Obesity and Biochemistry: Understanding the Molecular Causes of a Global Epidemic

We hear about obesity all the time, whether it's in the media or at the doctor's office, usually in connection with fast food, lack of exercise, and willpower. Most people begin discussions about weight with food choices and gym workouts, assuming those habits explain everything.

We're often led to believe that obesity comes down to personal choices. But in reality, it's not that simple. That narrative is appealing, but if we look more closely, we find that the problem is far more complex and deeply connected to biochemistry.

Leptin and insulin are hormones with significant roles in appetite control, energy use, and fat storage. They send chemical messages to the brain that help us know when we're full or when it's time to store energy instead of using it. If this messaging system breaks down, the brain begins to misinterpret the signals, and this is where many of our problems with appetite and obesity truly begin.

Leptin and Insulin Resistance

In a healthy body, leptin, produced by fat cells, signals the brain when there is enough stored energy. Insulin, released by the pancreas after meals, helps glucose enter cells and tells the brain that we are full. However, in many people with obesity, these systems no longer function as they should.

Take leptin, for example. One might assume that low levels of leptin lead to obesity, but in fact, many individuals with obesity have excessive amounts of this hormone. The issue is that the brain stops responding to leptin, similar to a phone set on silent mode. The signal is still present, but it goes unheard. As a result, even with abundant fat stores, the brain behaves as if the body is starving, leading to persistent hunger. This dysfunction sets the stage for the continuous accumulation of fat, with fat cells acting out of control. Insulin resistance occurs when the body no longer responds effectively to insulin. This results in elevated blood sugar levels. In response, the pancreas produces even more insulin, which leads to increased fat storage. This fat tends to accumulate in especially harmful areas, particularly around the abdomen and liver. This is not just surface-level belly fat. Fat surrounding the abdominal organs and liver is a key indicator of metabolic syndrome, a condition that affects about 34 per cent of adults in the United States.



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Cravings, Fat Signals, and the Brain

It is essential to understand that obesity is linked not only to hunger but also to food cravings. The types of food we consume, especially highly processed ones, are often engineered to be irresistible. These foods are not just unhealthy; they play a major role in the obesity epidemic. They are rich in refined flour and sugar, with artificial flavourings that disrupt normal brain chemistry.

Highly processed foods can interfere with neurotransmitters such as dopamine, which is involved in reward and motivation. These foods cause dopamine levels to spike, making us crave more even when we are full. Over time, this can create patterns that resemble addiction.

In addition, fat cells are not passive. They release substances called adipokines that contribute to inflammation and further disrupt hormonal signals. Visceral fat, which surrounds the organs, is particularly harmful. It can raise blood pressure, worsen insulin resistance, and lead to widespread inflammation.

Restoring Balance

The solution to obesity does not lie solely in eating less and exercising more. Instead, it involves restoring the body's internal balance. This includes improving the brain's response to hormones, reducing inflammation in the brain, and enhancing communication between organs through biochemical signals.

Future treatments focus on several areas. One is the development of leptin sensitisers, which help the brain respond to hunger signals again. Another is the use of anti-inflammatory therapies to reduce stress in the hypothalamus. A third approach involves targeting the gut microbiome, since gut bacteria influence how hormones are processed. Ultimately, viewing obesity through a biochemical lens allows us to rethink what is happening. It shifts the focus from personal blame to science-based solutions. When we understand how the body's internal systems are disrupted, we can begin to reset them. That brings us closer to a future that is not only healthier and happier but truly in balance.



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CITATIONS

Lustig, Robert H., et al. "Obesity I: Overview and Molecular and Biochemical Mechanisms." Biochemical Pharmacology, vol. 199, May 2022, p. 115012, https://doi.org/10.1016/j.bcp.2022.115012.

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