Diagnosing the Cause of Chest Pain

WILLIAM E. CAYLEY, JR., M.D., Eau Claire Family Medicine Residency, Eau Claire, Wisconsin

Chest pain presents a diagnostic challenge in outpatient family medicine. Noncardiac causes are common, but it is important not to overlook serious conditions such as an acute coronary syndrome, pulmonary embolism, or pneumonia. In addition to a thorough history and physical examination, most patients should have a chest radiograph and an electrocardiogram. Patients with chest pain that is predictably exertional, with electrocardiogram abnormalities, or with cardiac risk factors should be evaluated further with measurement of troponin levels and cardiac stress testing. Risk of pulmonary embolism can be determined with a simple prediction rule, and a D-dimer assay can help determine whether further evaluation with helical computed tomography or venous ultrasound is needed. Fever, egophony, and dullness to percussion suggest pneumonia, which can be confirmed with chest radiograph. Although some patients with chest pain have heart failure, this is unlikely in the absence of dyspnea; a brain natriuretic peptide level measurement can clarify the diagnosis. Pain reproducible by palpation is more likely to be musculoskeletal than ischemic. Chest pain also may be associated with panic disorder, for which patients can be screened with a two-item questionnaire. Clinical prediction rules can help clarify many of these diagnoses. (Am Fam Physician 2005;72:2012-21. Copyright © 2005 American Academy of Family Physicians.)
Determining whether chest pain is anginal, atypical anginal, or non-anginal is recommended to help determine a patient’s cardiac risk.

The Rouan decision rule is recommended to help predict which patients are at higher risk of MI.

A Wells score of less than 2 plus a normal D-Dimer assay should rule out PE.

In patients with an abnormal D-Dimer assay or a Wells score indicating moderate to high risk, helical CT and lower extremity venous ultrasound examination should be used to rule in or rule out PE.

The Diehr diagnostic rule is recommended to predict the likelihood of pneumonia based on clinical findings.

Patients should be screened for panic disorder using two set questions. Patients presenting with chest pain should have an ECG evaluation for ST segment elevation, Q waves, and conduction defects. Results should be compared with previous tracings.

Serum troponin–level testing is recommended to aid in the diagnosis of MI and help predict the likelihood of death or recurrent MI within 30 days.

Patients with chest pain and a negative initial cardiac evaluation should have further testing with stress ECG, perfusion scanning, or angiography depending on their level of risk.

The Duke treadmill score is recommended to help predict long-term prognosis for patients undergoing stress ECG testing.

**MI = myocardial infarction; PE = pulmonary embolism; CT = computed tomography; ECG = electrocardiography.**

A = consistent, good-quality patient-oriented evidence; B = inconsistent or limited-quality patient-oriented evidence; C = consensus, disease-oriented evidence, usual practice, expert opinion, or case series. For information about the SORT evidence rating system, see page 1949 or http://www.aafp.org/afpsort.xml.

### TABLE 1
Epidemiology of Chest Pain in Primary Care and Emergency Department Settings

<table>
<thead>
<tr>
<th>Diagnosis*</th>
<th>Primary care: United States</th>
<th>Primary care: Europe</th>
<th>Emergency department</th>
</tr>
</thead>
<tbody>
<tr>
<td>Musculoskeletal condition</td>
<td>36</td>
<td>29</td>
<td>7</td>
</tr>
<tr>
<td>Gastrointestinal disease</td>
<td>19</td>
<td>10</td>
<td>3</td>
</tr>
<tr>
<td>Serious cardiovascular disease†</td>
<td>16</td>
<td>13</td>
<td>54</td>
</tr>
<tr>
<td>Stable coronary artery disease</td>
<td>10</td>
<td>8</td>
<td>13</td>
</tr>
<tr>
<td>Unstable coronary artery disease</td>
<td>1.5</td>
<td>—</td>
<td>13</td>
</tr>
<tr>
<td>Psychosocial or psychiatric disease</td>
<td>8</td>
<td>17</td>
<td>9</td>
</tr>
<tr>
<td>Pulmonary disease‡</td>
<td>5</td>
<td>20</td>
<td>12</td>
</tr>
<tr>
<td>Nonspecific chest pain</td>
<td>16</td>
<td>11</td>
<td>15</td>
</tr>
</tbody>
</table>

*—Diagnoses are listed in order of prevalence in United States.
†—Including infarction, unstable angina, pulmonary embolism, and heart failure.
‡—Including pneumonia, pneumothorax, and lung cancer.

Adapted with permission from Klinkman MS, Stevens D, Gorenflo DW. Episodes of care for chest pain: a preliminary report from MIRNET. J Fam Pract 1994;38:349, with additional information from reference 3.
consistent finding and should not be relied upon. There is enough overlap among the clinical manifestations of different causes of chest pain to make “classic” symptoms unhelpful in differentiating among diagnoses and ruling out serious causes. However, there are several validated clinical decision rules that combine key groups of symptoms.

**HISTORY AND PHYSICAL EXAMINATION**

It is important to obtain a clear history of the onset and evolution of chest pain, with particular attention to details such as location, quality, duration, and aggravating or alleviating factors. Certain key symptoms and clinical findings can help rule in or out specific diagnoses (Table 2).4-15

Determining whether pain is (1) substernal, (2) provoked by exertion, or (3) relieved by rest or nitroglycerin helps to clarify whether it is typical anginal pain (has all three characteristics), atypical anginal pain (has two characteristics), or nonanginal

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**The Author**

WILLIAM E. CAYLEY, JR., M.D., M.DIV., is assistant professor at the Eau Claire (Wis.) Family Medicine Residency, University of Wisconsin, Eau Claire. He received his medical degree from the Medical College of Wisconsin, Milwaukee, and completed a residency at the Eau Claire Family Medicine Residency, Eau Claire.

Address correspondence to William E. Cayley, Jr., M.D., M.Div., Eau Claire Family Medicine Residency, 617 W. Clairemont, Eau Claire, WI 54701 (e-mail: bcayley@yahoo.com). Reprints are not available from the author.
Chest Pain Dx

The most common causes of chest pain seen in outpatient primary care are musculoskeletal conditions, gastrointestinal disease, stable coronary artery disease, panic disorder or other psychiatric conditions, and pulmonary disease.

At higher risk for MI (Table 3). However, because up to 3 percent of patients initially diagnosed with a noncardiac cause of chest pain suffer death or MI within 30 days of presentation, patients with cardiac risk factors such as male sex, greater age, diabetes, hyperlipidemia, prior CAD, or heart failure warrant close follow-up.

There are no individual signs or symptoms that reliably diagnose PE, but the simplified Wells scoring system (Table 4) is well validated for determining whether patients have low, moderate, or high likelihood of PE, and this guides further evaluation.

Findings that suggest pneumonia include fever, egophony, and dullness to percussion, but their absence does not rule out the diagnosis. Although chest pain in patients with chronic obstructive pulmonary disease and at least four previous acute exacerbations of chronic bronchitis is more likely to be caused by a recurrent exacerbation of bronchitis or pneumonia, these patients are also at greater risk for CAD or acute coronary syndrome. The Diehr diagnostic rule, developed in a large study from 1984, uses seven clinical findings to predict the likelihood of pneumonia.

Although heart failure alone is an uncommon cause of chest pain, it may accompany acute coronary syndrome, valvular disease, or MI. A displaced apical impulse and a history of MI also support this diagnosis. Almost all patients with heart failure have exertional dyspnea, so the absence of exertional dyspnea is helpful in ruling out this diagnosis.

Two simple questions are a highly sensitive screen for panic disorder:

• “In the past six months, did you ever have a spell or an attack when all of a sudden you felt frightened, anxious, or very uneasy?”
• “In the past six months, did you ever have a spell or an attack when for no reason your heart suddenly began to race, you felt faint, or you couldn’t catch your breath?”

A “yes” on either item is a positive screen.
Chest Pain Dx and a “no” on both items makes panic disorder unlikely. However, neither these questions nor a general clinical impression are specific enough to allow a definite diagnosis of anxiety-related noncardiac chest pain, and a positive screen should not preclude further cardiac testing in patients with cardiac risk factors.19

Gastrointestinal disease can cause chest pain, but the history and physical examination are relatively inaccurate for ruling in or ruling out serious gastrointestinal pathology,24 and it is important first to rule out immediately life-threatening cardiovascular and pulmonary causes of chest pain.

Diagnostic Testing

Once the clinical examination has narrowed the differential diagnosis, diagnostic testing helps determine whether the patient has a serious condition (Table 6).4,7,12,25,26 Most adults with chest pain should have at least an ECG and a chest radiograph, unless the history and physical examination suggest an obviously nonthreatening cause of chest discomfort.

**ACUTE CORONARY SYNDROME AND CAD**

Important diagnostic tests when evaluating for acute coronary syndrome include the 12-lead ECG, serum markers of myocardial damage, and cardiac testing with stress testing or nuclear imaging. ECG findings that most strongly suggest MI are ST segment elevation, Q waves, and a conduction defect, especially if such findings are new compared with a previous ECG. New T-wave inversion also increases the likelihood of MI.7,9 However, none of these findings is sensitive enough that its absence can exclude MI.

The most common markers of myocardial damage are creatine kinase, the MB isoenzyme of creatine kinase (CK-MB), troponin T, and troponin I. A CK-MB level greater than 6.0 ng per mL (6.0 mcg per L) within nine hours of presentation for emergency care modestly increases the likelihood of MI or death in the next 30 days.27

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**TABLE 4**

**Wells Model for Clinical Diagnosis of Pulmonary Embolism**

<table>
<thead>
<tr>
<th>Clinical finding*</th>
<th>Points</th>
</tr>
</thead>
<tbody>
<tr>
<td>Clinical signs and symptoms of DVT (i.e., objectively measured leg swelling or pain with palpation of deep leg veins)</td>
<td>3.0</td>
</tr>
<tr>
<td>PE as likely or more likely than an alternative diagnosis</td>
<td>3.0</td>
</tr>
<tr>
<td>Heart rate more than 100 beats per minute</td>
<td>1.5</td>
</tr>
<tr>
<td>Immobilization (i.e., bedrest except for bathroom access for at least three consecutive days) or surgery in the past four weeks</td>
<td>1.5</td>
</tr>
<tr>
<td>Previous objectively diagnosed DVT or PE</td>
<td>1.5</td>
</tr>
<tr>
<td>Malignancy (treatment for cancer that is ongoing, within the past six months, or palliative)</td>
<td>1.0</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Total points</th>
<th>Risk of PE</th>
<th>Probability of PE (%)&lt;sup&gt;21&lt;/sup&gt;</th>
</tr>
</thead>
<tbody>
<tr>
<td>&lt;2 points</td>
<td>Low</td>
<td>0.13</td>
</tr>
<tr>
<td>2 to 6 points</td>
<td>Moderate</td>
<td>1.82</td>
</tr>
<tr>
<td>&gt;6 points</td>
<td>High</td>
<td>6.75</td>
</tr>
</tbody>
</table>

<sup>*—Findings are listed in order of clinical importance.

DVT = deep venous thrombosis; PE = pulmonary embolism; LR+ = positive likelihood ratio.


**TABLE 5**

**Diehr Rule for Diagnosing Pneumonia in Adults with Acute Cough**

<table>
<thead>
<tr>
<th>Finding</th>
<th>Points</th>
</tr>
</thead>
<tbody>
<tr>
<td>Rhinorrhea</td>
<td>–2</td>
</tr>
<tr>
<td>Sore throat</td>
<td>–1</td>
</tr>
<tr>
<td>Myalgia</td>
<td>1</td>
</tr>
<tr>
<td>Night sweats</td>
<td>1</td>
</tr>
<tr>
<td>Sputum all day</td>
<td>1</td>
</tr>
<tr>
<td>Respiratory rate more than 25 breaths per minute</td>
<td>2</td>
</tr>
<tr>
<td>Temperature more than 100ºF (37.8ºC)</td>
<td>2</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Total points</th>
<th>Probability of pneumonia (%; overall probability = 3)</th>
</tr>
</thead>
<tbody>
<tr>
<td>–3</td>
<td>0.0</td>
</tr>
<tr>
<td>–2</td>
<td>0.7</td>
</tr>
<tr>
<td>–1</td>
<td>1.6</td>
</tr>
<tr>
<td>0</td>
<td>2.2</td>
</tr>
<tr>
<td>1</td>
<td>8.8</td>
</tr>
<tr>
<td>2</td>
<td>10.3</td>
</tr>
<tr>
<td>3</td>
<td>25.0</td>
</tr>
<tr>
<td>≥4</td>
<td>29.4</td>
</tr>
</tbody>
</table>

levels of either troponin T (i.e., higher than 2 ng per mL [2 mcg per L]) at least eight hours from presentation or troponin I (i.e., higher than 1 ng per mL [1 mcg per L]) at least six hours from presentation support the diagnosis of MI or acute coronary syndrome and increase the likelihood of death or recurrent MI within 30 days. A normal level of troponin T or troponin I between six and 72 hours after the onset of chest pain is strong evidence against MI and acute coronary syndrome, particularly if the ECG is normal or near-normal.25,28 In one study29 of 773 patients who each presented to an emergency department with chest pain and had a normal ECG, researchers found that only 0.3 percent of those with a normal troponin I at six hours and 1.1 percent of those with a normal troponin T at six hours experienced acute MI or death in the 30 days following presentation. Thus, individuals with chest pain who have a history that indicates low risk of cardiovascular disease, a normal or near-normal ECG, and normal troponin levels can safely be evaluated as outpatients.

Patients at low risk usually do not need further testing unless there are other risk factors in their family or medical history that markedly increase their likelihood of CAD. Patients at intermediate risk for CAD who can exercise and have no left bundle branch block, preexcitation, or significant resting ST depression on their ECG can be evaluated with an exercise stress ECG. Patients with baseline ECG abnormalities should have perfusion imaging performed along with a stress ECG, and patients who cannot exercise may be evaluated with a pharmacologic stress or vasodilator test (e.g., dobutamine [Dobutrex], adenosine [Adenocard]). Patients at high risk for CAD generally should proceed directly to angiography, which allows definitive assessment of coronary artery anatomy for patients in whom other testing is nondiagnostic and for patients who could benefit from revascularization.30

For patients undergoing stress ECG testing, the Duke treadmill score (Table 731) provides helpful prognostic information. Among 1,466 patients with a normal resting ECG, and 939 patients with ST-T abnormalities on a resting ECG, low-, intermediate-, and high-risk Duke treadmill scores accurately predicted seven-year survival rates for all-cause mortality.31

<table>
<thead>
<tr>
<th>TABLE 6</th>
<th>Accuracy of Chest Pain Diagnosis Using Diagnostic and Prognostic Tests</th>
</tr>
</thead>
<tbody>
<tr>
<td>Diagnosis* (overall outpatient probability)</td>
<td>Clinical finding</td>
</tr>
<tr>
<td></td>
<td></td>
</tr>
<tr>
<td>Myocardial infarction (2%)4</td>
<td>New Q wave7</td>
</tr>
<tr>
<td></td>
<td>New conduction defect7</td>
</tr>
<tr>
<td></td>
<td>Any ST segment elevation7</td>
</tr>
<tr>
<td></td>
<td>Any Q wave7</td>
</tr>
<tr>
<td></td>
<td>Any conduction defect7</td>
</tr>
<tr>
<td></td>
<td>New T-wave inversion7</td>
</tr>
<tr>
<td></td>
<td>Troponin T &gt;2 ng per mL (2 mcg per L) at least eight hours from presentation25</td>
</tr>
<tr>
<td></td>
<td>Troponin I &gt;1 ng per mL (1 mcg per L) at least six hours from presentation25</td>
</tr>
<tr>
<td>Heart failure (2%)12</td>
<td>Abnormal electrocardiogram26</td>
</tr>
<tr>
<td></td>
<td>Abnormal BNP level (cutoff 80 pg per mL [1 ng per L])12</td>
</tr>
</tbody>
</table>

LR+ = positive likelihood ratio; LR- = negative likelihood ratio; BNP = brain natriuretic peptide.

*—Diagnoses are listed in order of clinical importance.

Information from references 4, 7, 12, 25, and 26.
PULMONARY EMBOLISM

D-dimer testing has become an important part of the evaluation for PE and deep venous thrombosis (DVT). Quantitative enzyme-linked immunosorbent antibody assay (ELISA) D-dimer assays are more sensitive and have been more thoroughly tested in clinical settings than whole-blood agglutination assays.32 A low clinical suspicion for PE (e.g., Wells score less than 2) plus a normal quantitative ELISA D-dimer assay safely rules out PE, with a negative predictive value greater than 99.5 percent.20,32,33 If further testing is needed, helical computed tomography (CT), combined with clinical suspicion and other testing such as lower extremity venous ultrasound, can be used to rule in or rule out PE.33,34 A number of different sequential testing protocols have been proposed, all of which involve the same basic elements: (1) for patients with low clinical suspicion and a normal D-dimer, no further evaluation or treatment is needed unless symptoms change or progress; (2) for patients with low clinical suspicion and an abnormal D-dimer, or moderate to high clinical suspicion, helical CT and lower extremity venous ultrasound examination should be ordered; (3) for patients with moderate or high clinical suspicion and an abnormal CT scan or venous ultrasound result, treatment should be given for PE or DVT regardless of D-dimer; and (4) for patients with an abnormal D-dimer plus a normal CT scan and a normal venous ultrasound result, serial ultrasound should be considered if clinical suspicion is low to moderate, and pulmonary angiography should be considered if clinical suspicion is high.33,35 Patients in whom PE initially is ruled out by such an approach and who do not receive treatment have a less than 1 percent risk for PE occurring over the subsequent three months.33 An encounter form that takes this approach appears in the February 1, 2004, issue of American Family Physician and can be accessed online at http://www.aafp.org/afp/20040201/pocform.html.36

PNEUMONIA AND HEART FAILURE

Chest radiograph generally is considered the reference standard for patients suspected of having pneumonia, and it is the standard against which clinical evaluations for pneumonia are compared.10 An abnormal ECG and cardiomegaly on chest radiograph increase the likelihood of heart failure among patients with chest pain,26 and brain natriuretic peptide (also known as B-type natriuretic peptide) level has been found to be reliable for detecting heart failure in patients presenting with acute dyspnea. Brain natriuretic peptide level is particularly helpful for ruling in heart failure if it is more than 500 pg per mL (500 ng per L), and for ruling out heart failure if it is less than 100 pg per mL (100 ng per L).14,37

CHEST WALL PAIN

Chest wall pain usually can be diagnosed by history and physical examination if other etiologies have been excluded. Measurement of the sedimentation rate generally is not helpful in making the diagnosis13; in unusual situations, radiography may be helpful.38

Recommended Diagnostic Strategy

An algorithm illustrating the discussed diagnostic strategy is provided in Figure 1.4,5,7-12,14-17,20-22,25,26,28,29,32-35 When a patient presents with new chest pain, a typical or an atypical
Outpatient Diagnosis of Chest Pain

Figure 1. Algorithm for the outpatient diagnosis of causes of chest pain. (ECG = electrocardiography; CT = computed tomography.)

Information from references 4, 5, 7 through 12, 14 through 17, 20 through 22, 25, 26, 28, 29, and 32 through 35.

NOTE: This algorithm combines and simplifies diagnostic recommendations from multiple sources to provide an overview, and does not represent a validated decision rule.

Patient presents with chest pain.

Does the patient have a typical or atypical anginal pattern, pain radiation or diaphoresis, or cardiac risk factors?

Is there clinical suspicion for pulmonary embolism?

Does the patient have fever, egophony, or dullness to percussion?

Perform chest radiography to evaluate for pneumonia.

Has the patient had spontaneous fright, anxiety, palpitations, dyspnea, or faintness in the past six months?

Consider panic disorder.

Is the pain reproducible by palpation?

Consider chest wall pain.

Consider heart failure or gastrointestinal pathology.

Calculate Wells score (see Table 4)

Wells score less than 2

Wells score 2 or greater

Measure D-dimer.

No further testing

Venous ultrasound result positive

Treat for deep venous thrombosis.

CT scan positive

Treat for pulmonary embolism.

Venous ultrasound result normal

No

Perform serial ultrasound.

Wells score 2 to 6

Perform pulmonary angiography.

Wells score greater than 6

Perform CT and venous ultrasound.

CT scan and venous ultrasound result normal (and the D-dimer abnormal)

Yes

MEASURE TROPNIM LEVELS SIX TO 72 HOURS AFTER THE ONSET OF CHEST PAIN.

Are troponin levels normal?

Evaluate as an inpatient.

No

Consider outpatient evaluation.

For patients with low cardiac risk, perform stress ECG.

For patients with moderate risk or abnormal ECG, perform stress test with perfusion scan.

For patients with high risk, perform angiography.

No

Yes

Is the ECG normal or near normal?

Measure troponin levels.

Is the ECG normal or near normal?

Yes

No

Is the ECG normal or near normal?
anginal pattern, pain radiation or diaphoresis, cardiac risk factors, or ischemic ECG changes, serial measurement of troponins should be considered to determine whether hospitalization or outpatient evaluation with stress testing is warranted. If the probability of PE is low, based on the Wells score, a negative D-dimer result eliminates the need for further testing; an abnormal D-dimer or moderate to high probability of PE should prompt helical CT and venous ultrasound examination to guide further management. Fever, egophony, or dullness to percussion should prompt evaluation for pneumonia with chest radiograph. If life-threatening causes of chest pain are ruled out, then a history of spontaneous anxiety, palpitations, faintness, or dyspnea suggests panic disorder. A history of exertional dyspnea and a displaced apical impulse should prompt investigation for heart failure. Gastrointestinal symptoms should prompt further evaluation.

Data Sources: The PubMed database was searched using the following terms: chest pain, angina, acute myocardial infarction, coronary artery disease, heart failure, pulmonary embolism, chest wall pain, bronchitis, pneumonia, and peptic ulcer disease. Titles were reviewed to identify literature relevant to the outpatient diagnosis of chest pain. Additional searches were performed using the following databases: InfoPOEMs (http://www.infopoems.com), Agency for Healthcare Research and Quality (http://www.ahrq.gov), Cochrane Collaboration (http://www.cochrane.org), Database of Abstracts of Reviews of Effects (http://www.york.ac.uk/inst/crd/darehp.htm), and Institute for Clinical Systems Improvement (http://www.ici.org).

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REFERENCES