

## Hypoglycemia - Part Two

The next category of hypoglycemia to consider in the conventional model is hypoglycemia in patients with diabetes and impaired glucose tolerance.

### **Is it true hypoglycemia? Or postprandial syndrome?**

This typically occurs in the postprandial state rather than the fasted state. In conventional medicine, the presence of sympathoadrenal symptoms after meals isn't sufficient to diagnose hypoglycemia. The patient must also have blood glucose below 50 mg/dL within four hours of a meal, symptoms consistent with hypoglycemia, and relief of those symptoms when blood sugar is raised. So, again, we're talking about Whipple's triad.

However, the diagnosis of postprandial hypoglycemia is now controversial, and many clinicians and researchers believe that postprandial hypoglycemia is not a true hypoglycemic disorder. They now refer to it instead of postprandial syndrome.

The diagnostic workup is similar to what you do for fasting hypoglycemia in patients without diabetes. The goal is to document Whipple's triad and demonstrate low glucose below 50 mg/dL at the time symptoms are occurring, within four hours after a meal. The difference is patients with reactive hypoglycemia should be evaluated in the postprandial state after a mixed meal. OGTT, oral glucose tolerance test, is actually no longer recommended as the test for postprandial hypoglycemia for several reasons. One is that there is no correlation between blood glucose concentrations and symptoms during the test. Another is that many patients with postprandial sympathoadrenal symptoms have similar symptoms after the administration of placebo instead of glucose, and a third is that cortisol and epinephrine levels often do not increase when blood sugar levels are low or symptoms are present, which indicates that the fall in glucose is not sufficient for counterregulatory mechanisms to be activated.

While glucometers are not considered to be accurate—they have about a 10 to 15 percent variation per reading—they probably are the most practical way of doing this kind of testing.

Given that postprandial hypoglycemia most often occurs in the context of diabetes and other blood sugar disorders, the causes of it are similar to what we talked about in the hyperglycemia

presentation, so you can refer to that for more detail. However, in patients with diabetes who are taking medication, drug-induced hypoglycemia is something you have to consider. This can occur with a variety of medications as well as insulin, so make sure to take a full medication history during the intake.

| TESTS   | RESULT     | FLAG        | UNITS       | REFERENCE INTERVAL | LAB |
|---|------------|-------------|-------------|--------------------|-----|
| <b>CMP14+LP+TP+TSH+5AC+CBC/D/P...</b>   |            |             |             |                    |     |
| <b>Glucose, Serum</b>   | <b>45</b>  | <b>Low</b>  | mg/dL       | 65 - 99            | 01  |
| Specimen received in contact with cells. No visible hemolysis present. However GLUC may be decreased and K increased. Clinical correlation indicated. |            |             |             |                    |     |
| <b>Hemoglobin A1c</b>   | <b>6.0</b> | <b>High</b> | %           | 4.8 - 5.6          | 01  |
| Increased risk for diabetes: 5.7 - 6.4  |            |             |             |                    |     |
| Diabetes: >6.4  |            |             |             |                    |     |
| Glycemic control for adults with diabetes: <7.0   |            |             |             |                    |     |
| Uric Acid, Serum  | 5.7        |             | mg/dL       | 3.7 - 8.6          | 01  |
| Please Note:  |            |             |             |                    | 01  |
| Therapeutic target for gout patients: <6.0  |            |             |             |                    |     |
| BUN   | 14         |             | mg/dL       | 6 - 24             | 01  |
| Creatinine, Serum   | 0.95       |             | mg/dL       | 0.76 - 1.27        | 01  |
| eGFR If NonAfrican Am   | 96         |             | mL/min/1.73 | >59                |     |
| eGFR If African Am  | 111        |             | mL/min/1.73 | >59                |     |
| BUN/Creatinine Ratio  | 15         |             |             | 9 - 20             |     |
| <b>Sodium, Serum</b>  | <b>145</b> | <b>High</b> | mmol/L      | 134 - 144          | 01  |
| Potassium, Serum  | 3.8        |             | mmol/L      | 3.5 - 5.2          | 01  |
| Chloride, Serum   | 102        |             | mmol/L      | 97 - 108           | 01  |
| Carbon Dioxide, Total   | 24         |             | mmol/L      | 18 - 29            | 01  |
| Calcium, Serum  | 8.8        |             | mg/dL       | 8.7 - 10.2         | 01  |
| Phosphorus, Serum   | 3.6        |             | mg/dL       | 2.5 - 4.5          | 01  |
| Magnesium, Serum  | 2.2        |             | mg/dL       | 1.6 - 2.6          | 01  |
| Protein, Total, Serum   | 6.3        |             | g/dL        | 6.0 - 8.5          | 01  |
| Albumin, Serum  | 4.3        |             | g/dL        | 3.5 - 5.5          | 01  |
| Globulin, Total   | 2.0        |             | g/dL        | 1.5 - 4.5          |     |
| A/G Ratio   | 2.2        |             |             | 1.1 - 2.5          |     |
| Bilirubin, Total  | 0.8        |             | mg/dL       | 0.0 - 1.2          | 01  |
| Alkaline Phosphatase, S   | 63         |             | IU/L        | 39 - 117           | 01  |
| LDH   | 154        |             | IU/L        | 121 - 224          | 01  |
| AST (SGOT)  | 16         |             | IU/L        | 0 - 40             | 01  |
| ALT (SGPT)  | 20         |             | IU/L        | 0 - 44             | 01  |
| GGT   | 19         |             | IU/L        | 0 - 65             | 01  |

In practice, you may see mixed cases where fasting glucose is low, but other blood sugar markers indicate hyperglycemia, so the lab on this slide is a good example. Patient was a 47-year-old male with loose stools, abdominal pain, weight loss, and low energy. He was unaware of any blood sugar issue when he came to see me, and as you can see, his fasting glucose was 45, which is very low, but his hemoglobin A1c was 6.0. Now we know that hemoglobin A1c is not necessarily a reliable marker, but in this case, some of his post-meal blood sugar readings were high, and some of them were low, so his blood sugar was fluctuating wildly. This presentation can occur during the early stages of type 2 diabetes or type 1.5 diabetes, which is adult-onset autoimmune diabetes.

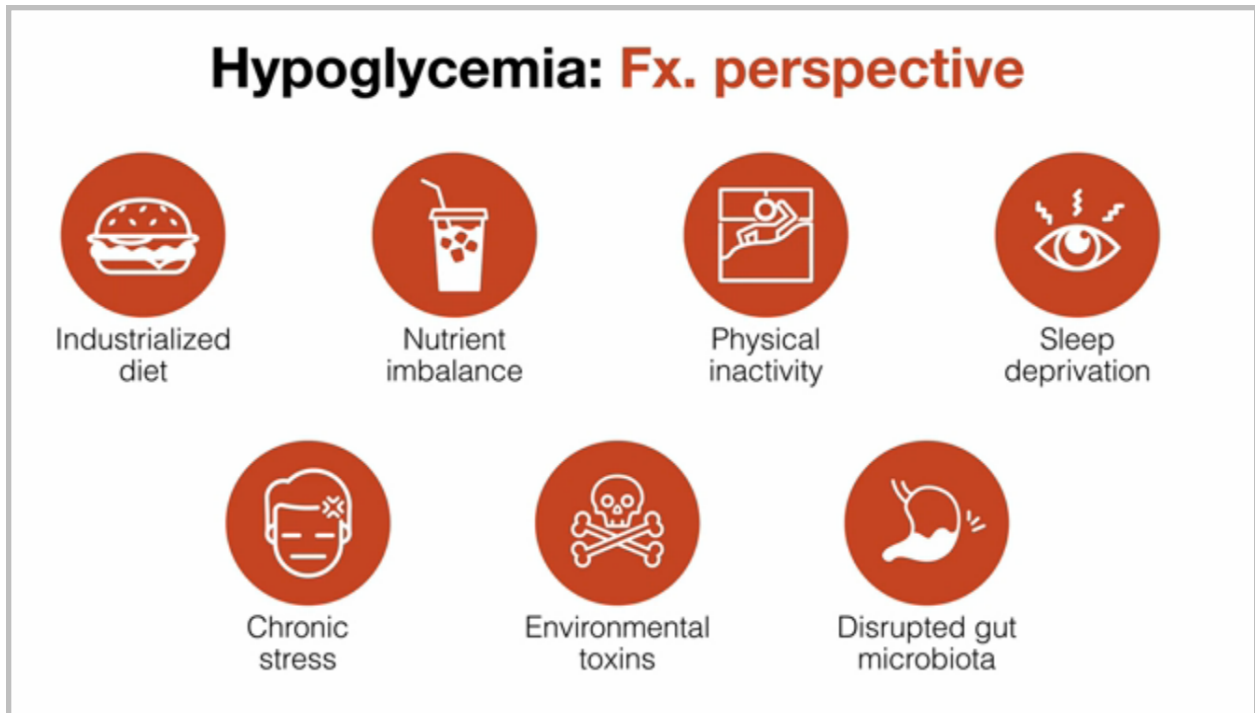
| DAY ONE                    |  |        |  |
|----------------------------|--|--------|--|
| # of hours fasted          | 12   |        |  |
| # of hours slept           | 8  |        |  |
| quality of sleep           | Good. Woke up 1 time   |        |  |
| what you ate for breakfast | Yams, white rice, porridge, 1 egg with no yolk,                    |        |  |
| what you ate for lunch     | White rice, boiled chicken, cabbage, celery, napa cabbage, carrots |        |  |
|                            | Time   | Result | How did you feel at the time of measurement (2-3 words)                                  |
| AM Fasting                 | 8:50am   | 96     | Good   |
| Before Lunch               | 12pm   | 111    | Good. Rested at home, didn't go out.   |
| After Lunch (45-min)       | 1pm  | 110    | Good. Rested at home, didn't go out.   |
| After Lunch (1-hour later) | 2:10pm   | 93     | Good. Rested at home, didn't go out  |
| After Lunch (1-hour later) | 3:10pm   | 80     | Good. Rested at home, didn't go out.   |
| DAY TWO                    |  |        |  |
| # of hours fasted          | 12   |        |  |
| # of hours slept           | 8  |        |  |
| quality of sleep           | average. not as well as yesterday                                  |        |  |
| what you ate for breakfast | yams, white rice, porridge, 1 egg with no yolk.                    |        |  |
| what you ate for lunch     | white rice, boiled chicken, cabbage, celery, napa cabbage, carrots |        |  |
|                            | Time   | Result | How did you feel at the time of measurement (2-3 words)                                  |
| AM Fasting                 | 8:15am   | 88     | Good <span style="float: right;"><b>Good</b></span>                                      |
| Before Lunch               | 12:10pm  | 77     | Hungry. <span style="float: right;"><b>Hungry.</b></span>                                |
| After Lunch (45-min)       | 1:15pm   | 134    |  |
| After Lunch (1-hour later) | 2:15pm   | 88     |  |
| After Lunch (1-hour later) | 3:15pm   | 84     |  |
| DAY THREE                  |  |        |  |
| # of hours fasted          | 12   |        |  |
| # of hours slept           | 8  |        |  |
| quality of sleep           | Not good. Woke up at 4-5am. Noisy neighbors                        |        |  |
| what you ate for breakfast | yams, white rice, porridge, 1 egg with no yolk.                    |        |  |
| what you ate for lunch     | white rice, boiled chicken, cabbage, celery, napa cabbage, carrots |        |  |
|                            | Time   | Result | How did you feel at the time of measurement (2-3 words)                                  |
| AM Fasting                 | 8am  | 69     | Hungry. Did not eat <del>to go to bed</del> <del>last night</del> for dinner last night. |
| Before Lunch               | 12pm   | 59     |  |
| After Lunch (45-min)       | 1:10pm   | 82     |  |
| After Lunch (1-hour later) | 2:10pm   | 85     |  |
| After Lunch (1-hour later) | 3:10pm   | 90     |  |

Here is the glucometer testing for this patient. Fasting blood sugar is highly variable, as you can see. It was 96 one day; 69 another day, which is borderline low on the functional range; 59 one day before lunch. Then one day after lunch, it went up to 134, which is not outside of the target, but it's getting up there. This patient moved recently in the past five years to the U.S. from China and ate a traditional Chinese diet. As you can see here on the page, this is why this form is so helpful. You ask the patient what he eats, and you see how his blood sugar responds to what he eats. In this case, given the amount of white rice that he eats and his blood sugar fluctuations, it's probably a good idea for him not to do that and to see how his blood sugar changes or stabilizes.

| Marker                    | Value | Functional Range | Lab Range    |
|---------------------------|-------|------------------|--------------|
| Glucose                   | 45    | 75 - 90          | 65 - 99      |
| Hemoglobin A1c            | 6.0   | 4.4 - 5.4        | 4.8 - 5.6    |
| Uric Acid                 | 5.7   | 3.7 - 6.0        | 3.7 - 8.6    |
| BUN                       | 14    | 13 - 18          | 6 - 24       |
| Creatinine                | 0.95  | 0.85 - 1.1       | 0.76 - 1.27  |
| BUN/Creatinine Ratio      | 15    | 8 - 19           | 9 - 20       |
| Sodium                    | 145   | 135 - 140        | 134 - 144    |
| Potassium                 | 3.8   | 4.0 - 4.5        | 3.5 - 5.2    |
| Chloride                  | 102   | 100 - 106        | 97 - 108     |
| C02                       | 24    | 25 - 30          | 18 - 29      |
| Calcium                   | 8.8   | 9.2 - 10.1       | 8.7 - 10.2   |
| Phosphorus                | 3.6   | 3.5 - 4.0        | 2.5 - 4.5    |
| Magnesium                 | 2.2   | 2.0 - 2.6        | 1.6 - 2.6    |
| Protein, total            | 6.3   | 6.9 - 7.4        | 6.0 - 8.5    |
| Albumin                   | 4.3   | 4.0 - 5.0        | 3.5 - 5.5    |
| Globulin                  | 2.0   | 2.4 - 2.8        | 1.5 - 4.5    |
| A/G ratio                 | 2.2   | 1.5 - 2.0        | 1.1 - 2.5    |
| Bilirubin, total          | 0.8   | 0.1 - 1.2        | 0.0 - 1.2    |
| Alkaline Phosphatase      | 63    | 42 - 107         | 39 - 117     |
| LDH                       | 154   | 140 - 180        | 121 - 224    |
| AST                       | 16    | 10 - 30          | 0 - 40       |
| ALT                       | 20    | 10 - 29          | 0 - 44       |
| GGT                       | 19    | 0 - 40           | 0 - 65       |
| TIBC                      | 245   | 250 - 350        | 250 - 450    |
| UIBC                      | 134   | 150 - 375        | 150 - 375    |
| Iron                      | 111   | 85 - 135         | 40 - 155     |
| Iron saturation           | 45    | 15 - 45          | 15 - 55      |
| Ferritin                  | 372   | 30 - 150         | 30 - 400     |
| Cholesterol, total        | 127   | 150 - 240        | 100 - 199    |
| Triglycerides             | 77    | 50 - 100         | 0 - 149      |
| HDL                       | 39    | 55 - 85          | > 39         |
| LDL                       | 73    | 0 - 175          | 0 - 99       |
| T. Chol / HDL Ratio       | 3.3   | < 3              | 0 - 5.0      |
| Triglycerides / HDL Ratio | 1.97  | < 2              |              |
| TSH                       | 1.290 | 0.5 - 2.5        | 0.45 - 4.50  |
| T4, total                 | 6.1   | 6.0 - 12         | 4.5 - 12     |
| T3 Uptake                 | 34    | 30 - 38          | 24 - 39      |
| T3, Total                 | 103   | 100 - 180        | 71 - 180     |
| Vitamin D, 25-hydroxy     | 29.1  | 35 - 60          | 30.0 - 100.0 |

This patient also had low vitamin D and iron overload, both of which can contribute to dysglycemia, as you know. Sodium was slightly high, and that can be caused by dehydration. It's likely unrelated to blood sugar.

Now let's talk about hypoglycemia from a functional perspective. We look at it differently in a number of ways. The first difference is how we view the etiology and pathogenesis. If you go to UpToDate or a similar resource online, it lists the causes of hypoglycemia as what I mentioned earlier: drugs, alcohol, critical illness, malnourishment, tumor, etc. There is no mention of the contribution of modern lifestyle, which is absolutely fundamental, in my opinion. The causes of hypoglycemia, especially postprandial, are virtually identical to the causes of hyperglycemia that were covered in the last presentation.



It's not that the conventional causes aren't valid. They are, but the functional perspective is broader and more inclusive. Many of your patients with hypoglycemia won't have any of the conventional causes at work, but they will have some or most of the ones on this slide. These include industrialized diet, nutrient imbalance, physical activity, sleep deprivation, chronic stress, environmental toxins, and disrupted gut microbiota.

Another difference between hypoglycemia from the conventional and functional perspectives is how it is defined. In conventional medicine, the patient must meet the strict definition of Whipple's triad, which again are symptoms consistent with hypoglycemia, a low plasma glucose concentration measured with a precise method when symptoms are occurring, and then relief of those symptoms after blood sugar is raised. In functional medicine, the definition is similar, but I'm less concerned with seeing a glucose level below the lab reference range. If I see a patient who has symptoms that are consistent with hypoglycemia, and those symptoms are resolved by eating, what do we do if their fasting blood sugar is 70? We should still rule out other conditions that can cause these symptoms, but if we do, I think it still makes sense to treat this patient as if he may

have a blood sugar disorder. This is especially true since treatment in functional medicine involves primarily dietary, lifestyle, and behavioral tweaks and possibly supplements and botanicals that are not likely to have significant side effects. In other words, if someone who didn't have hypoglycemia followed the functional treatment for it, he would likely feel better or unchanged, not worse, so there is very little risk in treatment even if the person doesn't have true hypoglycemia according to the conventional definition.

That said, I wouldn't diagnose functional hypoglycemia based on lab results alone. Those results should accompany symptoms, and those symptoms should resolve with interventions that regulate blood sugar. For example, if a patient has a fasting blood sugar of 70, his other blood sugar markers are normal, and he doesn't have any symptoms of hypoglycemia, I'm not going to be worried about that. I would just say that is good blood sugar control.