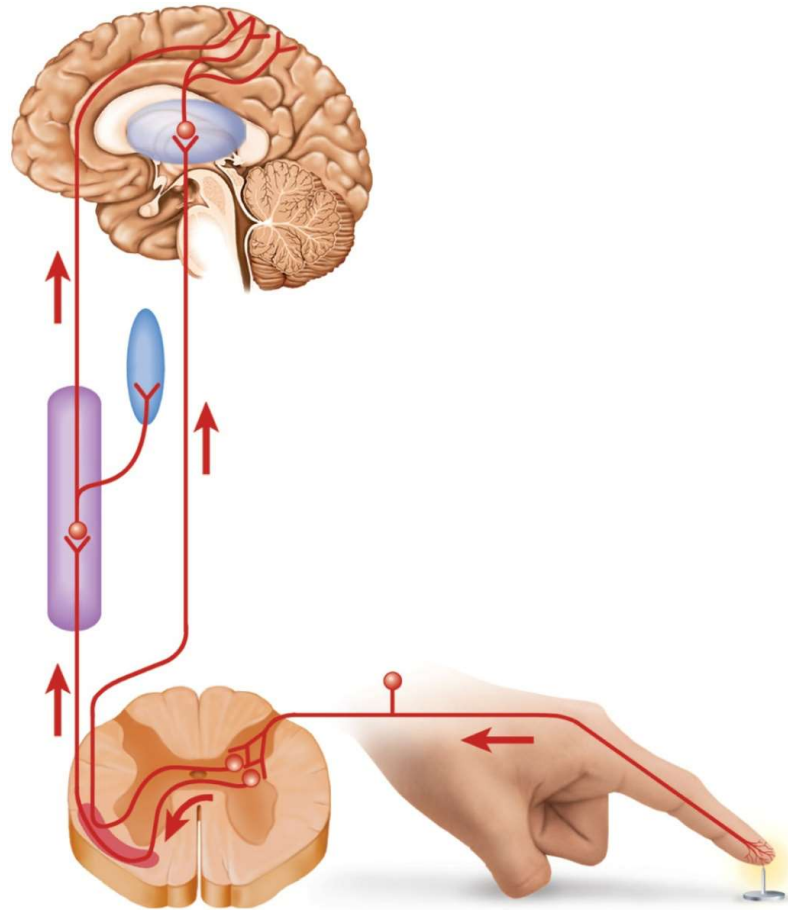


The Sense of Pain



Pain

- Discomfort caused by tissue injury or noxious stimulation that typically leads to evasive action
 - tells us something is wrong!
 - protect us from sustaining more tissue damage
 - lost of pain in diabetes mellitus = **diabetic neuropathy**
 - if you loose sense of pain then you can not detect tissue damage (e.g. Leprosy)
- **Nociceptors** = pain receptors (unencapsulated dendrites)
 - two types providing different pain sensations
 - **fast pain (alpha)** travels in myelinated fibers at 12 - 30 m/sec /// sharp, localized, stabbing pain perceived with injury
 - **slow pain (delta)** travels unmyelinated fibers at 0.5 - 2 m/sec /// longer-lasting, dull, diffuse feeling
 - Both type in skin // damage skin first detect by alpha fibers followed by delta fibers

Pain

- **somatic pain** - from skin, muscles and joints
- **visceral pain** - from the viscera - stretch, chemical irritants or ischemia of viscera (poorly localized)
- injured tissues release chemicals that stimulate pain fibers
 - **bradykinin** - most potent pain stimulus known
 - makes us aware of injury and activates cascade of reactions that promote healing
 - histamine, prostaglandin, potassium ions, ATP, and serotonin are molecules that also stimulate nociceptors

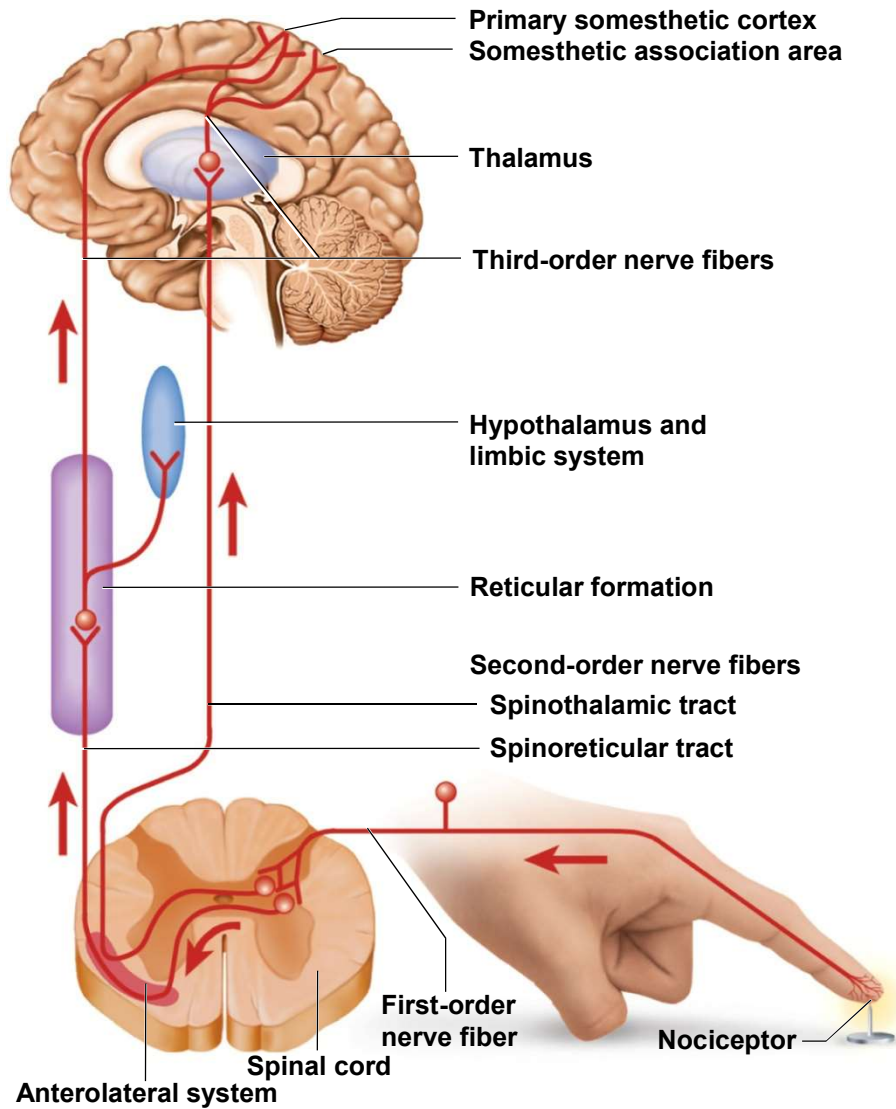
Projection Pathway for Pain

- Ascending and descending tracts (plus multiple sub-routes)
 - Ascending pain neurons route to somatic sensory gyrus:
 - **first-order neuron** cell bodies in dorsal root ganglion of spinal nerves or cranial nerves V, VII, IX, and X
 - **second-order neurons** decussate and send fibers up spinothalamic tract (somatic pain) or through medulla to thalamus /// 2nd order gracile fasciculus carries visceral pain signals
 - **third-order neurons** from thalamus reach postcentral gyrus of cerebrum (somatic sensory gyrus)

Projection Pathway for Pain

- Pain signals travel by way of four ascending tracts:
 - **spinothalamic tract** – most significant pain pathway /// carries most **somatic pain signals**
 - **spinoreticular tract** – carries pain signals to reticular formation /// activate visceral, emotional and behavioral reactions to pain
 - **gracile fasciculus** – carries signals to the thalamus for **visceral pain** -- lower extremities
 - **cuniate fasciculus** – carries signals to the thalamus for **visceral pain** -- upper extremities

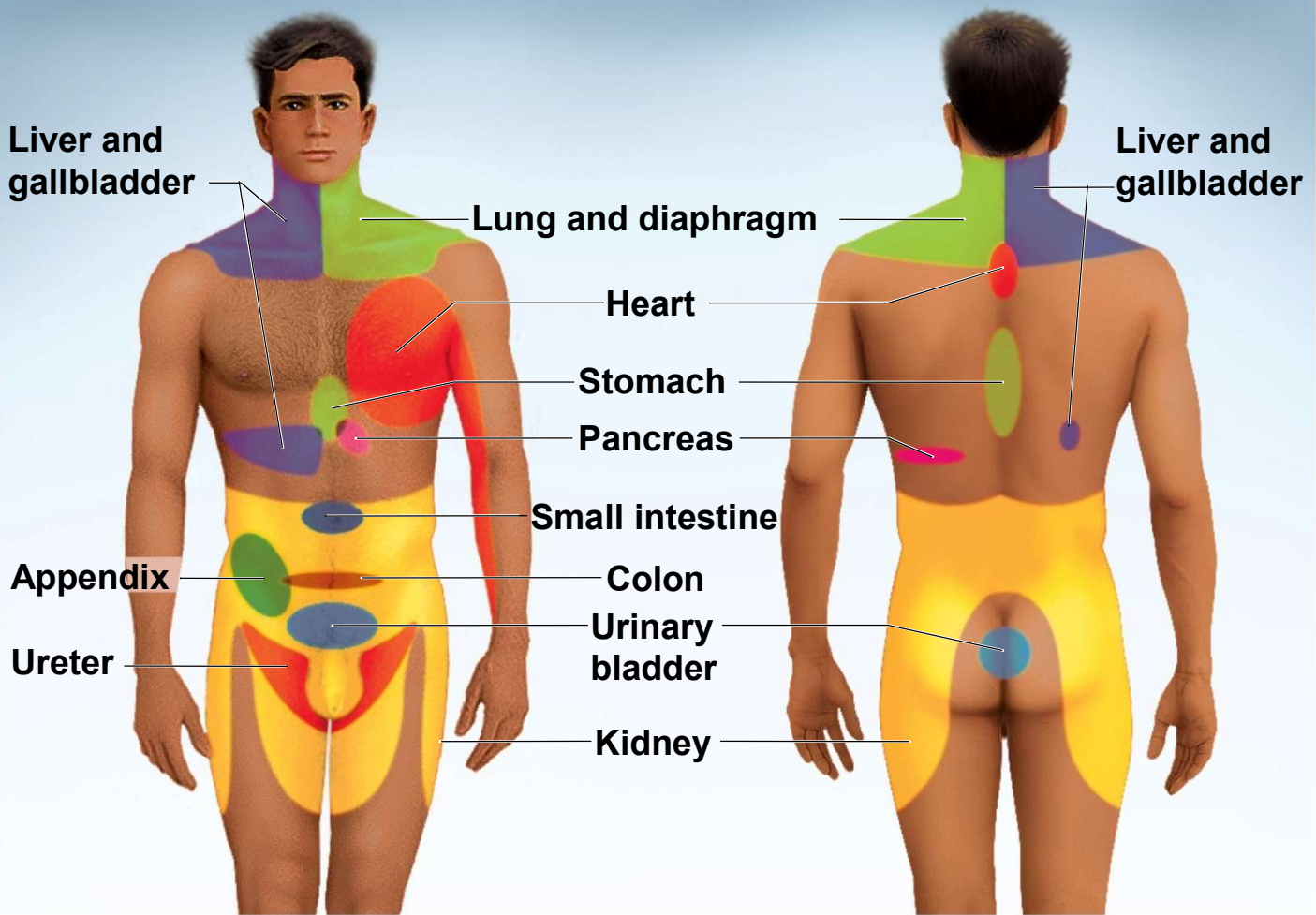
Pain Signal Destinations



Referred Pain

- Pain originates in deep viscera but mistakenly thought to come from the skin or another superficial site
 - results from convergence of neural pathways in CNS
 - brain “assumes” visceral pain is coming from skin
/// brain can not distinguish true source of pain
 - pain originating in heart pain felt in shoulder or arm because both send pain input to spinal cord segments T1 to T5

Referred Pain



CNS Modulation of Pain

- **analgesic** (pain-relieving) // target mechanisms in CNS /// just beginning to be understood
 - tied to receptor sites located in the brain for drugs like opium, morphine & heroin
 - **enkephalins** - two analgesic oligopeptides with 200 times the potency of morphine /// made naturally by our bodies
 - **endorphins**
 - **dynorphins**
 - other larger analgesic neuropeptides have been discovered

CNS Modulation of Pain

- **endogenous opioids** /// internally produced opium-like substances
 - enkephalins, endorphins, and dynorphins
 - secreted by the CNS, pituitary gland, digestive tract, and other organs
- **neuromodulators**
 - enkephalins can block the transmission of pain signals
 - produce feelings of pleasure and euphoria

Spinal Gating

- Stops pain signals at the posterior horn of the spinal cord
 - descending analgesic fibers arise in brain stem
 - travel down the spinal cord in the **reticulospinal tract**
 - block pain signals from traveling up the cord to the brain

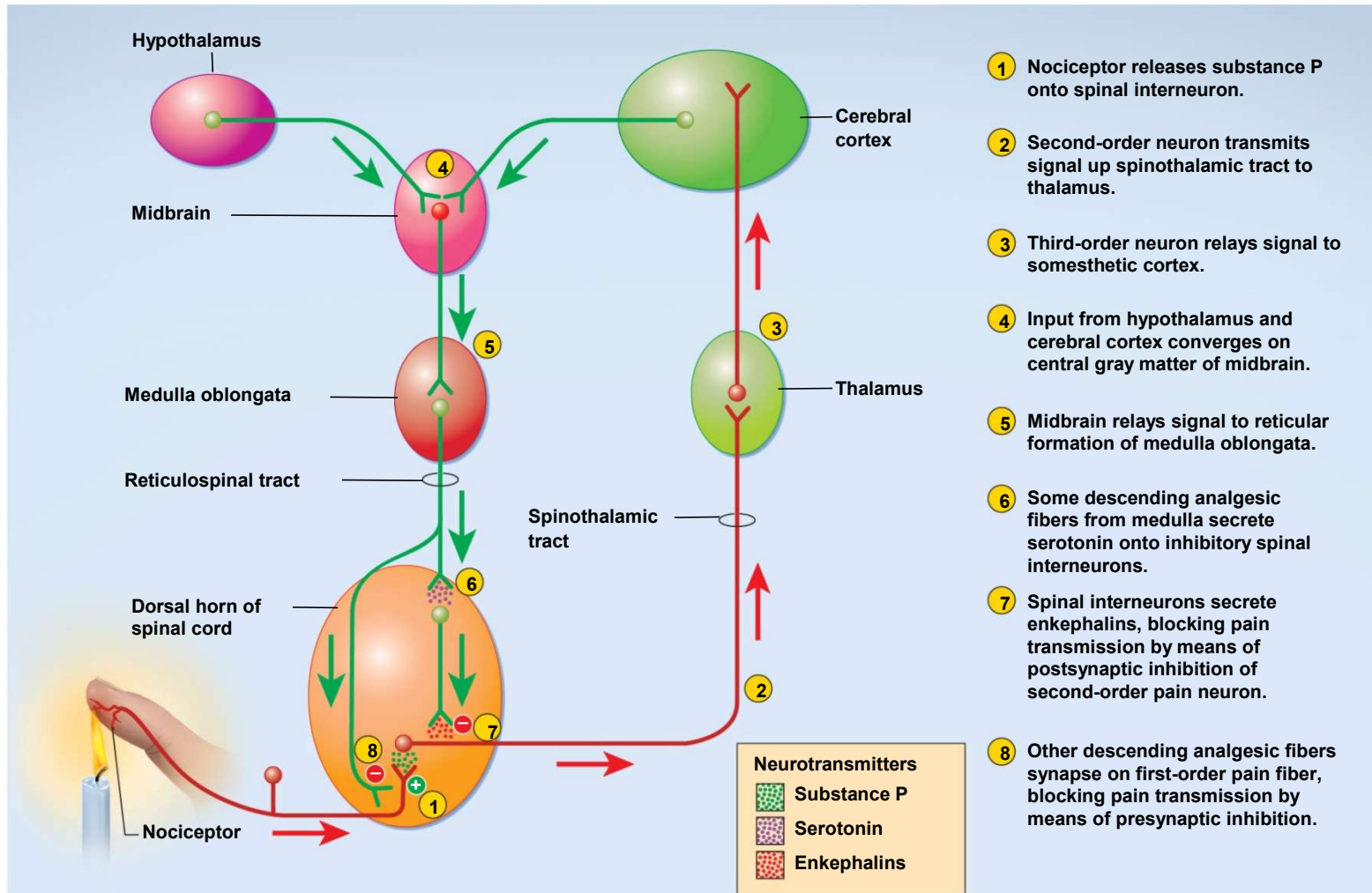
Spinal Gating

- Normal pain pathway (ascending)
 - **nociceptor** stimulates second-order nerve fiber
 - **substance P** is neurotransmitter at this synapse
 - second-order fiber transmits signal up the **spinothalamic tract** to the thalamus
 - **thalamus** relays the signals through third order neurons to the cerebral cortex where one becomes conscious of the pain

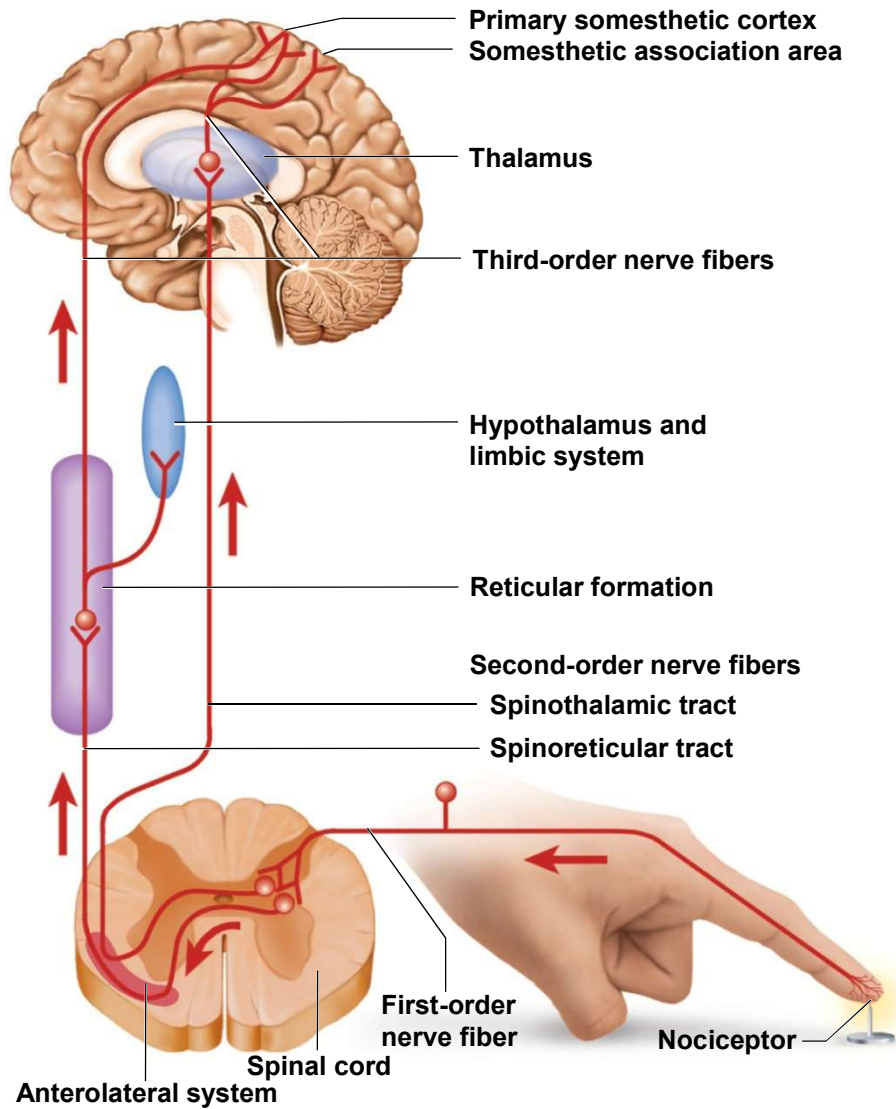
Spinal Gating: Pathway for Pain Blocking

- signals from the hypothalamus and cerebral cortex feed into the central gray matter of the midbrain // allows both autonomic and conscious influences on pain perception
- midbrain relays signals to certain nuclei in the reticular formation of the medulla oblongata
- medulla issues descending, serotonin-secreting analgesic fibers to the spinal cord // terminate in the posterior horn at all levels of the spinal cord
- in posterior horn, descending analgesic fibers synapse on short spinal interneurons (i.e. local circuit neurons)
- the interneurons synapse on the second-order pain fiber // secrete enkephalins to inhibit the second-order neuron
- some fibers from the medulla also exert presynaptic inhibition by synapsing on the axons of nociceptors and blocking the release of substance P

Spinal Gating of Pain Signals



Pain Signal Destinations (Ascending)



How to Interrupt Spinal Gating

- Rubbing or massaging injury
 - another pathway of spinal gating
 - pain-inhibiting neurons of the posterior horn receive input from mechanoreceptors in the skin and deeper tissues
 - rubbing stimulates mechanoreceptors which stimulates spinal interneurons to secrete enkephalins that inhibit second-order pain neurons