RETROGRADE ENDOCANNABINOID SIGNALING Synaptic cleft Neuron (Postsynaptic cell) Neuron (Presynaptic terminal) Ca Ca Calcium signaling pathway Depolarization Na + _**െ**²⁺ Endocytosis **Excitatory terminal** Synaptic vesicle cycle Glu ಄ Glu Depolarization-induced suppression of excitation (DSE) NAT PhosC Exocytosis 0 —O◀ NAPE Glutamatergic Glu 🖷 ි**ර**ි PhosEA AEA, synapse Gene expression Gene expression phospho-AEA 2-AG cAMP **○**◀ K02677... Action potential DAG Ţ<mark>G</mark>βγ K13700 K04997... PG-G **►⊙**7 ʹĠβу OGL OEA AEA ECT K04277 **▶©**2-AG 2-AG ΑĒΑ K04277 Gβy **⊌**Gβγ K04371... ▲ Gene Gene expression K04344... **P**O Inhibitory terminal cAMP Depolarization-induced suppression of inhibition (DSI) ^ලල34 eCB-LTD GABA GABAergic Cl -K05175... synapse 0 K15291 <u>O4</u> Synaptic vesicle cycle GABA. GABA Action potential Endocytosis Neuron (Presynap tic terminal) 04723 10/23/15 (c) Kanehisa Laboratories