



BST

Microbial Metabolism

Production and Regulation of Primary and Secondary Metabolites

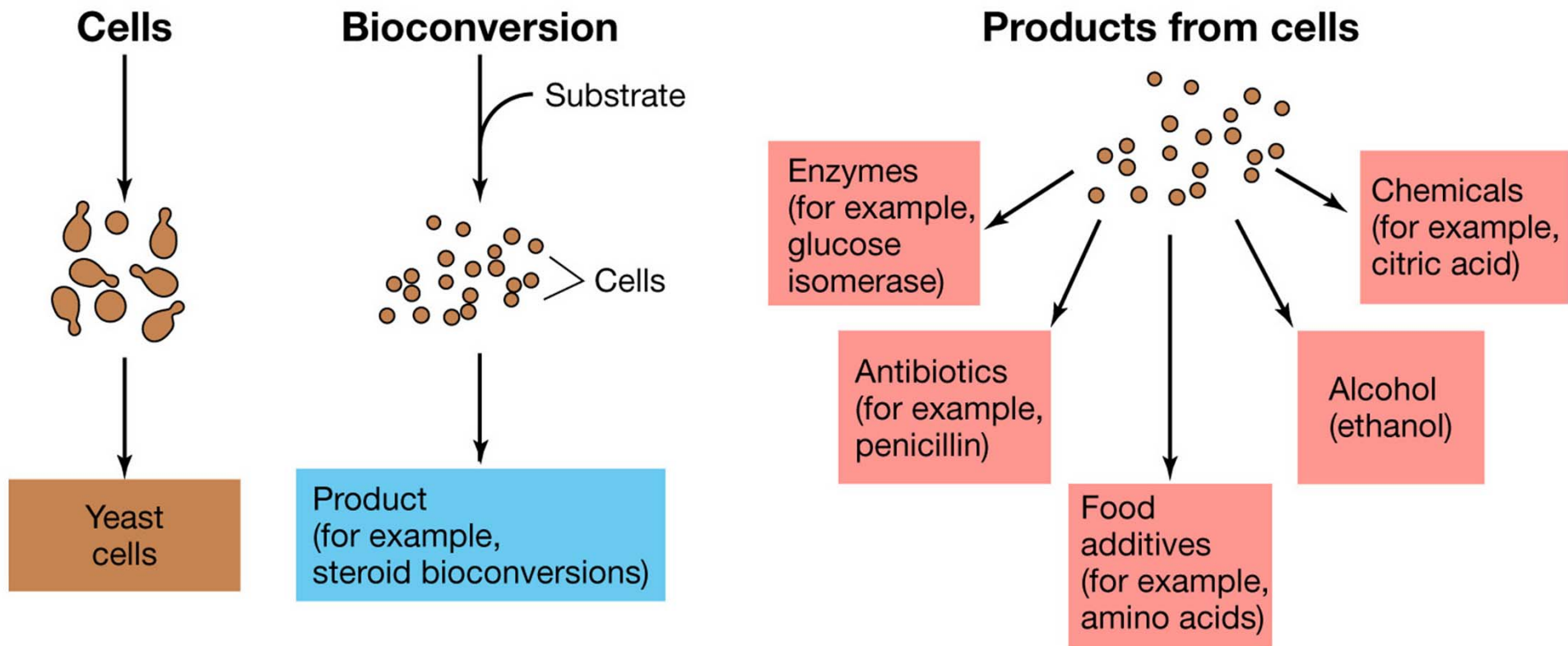
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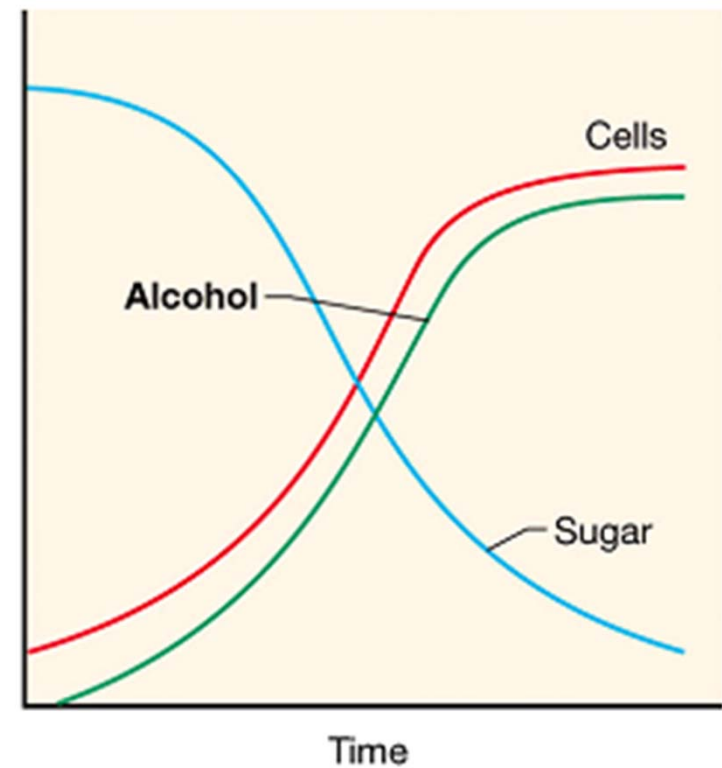
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Products of Microbiology



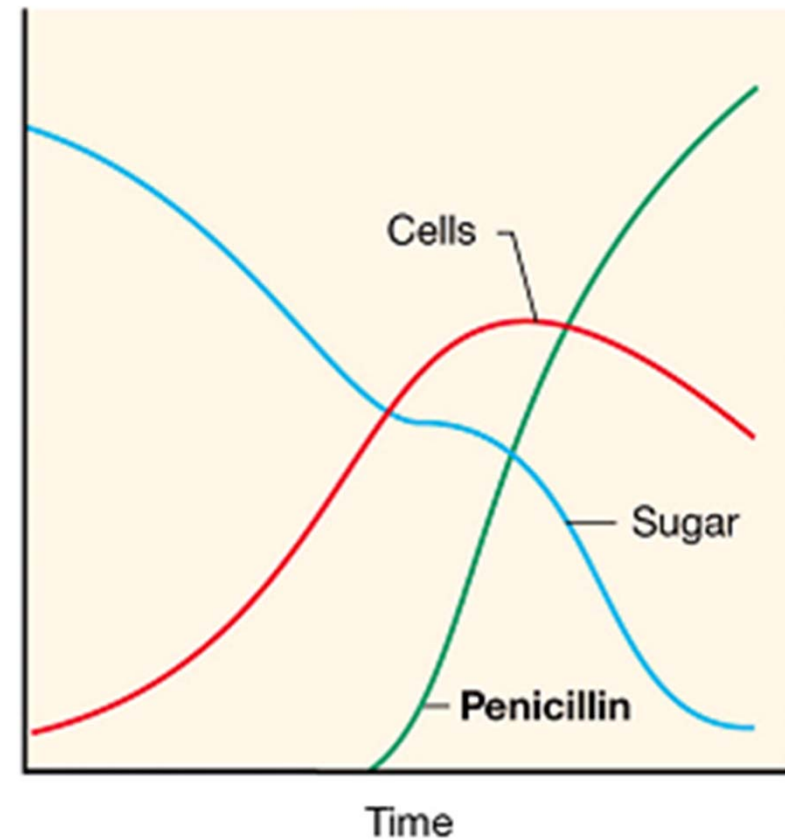
Primary Metabolites

- ➔ Small molecules of living cells
- ➔ Intermediates or end products of the pathway
- ➔ **Related to synthesis** of microbial cells in the growth phase
- ➔ Include alcohols, amino acids, nucleotides, organic acids, polyols, vitamins, and enzymes



Secondary Metabolites

- Accumulate following active growth
- Have **no direct relationship to synthesis** of cell material and natural growth
- Include antibiotics and toxins



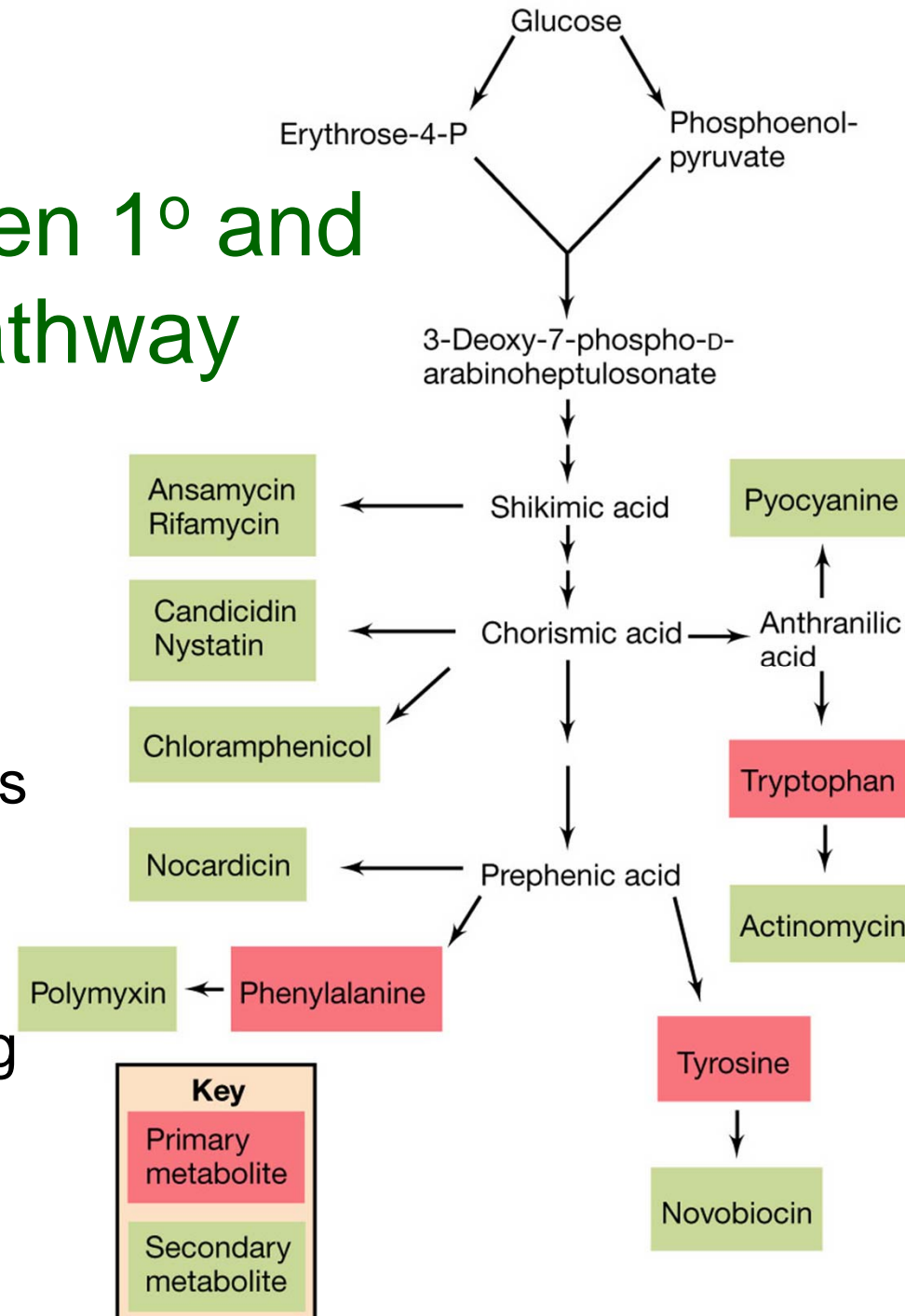
Relation between 1° and 2° metabolic pathway

Primary metabolic pathway

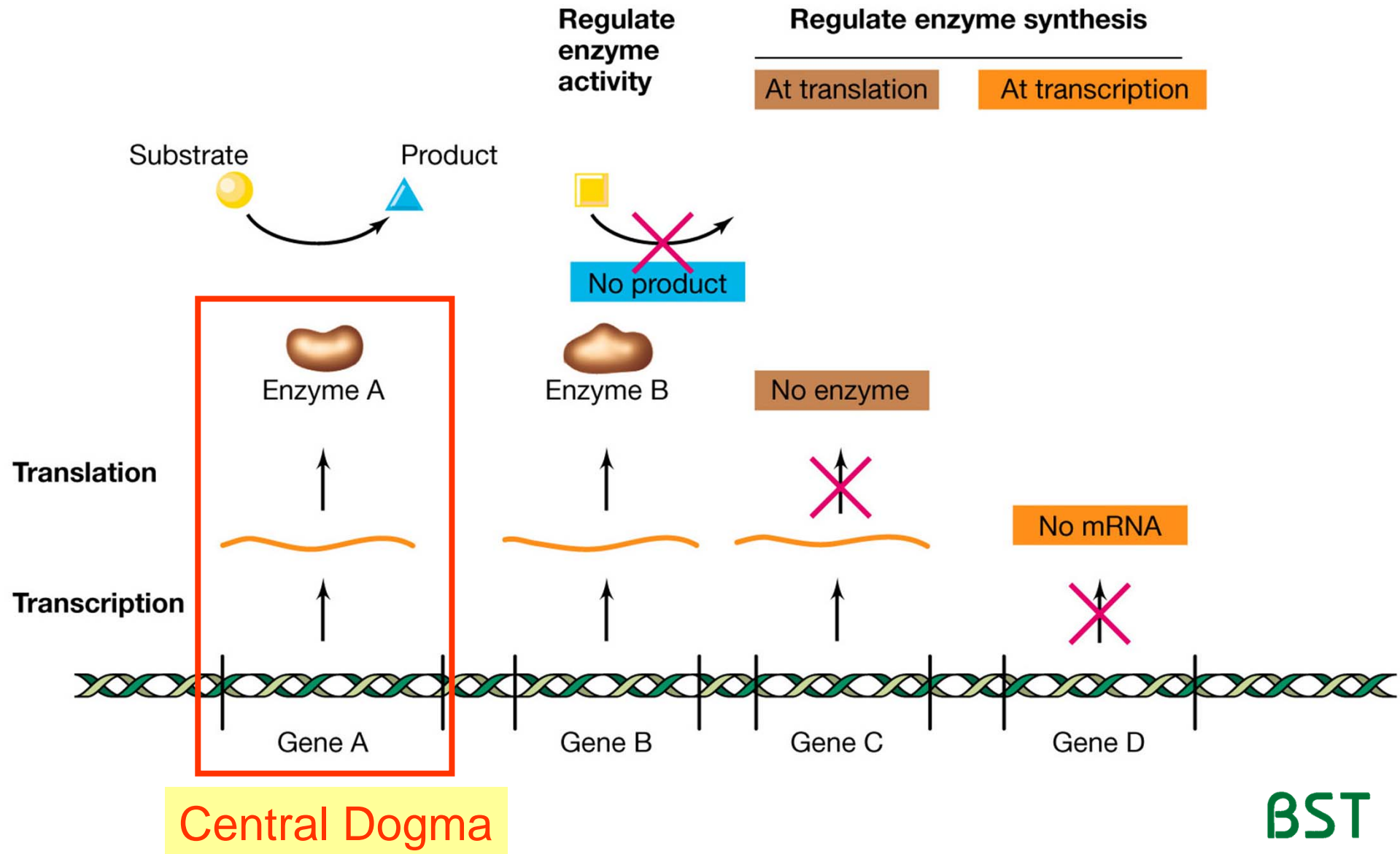
➔ for the synthesis of aromatic amino acids

The secondary metabolite

➔ antibiotics containing aromatic rings.



Regulation Mechanisms



Modification of Gene Expression

- allows for **overproduction** of a product, production of more than one product by the same organism, or synthesis of modified products

Pathway architecture

- analysis, design, and modification of **biochemical pathways** to increase process efficiency

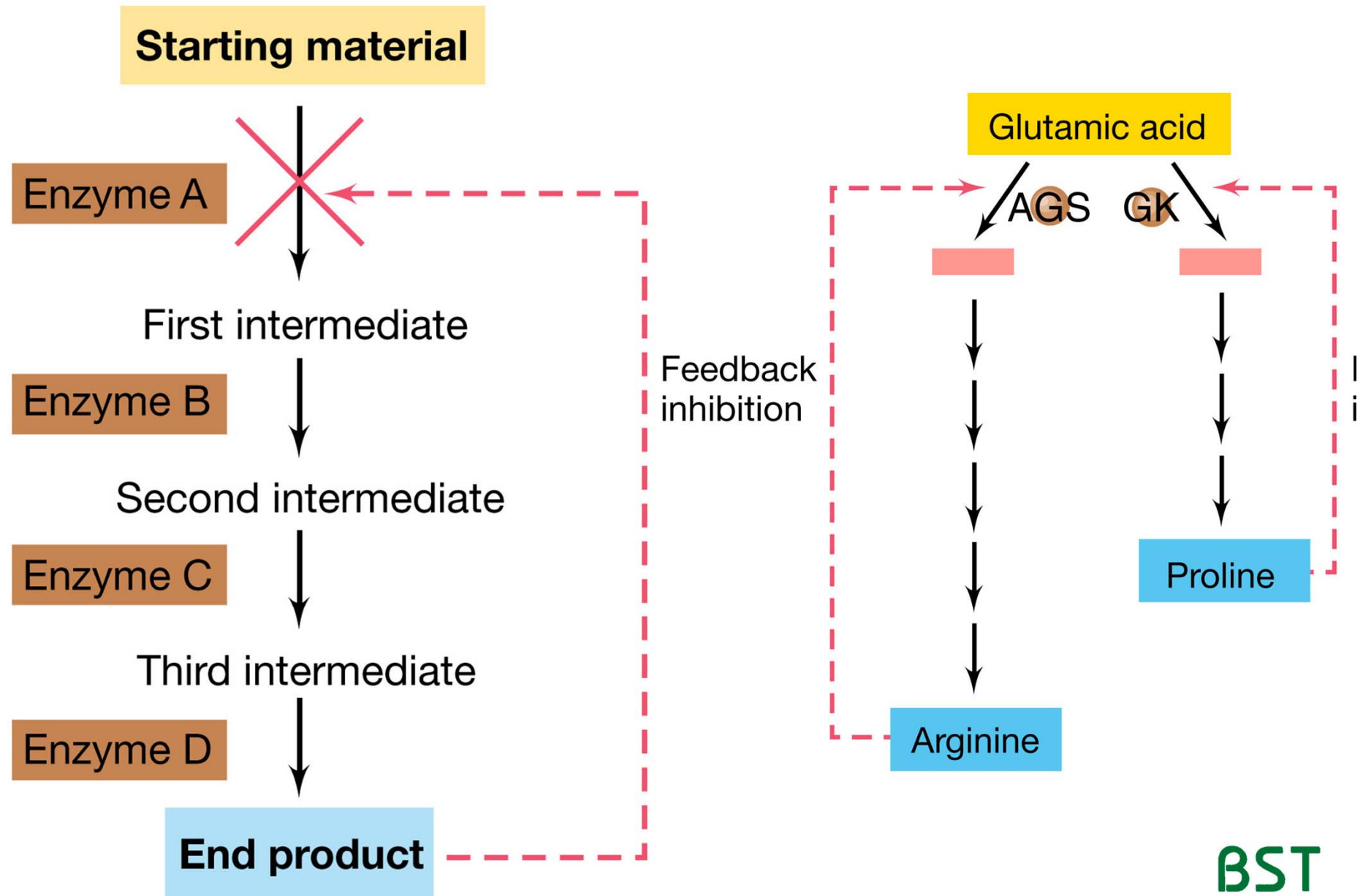
Metabolic pathway engineering

- intentional alteration of **metabolic pathway** by inactivation of specific genes

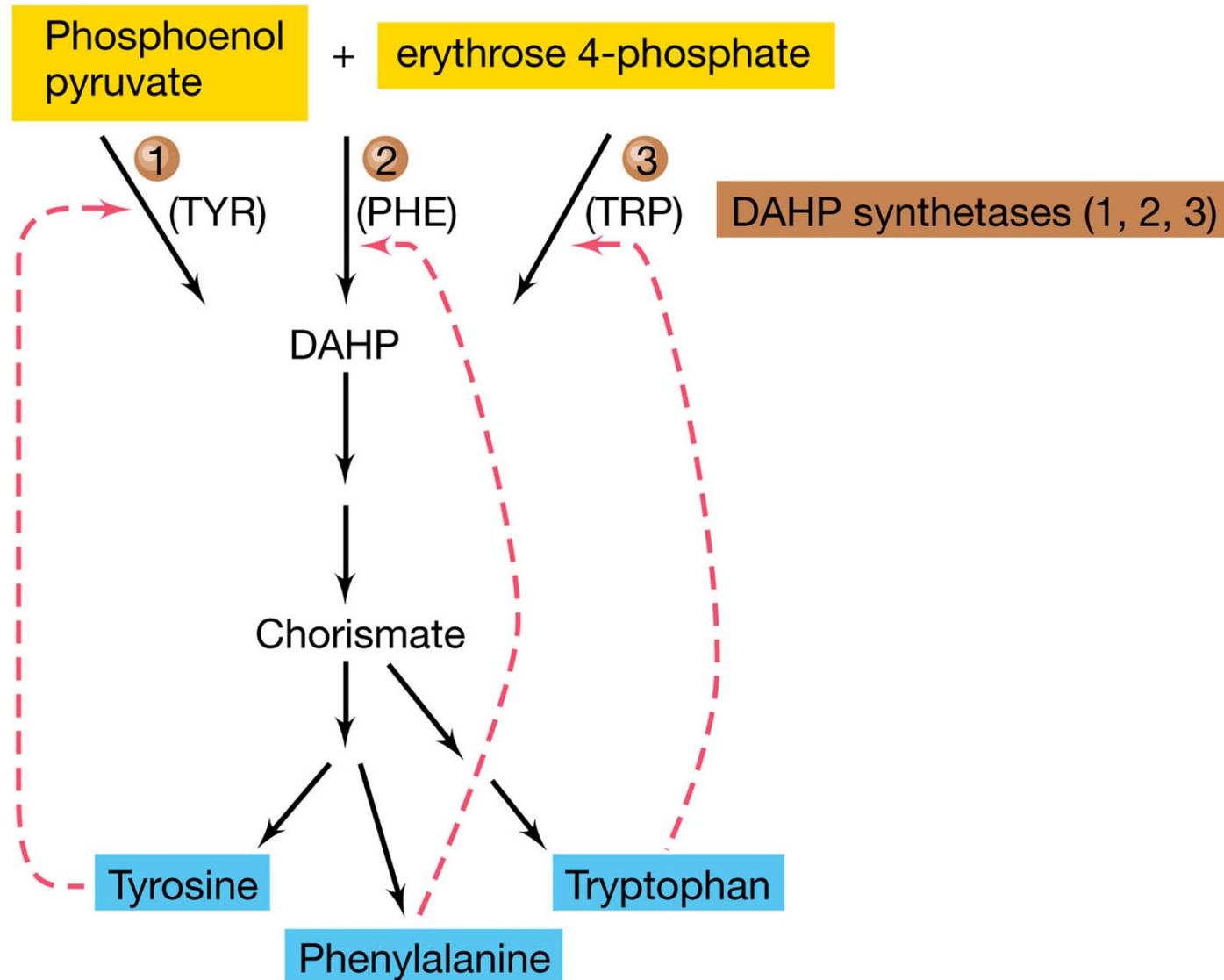
Metabolic control engineering

- alteration of **control mechanisms** of specific genes

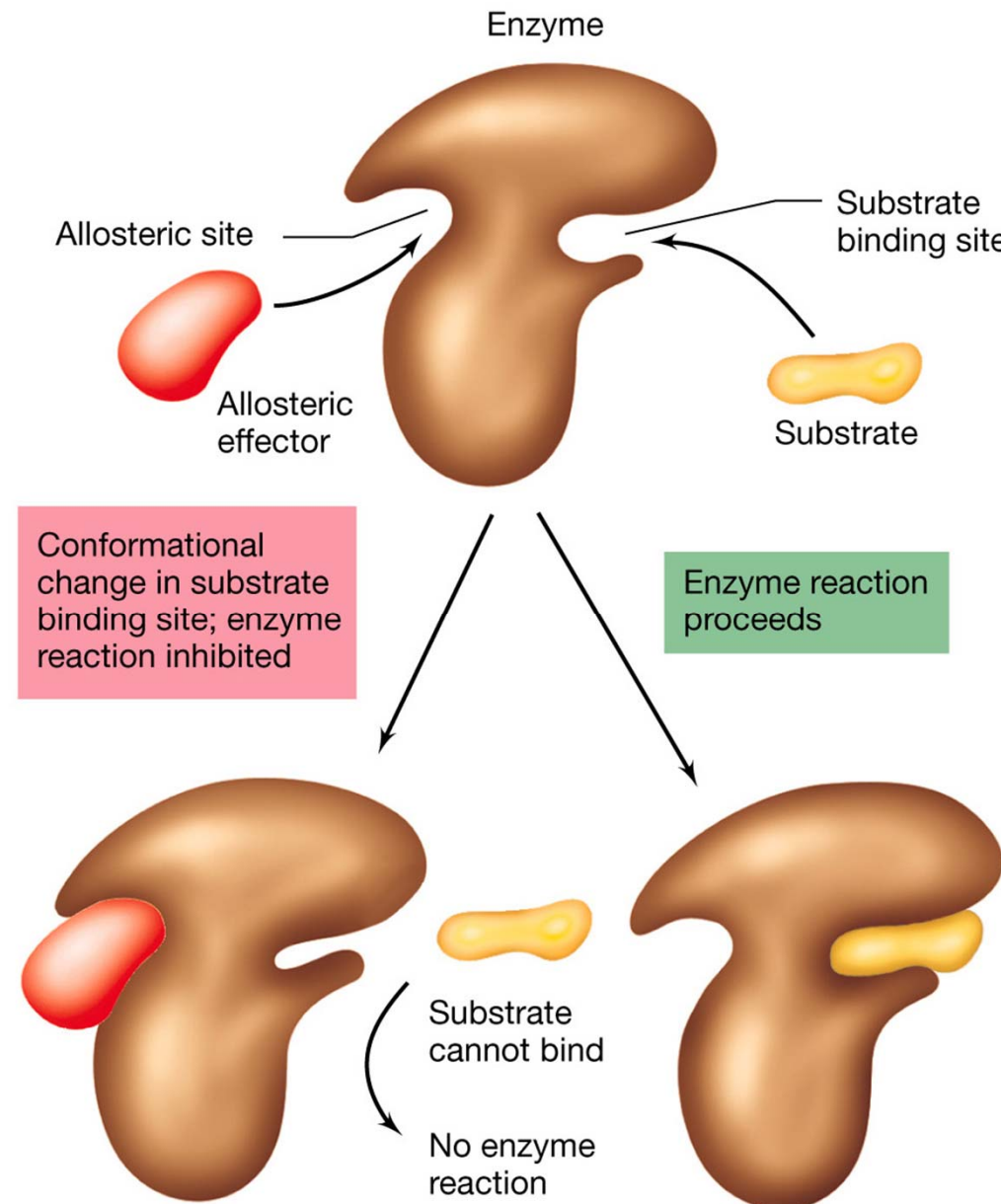
Feedback inhibition of enzyme activity



Specific Feedback Inhibition

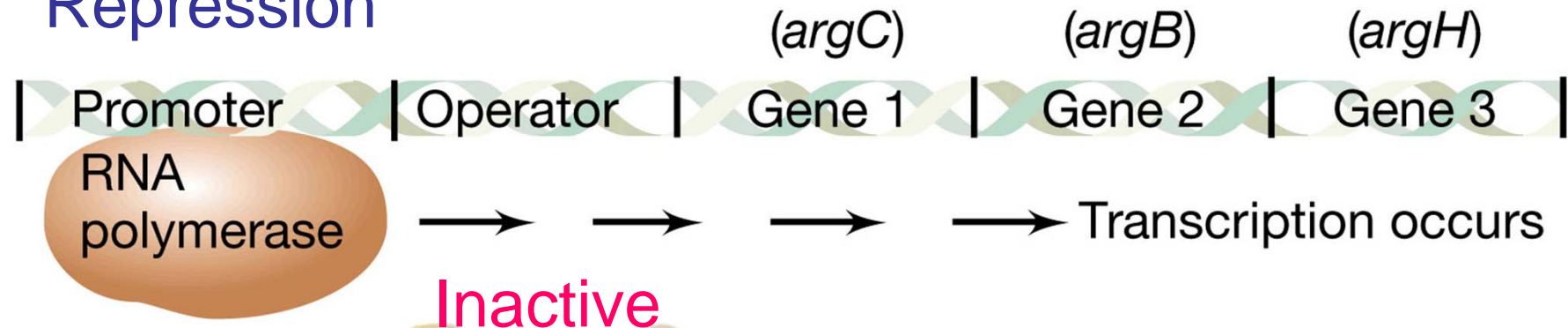


Enzyme Inhibition by Allosteric Effector

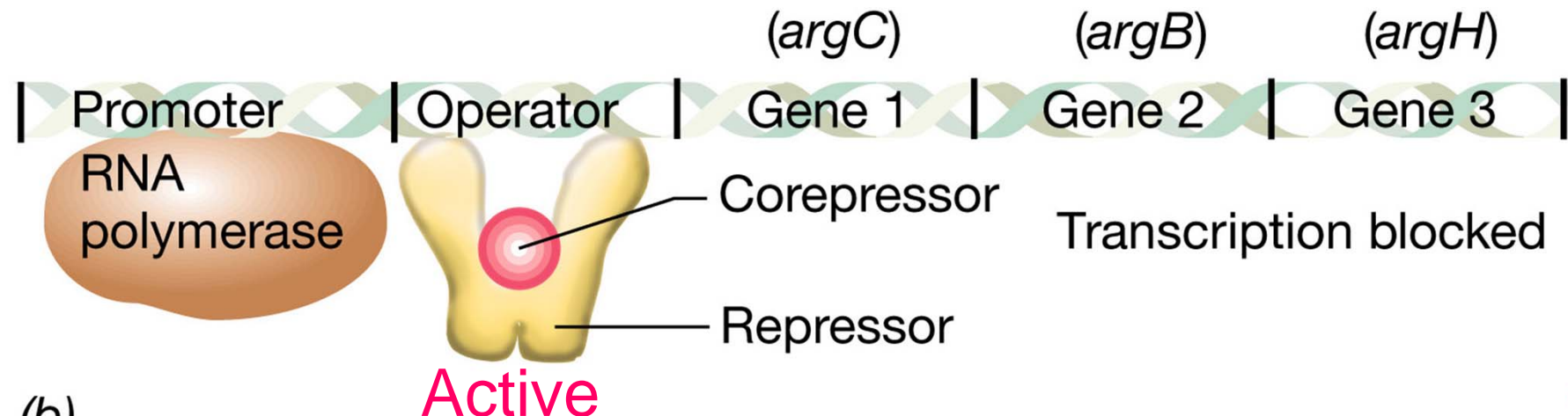


Negative Control of Transcription

Repression



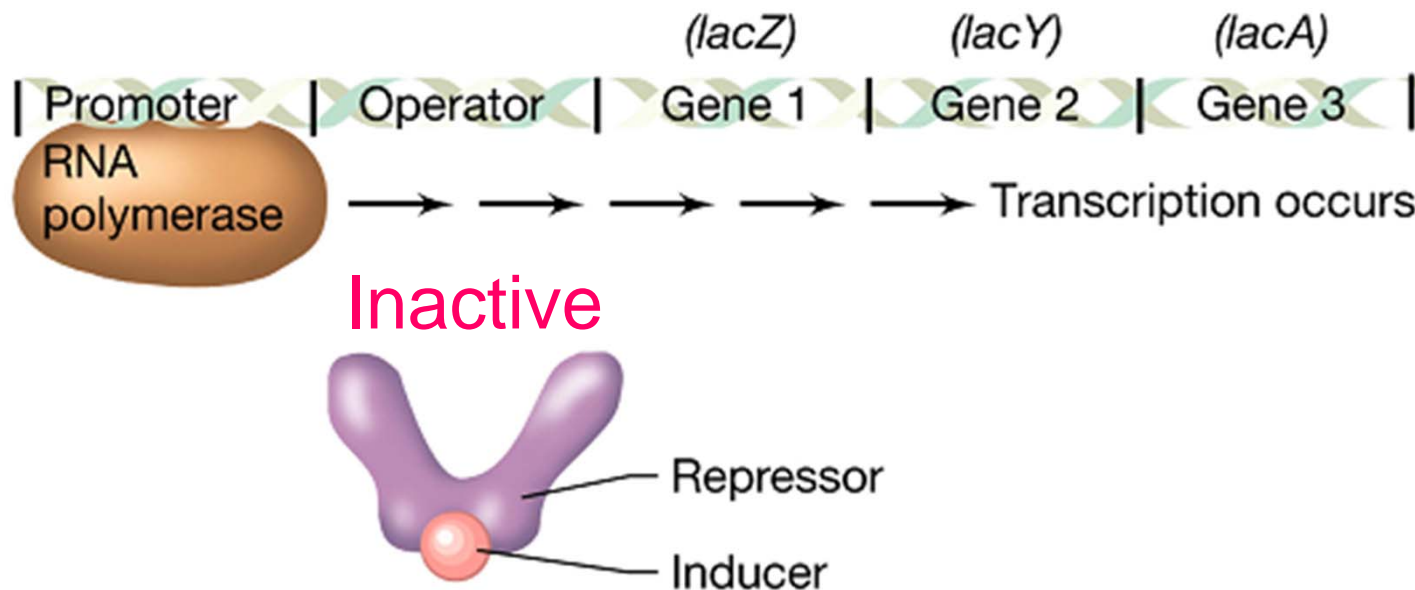
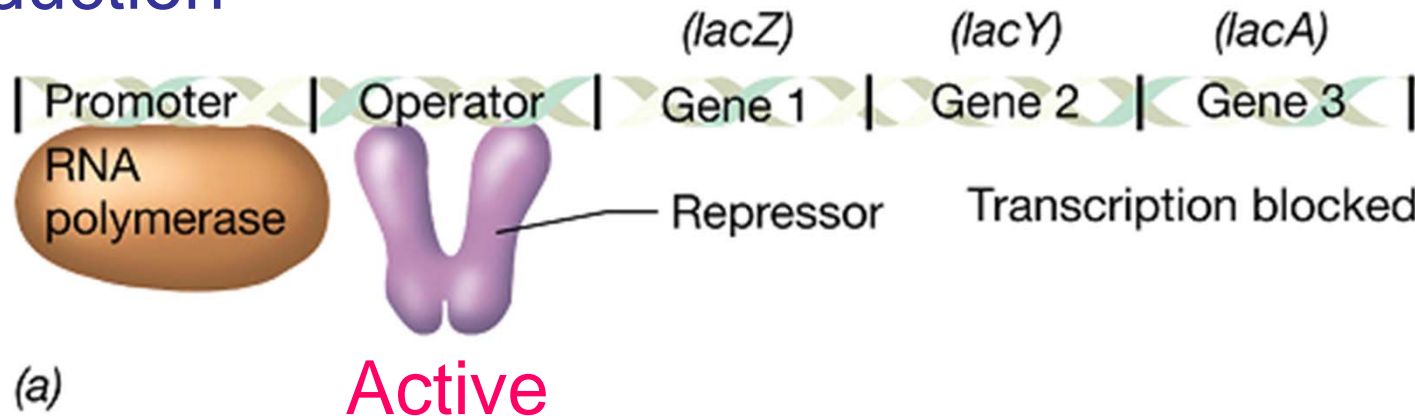
(a)



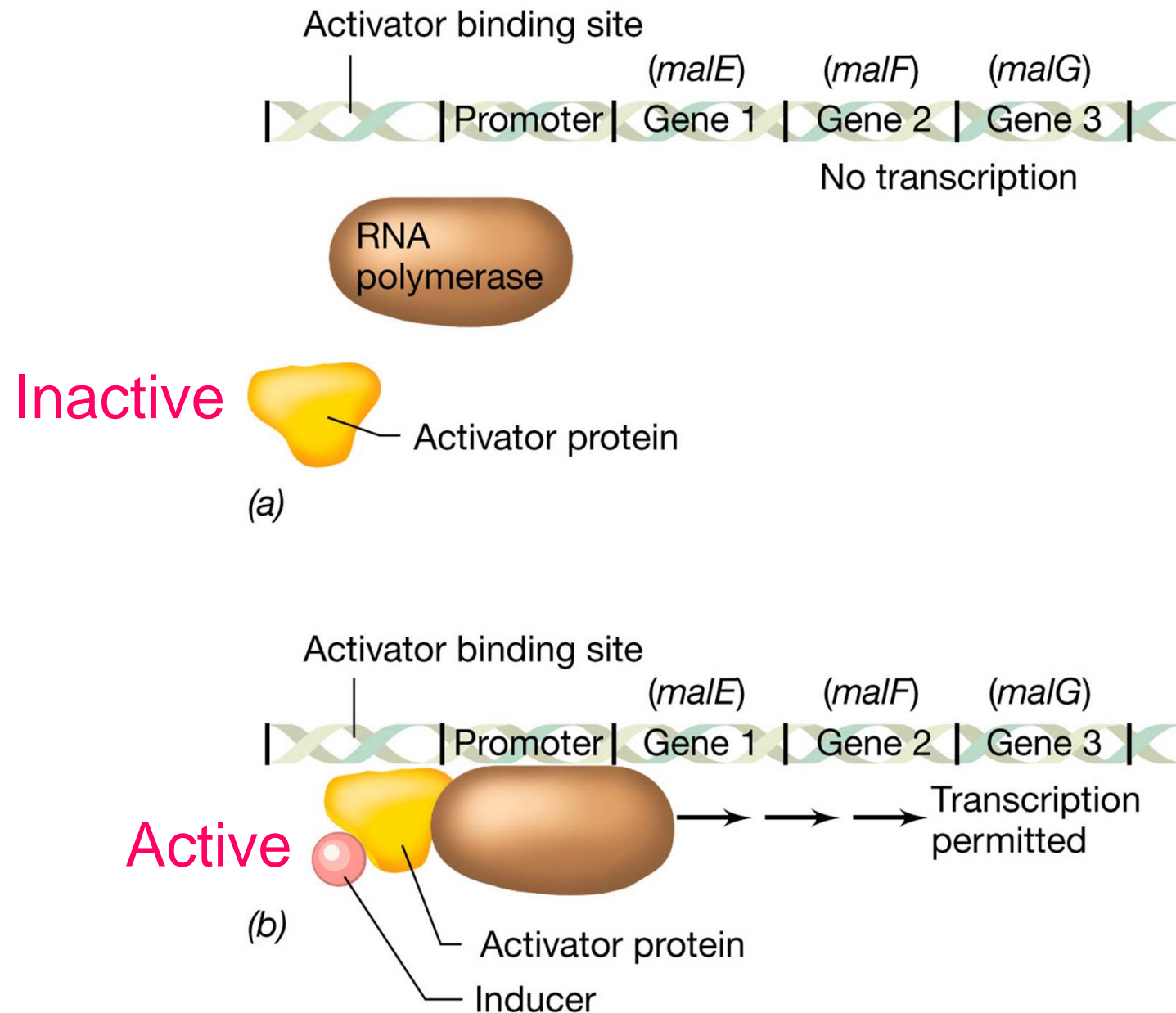
(b)

Negative Control of Transcription

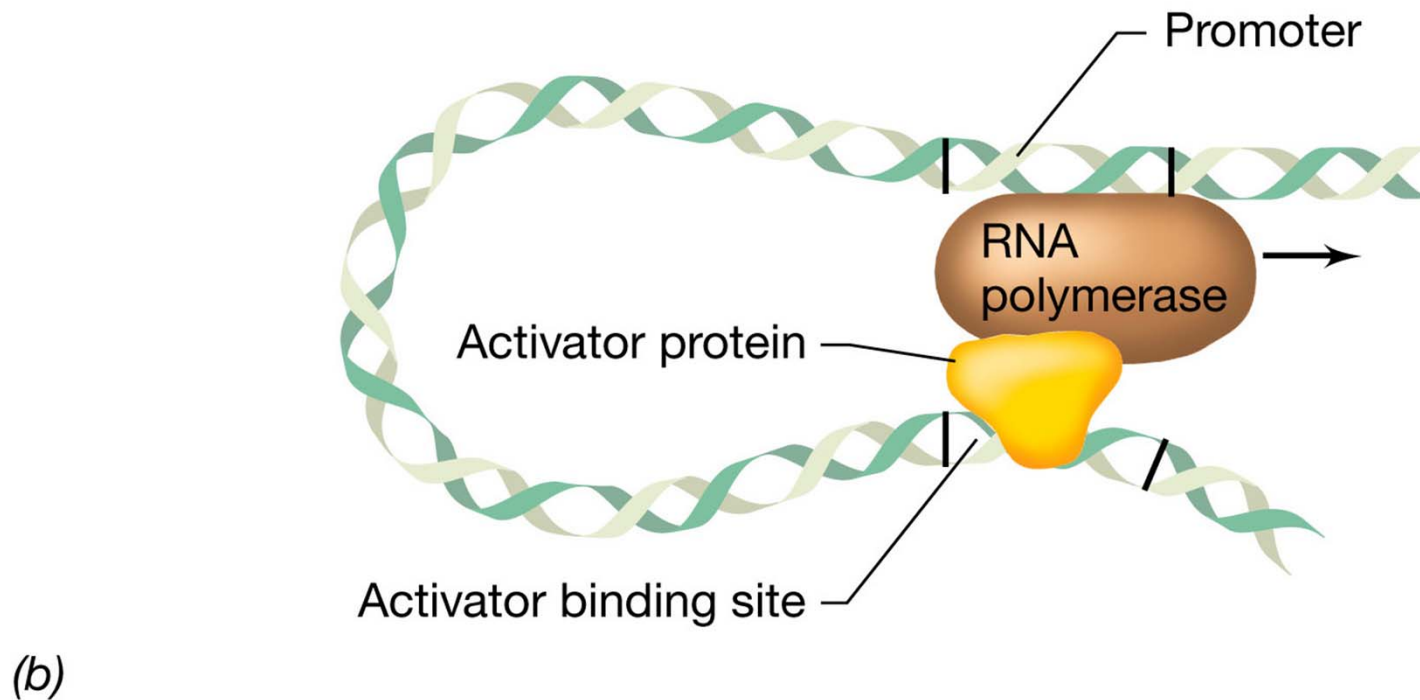
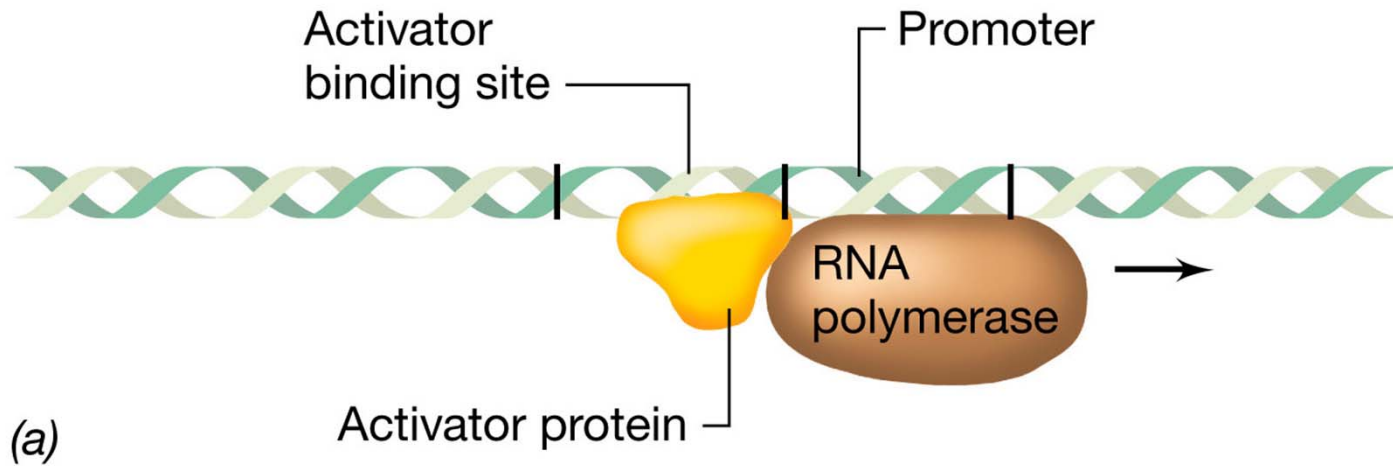
Induction



Positive Control of Enzyme Induction

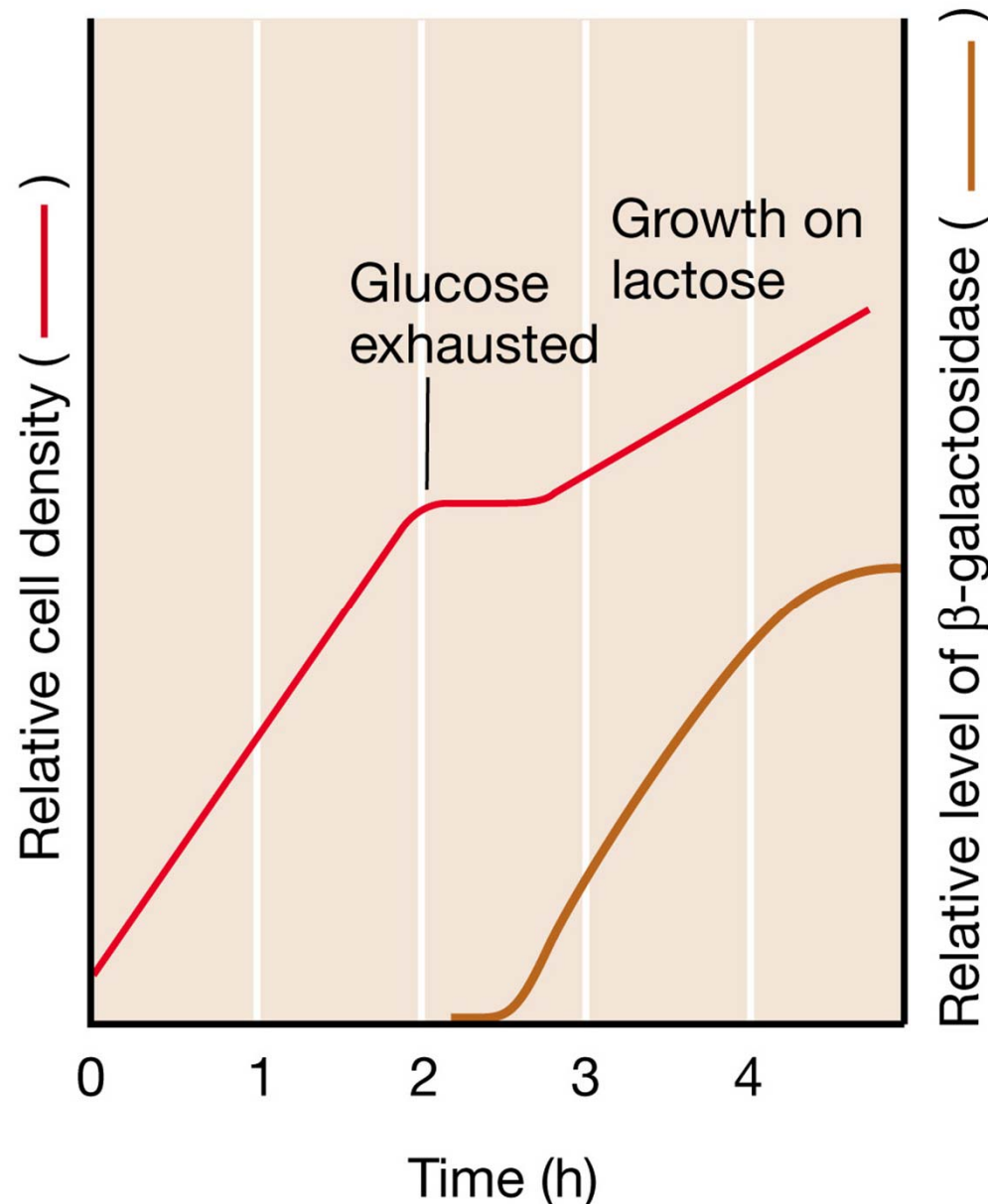


Positive Control of Transcription



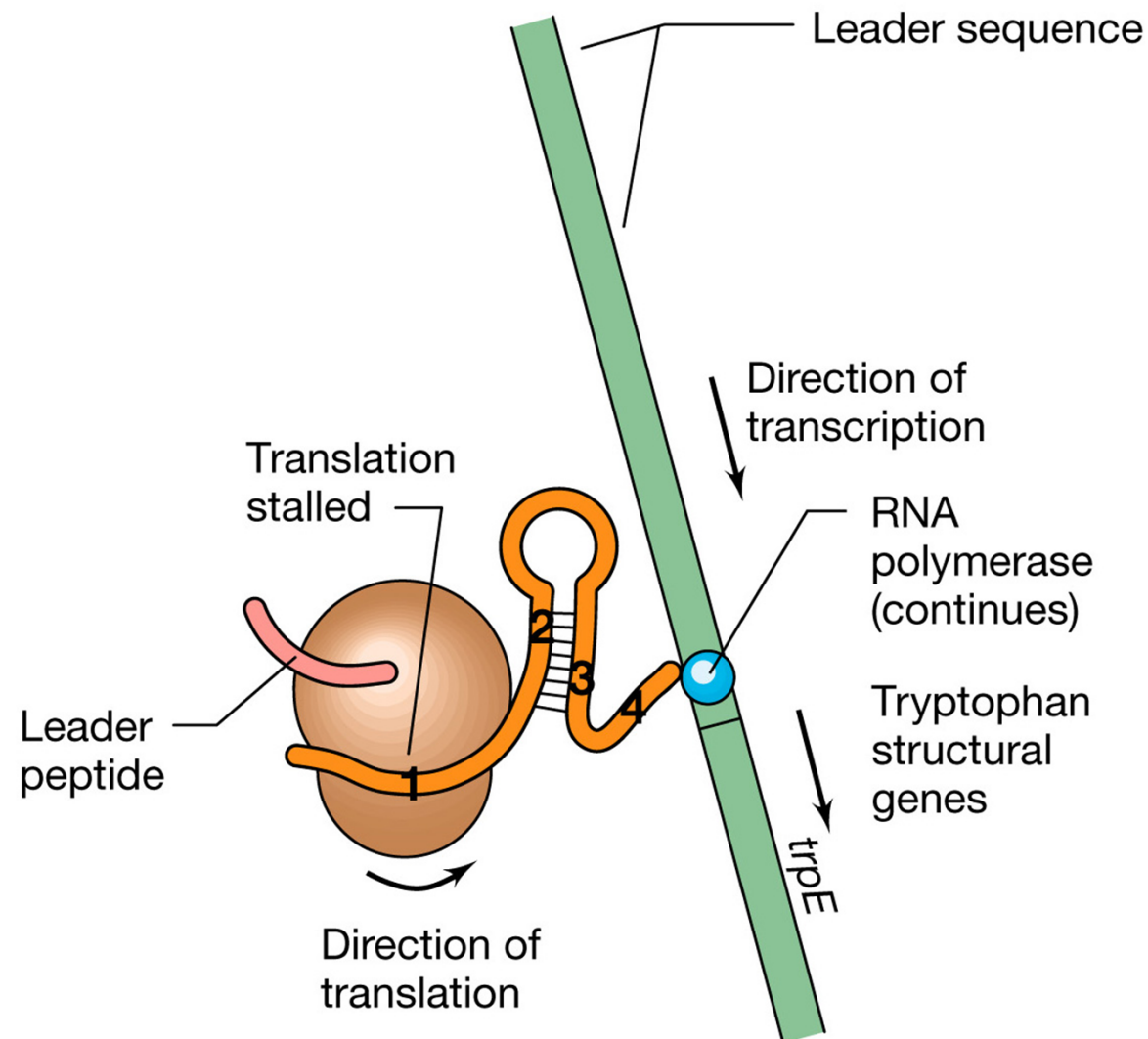
Catabolite Repression

- Occurs when operon is under control of **catabolite** other than initial substrate
- Allows **preferential** use of one carbon source over another



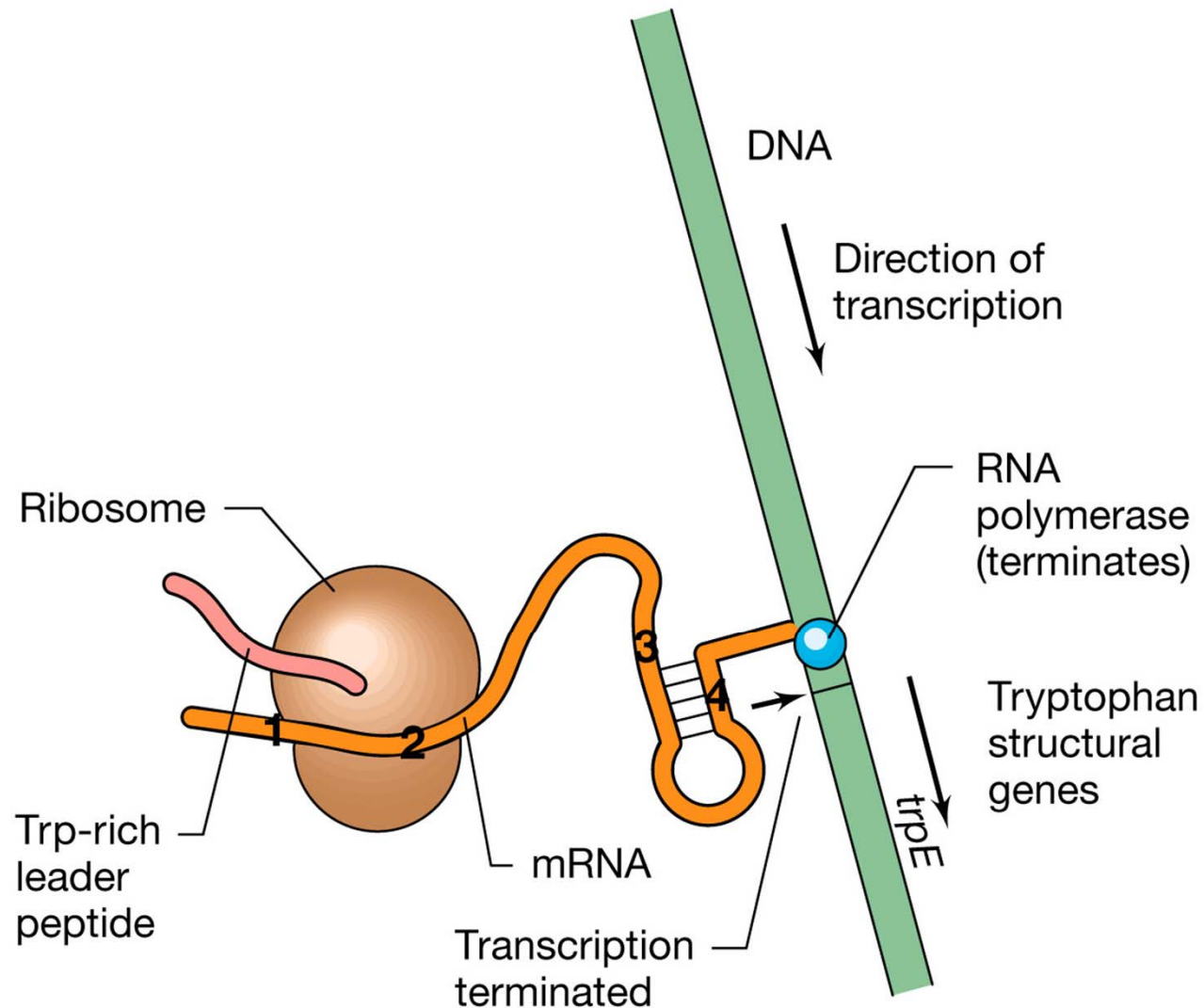
Transcription of *trp* operon by attenuation

Tryptophan-starved: transcription not terminated

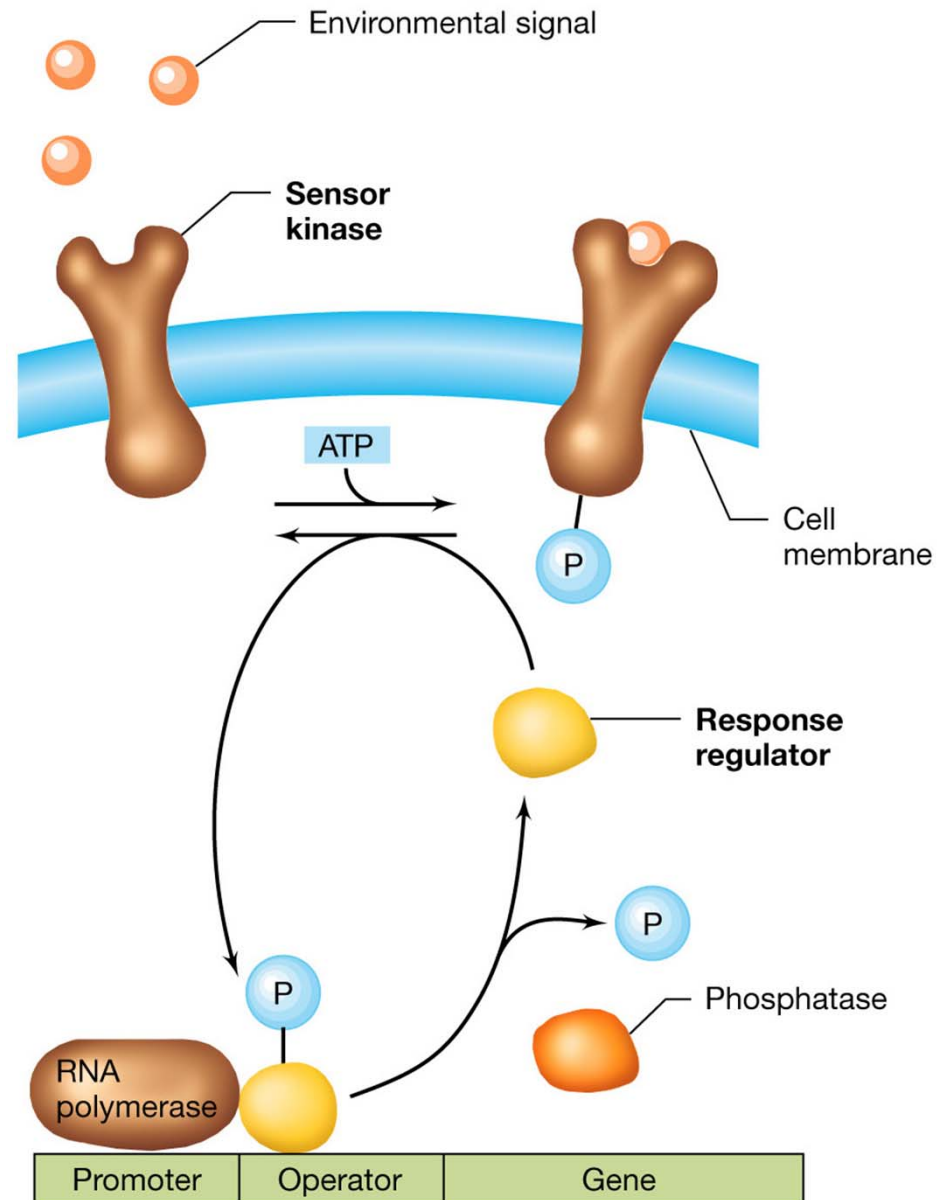


Transcription of *trp* operon by attenuation

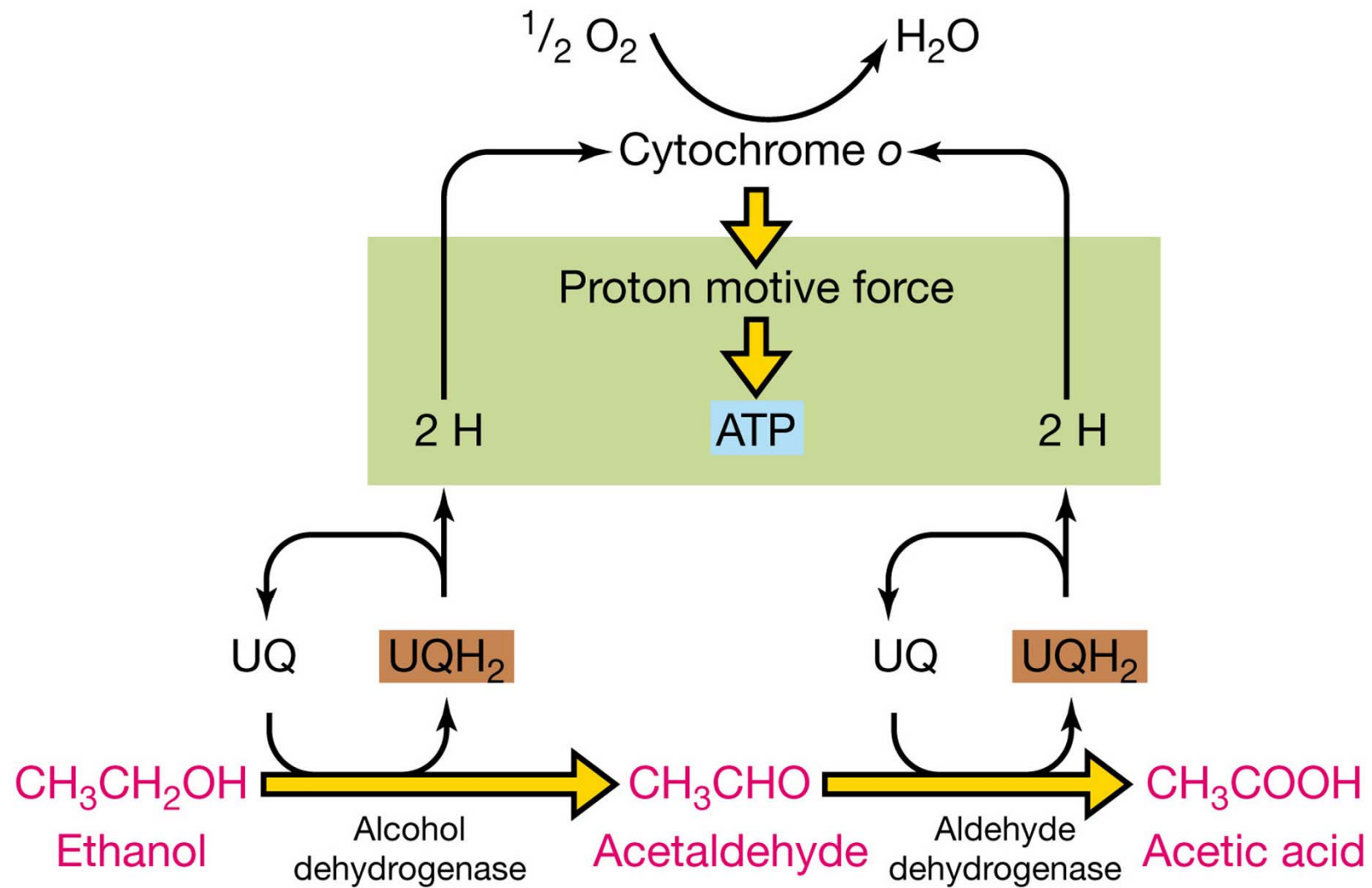
Excess tryptophan: transcription terminated

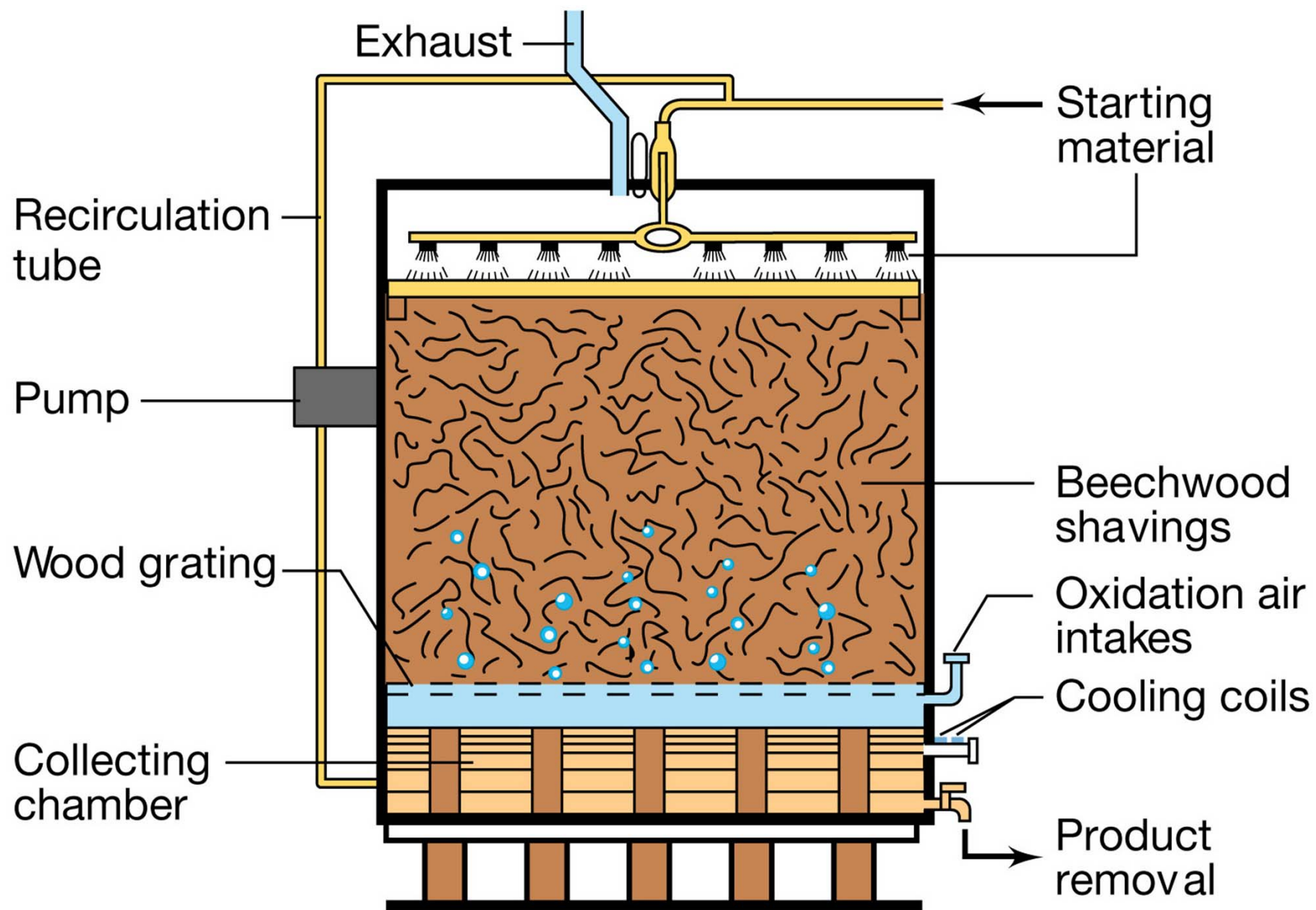


Two-Component System

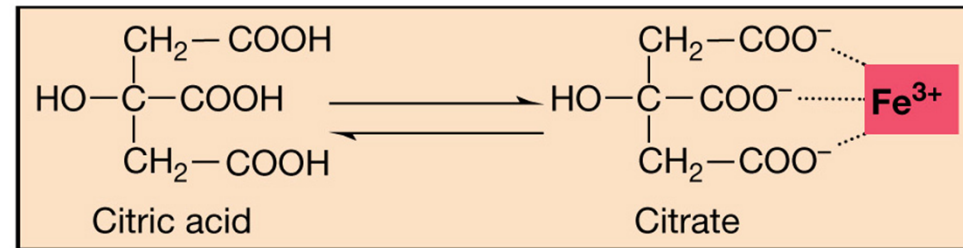


Production of Vinegar

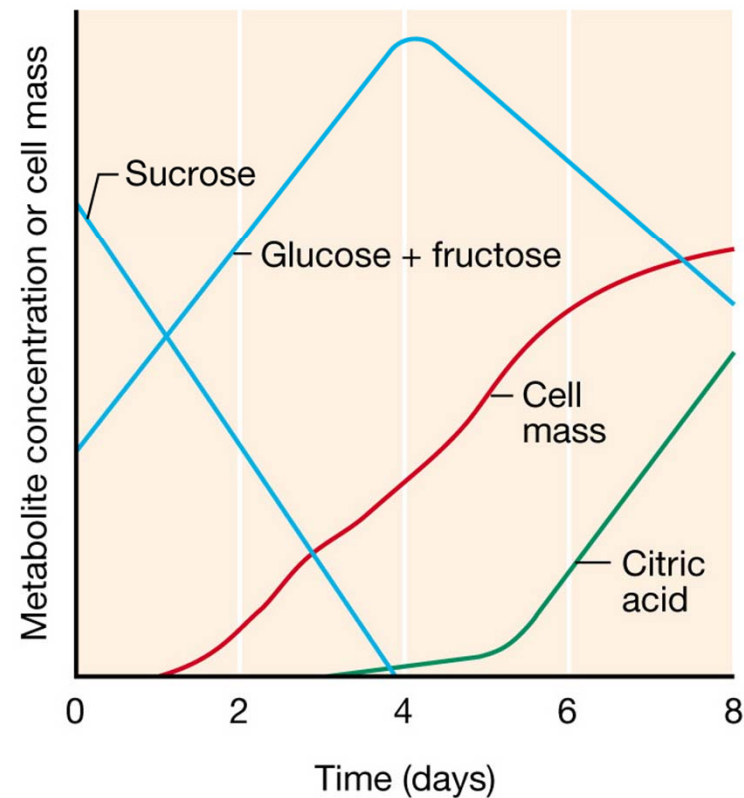




Citric Acid Fermentation



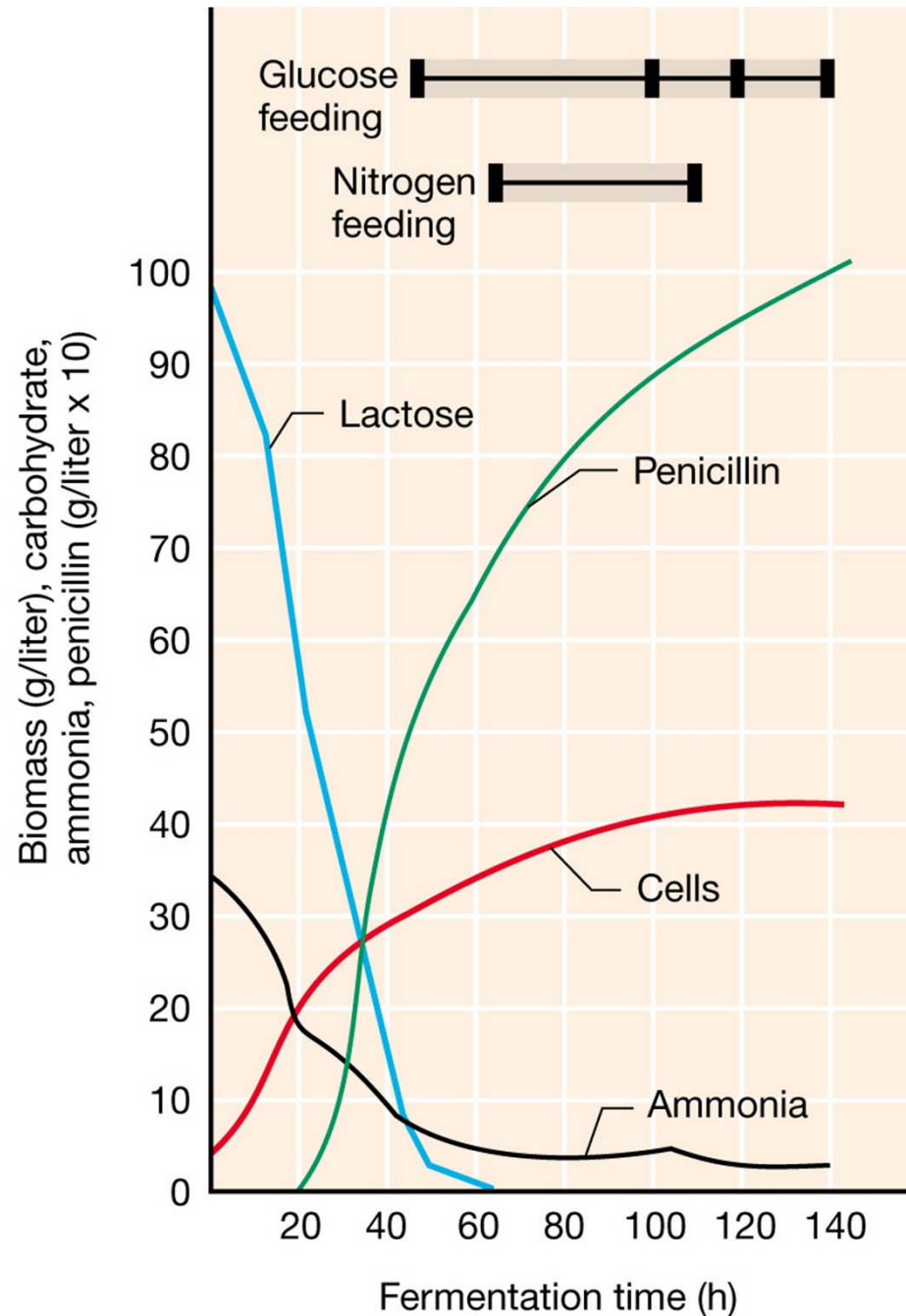
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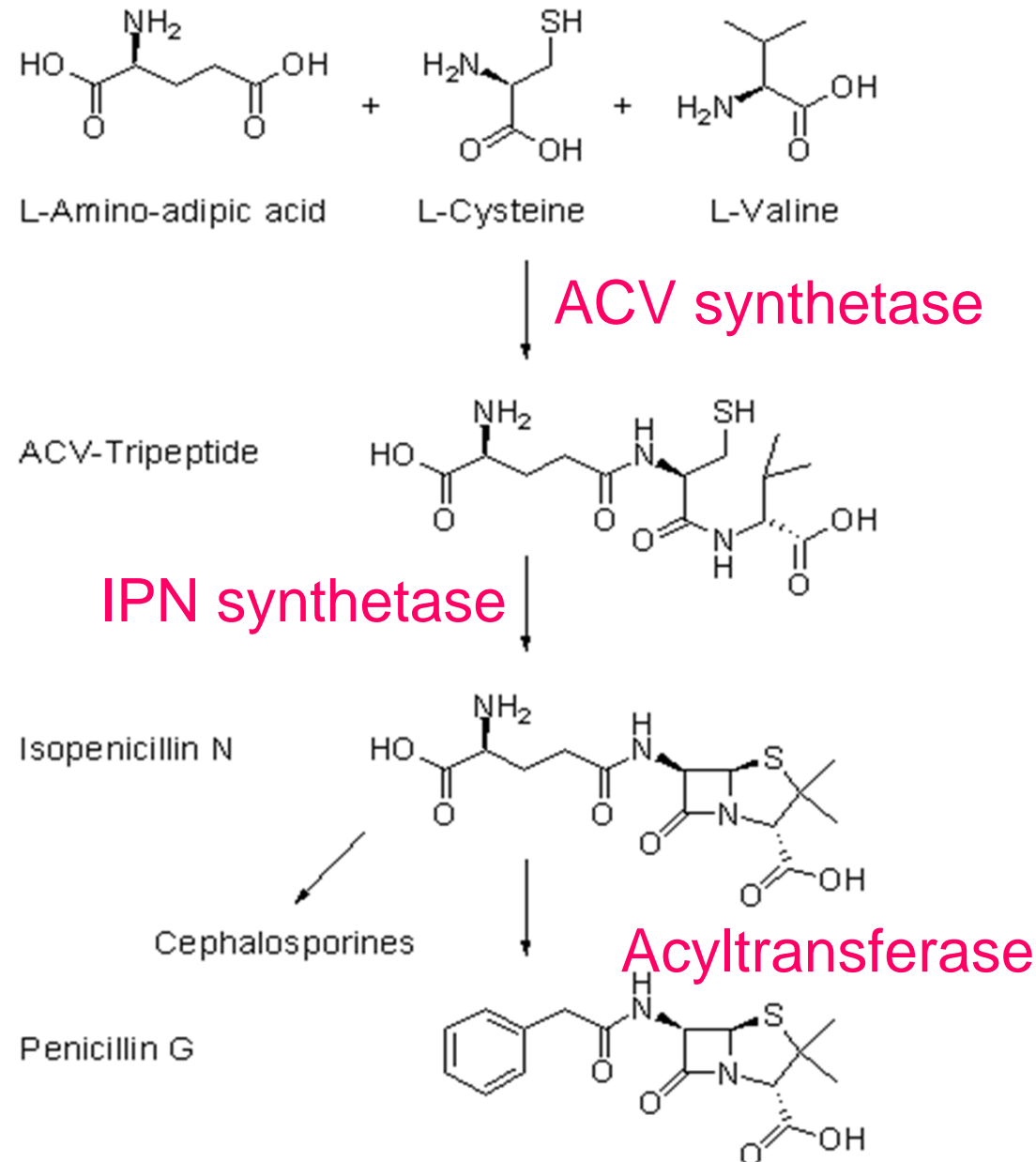
(b)

Antibiotics

- ➔ Requires precise control of nutrients
- ➔ Final product can be modified to yield a variety of **semisynthetic** penicillins



Biosynthetic Pathway of Penicillin



Industrial Production of Penicillins

