THE CASE
A 6-year-old girl was brought to the emergency department (ED) by her mother after the child had bumped her head while playing. While the physician examined the child’s head, the mother remarked that her daughter had recently developed bruises that appeared suddenly and only after minor, if any, known trauma. The ED physician determined that the child’s bump to the head was nothing to worry about, attributed the bruising to the child being a “healthy, active 6-year-old,” and sent her home.

Two days later the child was brought to our office because the mother was still concerned about her daughter’s easy bruising. The mother pointed out ecchymosis scattered across her daughter’s extremities and torso. The child denied any pain or other complaints, including any active or recurrent bleeding. Upon further questioning, the mother mentioned that her daughter had recovered from a cold-like illness several weeks earlier.

THE DIAGNOSIS
We ordered a complete blood count (CBC) and peripheral smear, which were normal except for the platelet count, which was 7000/mcL (normal, 150,000-450,000/mcL). Based on the child’s easy bruising and isolated thrombocytopenia, we diagnosed immune thrombocytopenia, which is also known as idiopathic thrombocytopenic purpura (ITP).

DISCUSSION
In ITP, autoantibodies are directed against platelets, leading to their sequestration and destruction in the spleen and a resultant drop in platelet count.1 Children with ITP typically present between the ages of 2 and 10 years, with a peak incidence between 2 and 5 years.2 The incidence is estimated to be as high as 8 per 100,000 children.3 However, this estimate primarily reflects symptomatic children, and the true incidence of childhood ITP may be much higher because asymptomatic children may not be brought in to see a doctor. For the majority of patients, ITP resolves within 3 months. However, for 20% to 30% of patients, thrombocytopenia will last beyond 6 months, with or without treatment.4 In 1% of cases, patients will have a recurrence of ITP.3

In addition to easy bruising, nearly all patients who present with possible ITP will complain of cutaneous bleeding, typically a nose bleed or bleeding in the oral cavity.2 Upon questioning, 60% of patients will report a history of recent infection.4 Not surprisingly, bleeding severity correlates inversely with platelet count; severe bleeding is seen in patients with a platelet count <10,000/mcL.

While rare, the more worrisome complications include intracranial hemorrhage, with an incidence of 0.1% to 0.8%, and other serious hemorrhages that would require transfusion, with an estimated incidence of 2.9%.2
The incidence of ITP may be higher during the winter months, when infections are more common.

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**Vast differential seen in child bruising**

When a child presents with bruising, perform a thorough history, including birth and prenatal course, as well as a physical to exclude other potential causes, such as physical abuse, use of herbal remedies or other natural supplements that may not be disclosed as medication, or even environmental exposure. When bruising is present in a child who has isolated thrombocytopenia, the diagnosis of ITP may be straightforward. However, many conditions may share thrombocytopenia in their disease process and should be considered in the differential diagnosis of a child who you suspect may have ITP.

- **Suspect physical abuse** in a bruised child who does not have thrombocytopenia, whose mood is flat or depressed, or who has experienced recurrent injuries or bruising.

- **Leukemia**, particularly acute lymphoblastic leukemia (ALL), the predominant leukemia found in children, should be ruled out, as well. Symptoms that may distinguish a child with ALL from one with ITP include fever, weight loss, and joint pain, as well as signs such as lymphadenopathy, hepatosplenomegaly, anemia, and leukocytosis. A peripheral smear may be ordered to help confirm or exclude a diagnosis of ALL should any of the above be present in a child with thrombocytopenia. It may show lymphoblasts and/or atypical cells in a patient with ALL.

- **Infections** should also be included in a differential when a patient is suspected of having ITP, particularly if he or she has systemic symptoms. Viral infections that may cause thrombocytopenia include mononucleosis, dengue virus, human herpesvirus-6, and human immunodeficiency virus.

ITP often follows an infection, and the incidence of ITP may be higher during winter months, when infections are more common. However, infection may not always be the cause of ITP. Sepsis may also lead to thrombocytopenia, but a child with sepsis would present very differently from a child who has only ITP. A septic child would present acutely ill with signs and symptoms of severe systemic illness, such as high fever, altered mental status, tachycardia, pallor, diaphoresis, and hypotension.

- **Drug-induced thrombocytopenia (DIT)** should be considered in any child who is taking or recently took a medication that may cause thrombocytopenia. Medications that can cause thrombocytopenia include heparin, quinine, vancomycin, trimethoprim-sulfamethoxazole, rifampin, carbamazepine, phenytoin, piperacillin, linezolid, and valproic acid. The measles, mumps, and rubella vaccine also can cause thrombocytopenia. A careful medication history may determine if the child is at risk for DIT.

- **To narrow the differential**, obtain a CBC and peripheral smear when evaluating a patient you suspect may have ITP (strength of recommendation [SOR]: A). A CBC will determine the patient’s platelet count and a peripheral smear should be obtained to exclude other possible diagnoses.

If there are any questions regarding the results of a peripheral smear, it may be necessary to perform a bone marrow aspiration. This, however, is not usually necessary in an otherwise typical case of ITP. Bone marrow aspiration may, however, be necessary to re-evaluate the initial diagnosis for a child who does not respond to treatment for ITP.

**Corticosteroids, IVIg are usually effective**

The first step in treating a patient with ITP is to limit the risk of further injury or bleeding, by stopping nonsteroidal anti-inflammatory drugs or ending participation in contact sports (SOR: C). The next step is to determine if pharmacologic therapy is warranted.

Medication, if necessary, is the mainstay of treatment for patients with ITP, particularly those experiencing significant bleeding. Corticosteroids, intravenous (IV) immunoglobulin (IVIg), and IV Rh(D) immune globulin (also known as anti-D) are the medications typically used to treat a child with ITP.

**Strength of recommendation (SOR)**

- **A** Good-quality patient-oriented evidence
- **B** Inconsistent or limited-quality patient-oriented evidence
- **C** Consensus, usual practice, opinion, disease-oriented evidence, case series
To start ITP treatment, limit the patient’s risk of further injury or bleeding by stopping NSAIDs and ending participation in contact sports.

<table>
<thead>
<tr>
<th>Treatment</th>
<th>Response rate</th>
<th>Adverse effects</th>
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<tbody>
<tr>
<td>IV Rh(D) 50-75 mcg/kg</td>
<td>50%-77% of patients achieve a platelet response, depending on dose</td>
<td>Headache, fever, chills (less common than with IVig)</td>
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<tr>
<td>IVlg 800 mg/kg on Day 1</td>
<td>Effective in &gt;80% of patients</td>
<td>Headache (which can be severe), fever</td>
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<tr>
<td>Prednisone 1-2 mg/kg/d for a maximum of 14 days, or 4 mg/kg/d for 3-4 days</td>
<td>Up to 75% of patients will respond, depending on dose</td>
<td>Transient mood changes, gastritis, and weight gain. Exercise caution in active infection</td>
</tr>
<tr>
<td>Watch and wait (with activity restriction)</td>
<td>Approximately 66% of ITP patients will improve spontaneously within 6 months</td>
<td>Hemorrhage (preventable), anxiety</td>
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ITP, idiopathic thrombocytopenic purpura; IV, intravenous; IVlg, intravenous immunoglobulin.
Soon after, she went to our ED with a headache, nausea, and fever of 102°F. A computed tomography scan of her head was normal; a repeat CBC showed no elevation in white blood cells but her hemoglobin had decreased from 11.9 g/dL to 9.7 g/dL. (Her platelets were 254,000/mcL.) The patient’s complaints were likely adverse effects of the IVIg. The CBC abnormalities, fever, headache, and malaise resolved shortly thereafter and the patient remains asymptomatic with no recurrence of ITP.

THE TAKEAWAY
Suspect ITP in a child who bruises easily and who also has thrombocytopenia. Order a CBC and peripheral blood smear to rule out other potential illnesses. Pharmacotherapy, if needed, typically consists of an oral or IV corticosteroid or IVIg; IV Rh(D) Ig may be used in patients who are Rh(D)-positive who don’t respond to other treatments. Patients with ITP should have their platelet count monitored at least once weekly until platelets have increased to 150,000/mcL or higher. Frequency of monitoring may be reduced as the clinical picture improves and the patient remains stable. More frequent monitoring may be necessary based on severity, complications, and response to treatment.

References