIT CRISPR/Cas9 Lentivirus (Integrating)	78058	500 µl x 2
			TIGIT CRISPR/Cas9 Lentivirus (Non-Integrating)	78065	500 µl x 2



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