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## ***Neurotransmitters That Govern Hunger and Satiety***

There are many neurotransmitters and neuropeptides involved in the regulation of **hunger** and **satiety**. These signals primarily interact within hypothalamic circuits—especially the arcuate nucleus (ARC), paraventricular nucleus (PVN), and lateral hypothalamus (LH).

### **Neurotransmitters That Increase Hunger (Orexigenic)**

These neurotransmitters and related signals promote feeding behavior and increase appetite.

#### **1. Neuropeptide Y (NPY)**

- One of the most potent appetite-stimulating neurotransmitters.
- Released by NPY/AgRP neurons in the arcuate nucleus.
- Activated during fasting and energy deprivation.
- Increases food intake and reduces energy expenditure.

#### **2. Agouti-related peptide (AgRP)**

- Co-expressed with NPY; antagonizes melanocortin receptors (MC3R/MC4R).
- Strongly increases appetite.
- Activated when leptin is low (e.g., during weight loss).

#### **3. Ghrelin**

- A hormone from the stomach but acts as a neurotransmitter in the brain.
- The only known peripheral hormone that increases hunger.
- Stimulates NPY/AgRP neurons.
- Peaks before meals.

#### 4. Orexins (Hypocretins)

- Produced in the lateral hypothalamus.
- Increase appetite and also enhance arousal, wakefulness, and reward-driven eating.

#### 5. Dopamine

- Governs reward-driven eating (wanting food, cravings).
- Not a direct hunger signal but increases motivation to eat palatable foods.

### Neurotransmitters That Reduce Hunger (Anorexigenic / Satiety Signals)

These neurotransmitters and related signals suppress appetite, promote fullness, and support energy balance.

#### 1. Pro-opiomelanocortin (POMC) & $\alpha$ -MSH

- POMC neurons produce  $\alpha$ -MSH, which activates MC4 receptors to reduce appetite.
- Activated by leptin, insulin, and high energy stores.

#### 2. CART (Cocaine- and Amphetamine-Regulated Transcript)

- Co-expressed with POMC neurons.
- Suppresses appetite and increases energy expenditure.

#### 3. Serotonin (5-HT)

- Major satiety-promoting neurotransmitter.
- 5-HT<sub>2C</sub> receptor activation increases POMC activity.
- Drugs that raise serotonin (e.g., fenfluramine, lorcaserin) suppress appetite.

#### 4. Leptin

- A hormone from fat cells but acts centrally as a neurotransmitter-like signal.
- Inhibits NPY/AgRP neurons; activates POMC/CART neurons.
- Signals long-term energy status.

#### 5. Insulin

- From the pancreas; acts in the brain to reduce appetite.
- Increases POMC activity and decreases NPY/AgRP signaling.

#### 6. Cholecystokinin (CCK)

- Gut hormone acting via vagal afferents to produce meal-ending satiety.
- Quick-acting “stop eating now” signal.

## 7. GLP-1 (Glucagon-like peptide-1)

- Gut/brain peptide that increases satiety.
- Slows gastric emptying.
- Basis of GLP-1 weight-loss medications (e.g., semaglutide).

## 8. Peptide YY (PYY3-36)

- Released from the ileum/colon after meals.
- Inhibits NPY/AgRP neurons, reducing hunger.

## Simplified Summary

### Hunger Signals

- NPY
- AgRP
- Ghrelin
- Orexins
- Dopamine (reward-driven eating)

### Satiety Signals

- POMC /  $\alpha$ -MSH
- CART
- Serotonin
- Leptin
- Insulin
- CCK
- GLP-1
- PYY