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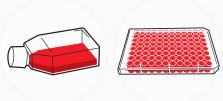
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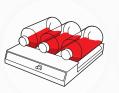
A Check-up Before you Scale up

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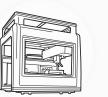
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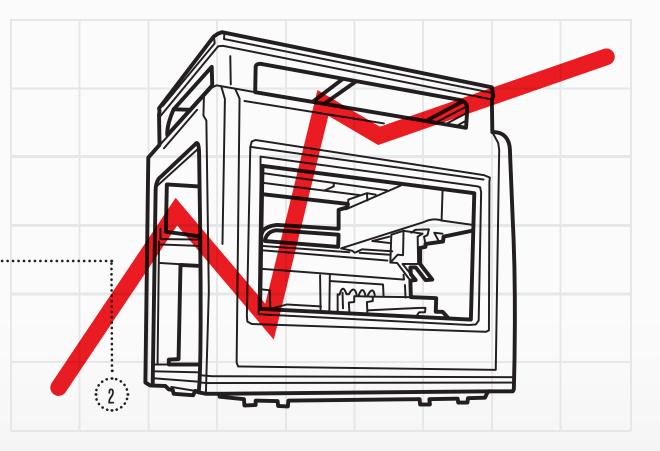






Cell-Line Engineering: A Check-up Before you Scale up

Bioproduction is a booming industry, with more therapeutics being produced in cell culture than ever before. With increased workflow flexibility, decreased variability, and high product yields, cell-line engineering is here to stay. But before you take your bioproduction to the next level, you'll need to know a bit more about your product, your cells, and their health. It's time for a cell-line engineering check-up!



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Transfection

There are several transfection methods available, including electroporation, lipofection, calcium phosphate, polymerbased methods, and viral transduction. Choose one that is optimized for your cell line, vector size, and throughput. Screen for successful, stable transfectants before moving on to optimize your media and culture conditions.

Selection

Select for cells that have taken up the plasmid by applying selective ressure, either an antibiotic or medium lacking an essential amino acid. Cells that survive will have incorporated both the gene of interest and the selection marker.

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Cell Viability Cell-viability analysis is essential for bioproduction workflows, since dead or dying cells don't only stop producing, but they may introduce proteolytic breakdown products into the culture. For optimal cellviability analysis during bioproduction, trypan blue dye exclusion

is the gold standard method, although other methods, like flow

cytometry, can be used to determine the type of cell death.

cell death Cell death is not a one-size-fits-all rocess. There are several different aths to cell death, including necrosis poptosis, mitotic catastrophe, or failur of autophagy. Knowing which is impacting

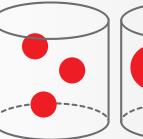
your culture can help you with preventin

the same fate in future batches.

Documenting

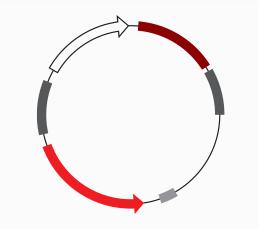
Cell Volume

The volume of a cell changes throughout protein expression, so cell-volume analysis can provide information on transfection efficiency. One of the most accepted methods to measure cell volume is using the Coulter Principle.



Expression Vector

For optimal expression of your product, choose an expression vector with proven success in your chosen cell line. Ensure that you include the genes encoding your product, strong promoters and enhancers, selection marker(s), and the required equipment for plasmid replication in your cell line of choice.



Which one is right for my product?

Scale up involves increasing the batch size of your culture, with the end result of increasing the amount of product derived from each batch. Mass-produced drugs are scaled up.



Scale out refers to the process of production on a patient-bypatient basis. Scale out often refers to the bioproduction of autologous cells for cell therapy.













Cell Health

Bioproduction can be stressful for cells, so monitoring cell health throughout the process can help prevent unexpected protein modifications or off-target products. Two common ways to monitor cell health during bioproduction are:

Morphology tracking: Follows changes in cell phenotype throughout bioproduction

Electrochemical tracking:

Measures the extracellular changes

created by cell activity





Dissolved 0₂

• Dissolved CO₂

Influences

What factors can influence cell

health during bioproduction?

Temperature

Agitation rate

 Other cellular functions





