Sites of modulation of opioid tolerance

Receptor

Desensitization
Internalization/trafficking

Transduction systems

Adenylyl cyclase G protein regulation RGS proteins Guanylyl cyclase

Neurotransmitters and their circuits

Cholecystokinin δ Opioid receptor

Enkephalin

Glutamate transporter GLT-1
Glutamine synthase

Neuropeptide FF

NO

NMDA receptors/glutamate Orphanin FQ/nociceptin

Superoxide dismutase

Pharmacokinetic

P-glycoprotein

RGS, regulator of G-protein signaling.