

Table S1 List of drug hits decreasing yeast extract productivity

Drug	Mode of action	Targets	
Hygromycin B	Antibiotic	Ribosomes	30S rRNA
	inhibits protein synthesis	Ribosome-dependent ATPase	RbbA
Doxorubicin (Adriamycin)	Antibiotic	DNA Topoisomerase	TOP1, TOP2A, TOP2B
	induces DNA damage and apoptosis	Nucleic acids	DNA
		Nucleolar phosphoprotein	NOLC1
Puromycin 2HCl	Antibiotic	Ribosomal proteins	RPL10L, RPL13A, RPL23, RPL15, RPL19, RPL23A, RSL24D1, RPL26L1, RPL8, RPL37, RPL3, RPL11,
	inhibits protein synthesis	tRNA-protein transferase	aat
Proflavine Hemisulfate	Antiseptic intercalates DNA	Nucleic acids	DNA
Ethacridine lactate monohydrate	Antiseptic intercalates DNA	Nucleic acids	DNA
Mitoxantrone HCl	type II topoisomerase inhibitor	DNA Topoisomerase	TOP2A
		Nucleic acids	DNA
	intercalates DNA		
Aprotinin	Serine protease inhibitor	Serine protease	PRSS1, CTRB1, PLG, KLK1
Gentamicin Sulfate	Antibiotic inhibits protein synthesis	Ribosomes	rpsL, 16S rRNA, 23S rRNA,
		Membrane	
		Synthetase	nadE
		Reductase	DHFR
Pentamidine	Antiprotozoal agent	PRL Phosphatases	PTP1B

	inhibits synthesis of DNA, RNA, phospholipids and proteins	tRNA-methyltransferase Nucleic acids	TRDMT1 DNA
Heparin sodium	Anticoagulant	Serpin protease inhibitor	SERPINC1
	Competitive inhibitor of RNA synthesis	Coagulation factor X	F10
		Selectin	SELP
		Tyrosine kinase	FGFR1, FGFR2, FGFR4
		Fibroblast growth factor	FGF1, FGF2, FGF4, FGF19
		Cytokine	PF4
		Polymerases	T7RNAP, polA
Paromomycin Sulfate	Antibiotic inhibits protein synthesis	Ribosomes	16S rRNA, rpsJ, RPSA, RPL10L
Calcium Levofolinate	Calcium salt of folinic acid and analogue of tetrahydrofolate	Serine hydroxymethyltransferase	glyA
Carbenoxolone Sodium	Synthetic derivative of glycyrrhetic acid	Pannexon membrane channels and related innexon channels	11 β -HSD