TOPIC: COMPETITIVE INHIBITORS OF METABOLIC PATHWAYS

Sulfa Drugs & Folic Acid Synthesis

◆ Recall: bacterial cells synthesize	acid → humans get it from their
РАВА	 ◆ Sulfonamides (Drugs): block folic acid production. → inhibition. ◆ Structurally similar to para-aminobenzoic acid ().
Dihydrofolic acid Enzyme A	 , widely used antibacterial. Used less today, but still used for some & other infections.
Tetrahydrofolic acid Enzyme B	 ◆ Trimethoprim: competitively inhibits the step in folic acid synthesis. ◆ Not a drug.
T	◆ Often used together
DNA and RNA	► E.g. sulfamethoxazole-trimethoprim (Bactrim®):
nucleotides	- More Reduces spread of resistance.

EXAMPLE

For each statement below, determine which of the two words best completes the sentence.

- 1. Sulfa drugs and trimethoprim block the synthesis of (ATP / Folic Acid).
- 2. The combination drug sulfamethoxazole-trimethoprim exhibits a(n) (synergistic / antagonistic) effect.
- 3. Sulfonamides work by (competitive inhibition / non-competitive inhibition).
- 4. Sulfonamides are structurally similar to (PABA / dihydrofolic acid).
- 5. Trimethoprim is structurally similar to (PABA / dihydrofolic acid).

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PRACTICE

Why are sulfa drugs & trimethoprim often used together?

- a) Sulfa drugs target Gram-positive bacteria, while trimethoprim targets Gram-negative bacteria.
- b) Trimethoprim destroys naturally occurring enzymes that inhibit sulfa drugs.
- c) They exhibit synergism by inhibiting different steps of the same metabolic pathway.
- d) Trimethoprim blocks sulfa drugs from affecting folic acid synthesis in human cells.

PRACTICE

Which mechanism below best describes the method by which sulfa drugs inhibit the folic acid production pathway?

- a) Sulfa drugs block the conversion of tetrahydrofolic acid into folic acid.
- b) Sulfa drugs bind to PABA, preventing it from being converted to tetrahydrofolic acid.
- c) Sulfa drugs denature the enzyme that converts dihydrofolic acid to tetrahydrofolic acid.
- d) Sulfa drugs bind to the same enzyme that converts PABA to dihydrofolic acid.