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Methotrexate resistance in leukocytes. Methotrexate (MTX) is used to treat various diseases including rheumatoid arthritis (RA), juvenile idiopathic arthritis, psoriasis, and lymphoma. This drug is widely used due to its relatively low toxicity and low cost. MTX enters the cell by forming an active metabolite, MTX-7-ols, and then further converted to the tetrahydrofolate analogs. Methotrexate is a folate analogue that inhibits dihydrofolate reductase. MTX is generally administered orally, and its intracellular concentrations are therefore thought to be determined by folate uptake and intracellular MTX-7-ols production. MTX is widely used for the treatment of autoimmune diseases. As a result, patients receiving oral MTX therapy often develop severe mucositis and/or severe hair loss, which limits the administration of MTX for the treatment of rheumatoid arthritis. MTX has been used for a long time with a relatively low dose; however, this drug is also known to cause serious adverse effects. MTX resistance is one of the mechanisms responsible for its side effects and severe toxicity. To treat MTX-resistant patients, MTX is usually administered together with folic acid (FA) and/or leucovorin (LV). FA competes with MTX for the binding site on dihydrofolate reductase (DHFR), and also inhibits the MTX-7-ols formation. LV is a synthetic derivative of LV-FA and is expected to promote the regeneration of dihydrofolate and to increase the intracellular concentration of MTX.

Q: Why do I get a segmentation fault? I'm writing a code, and I encountered a segmentation fault problem. I'll post the code, and the stacktrace, here. Note that my code works, but without the segmentation fault. Here's the code: `int main(){ char *str = malloc(sizeof(char)*20); strcpy(str, "hello"); return 0; }`

Here's the stacktrace: Program received signal SIGSEGV, Segmentation fault. 0x7ff 82157476af

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