Acute Stroke Diagnosis

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Stroke can be categorized as ischemic stroke, intracerebral hemorrhage, and subarachnoid hemorrhage. Awakening with or experiencing the abrupt onset of focal neurologic deficits is the hallmark of ischemic stroke diagnosis. The most common presenting symptoms for ischemic stroke are difficulty with speech and weakness on one half of the body. Many stroke mimics exist; two of the most common are a postictal seizure and hypoglycemia. Taking a detailed history and performing ancillary testing will usually exclude stroke mimics. Neuroimaging is required to differentiate ischemic stroke from intracerebral hemorrhage, as well as to diagnose entities other than stroke. The choice of neuroimaging depends on its availability, eligibility for acute stroke interventions, and the presence of patient contraindications. Subarachnoid hemorrhage presents most commonly with severe headache and may require analysis of cerebrospinal fluid when neuroimaging is not definitive. Public education of common presenting stroke symptoms is needed for patients to activate emergency medical services as soon as possible after the onset of stroke. (*Am Fam Physician.* 2009;80(1):33-40. Copyright © 2009 American Academy of Family Physicians.)

ILLUSTRATION BY ENID HATTON

The symptoms of stroke can sometimes be misleading and misinterpreted by physicians and patients. Family physicians are on the front line in their communities to recognize and manage acute cerebrovascular diseases. Accurate and prompt evaluation of cerebrovascular disease will increase eligibility of patients to receive acute therapy for stroke.

Classifying Stroke

Stroke can be subclassified by pathologic process and the vascular distribution affected. Defining the overall pathologic process is critical for decisions regarding thrombolysis, inpatient therapy, and prognosis. In the United States, 87 percent of all strokes are ischemic secondary to large-artery atherosclerosis, cardioembolism, small-vessel occlusion, and other or undetermined causes.[1,2] The remaining 13 percent of strokes are hemorrhagic in intracerebral or subarachnoid locations.[1,2] A common means of subclassifying ischemic stroke is by vascular distribution. Clinical determination of the affected vascular territory may aid rational evaluation and individualization of therapy.[3] However, this type of subclassification has only fair to good interobserver agreement among stroke experts.[4,5] *Table 1* lists stroke subtypes by vascular distribution.[6] The cause of subarachnoid hemorrhage is attributed to an aneurysm in approximately 85 percent of cases,[7] with rarer causes accounting for the rest.

Clinical Diagnosis

HISTORY AND PHYSICAL EXAMINATION

History and physical examination remain the pillars of diagnosing stroke. The most common historical feature of an ischemic stroke is its acute onset; the most common physical findings of ischemic stroke are focal weakness and speech disturbance.[8] The most common and reliable symptoms and signs of ischemic stroke are listed in *Table 2*.[4,8,9] Primary care physicians practicing in an emergency setting had a 92 percent sensitivity for diagnosis of stroke and transient ischemic attack (TIA) in a community-based study of

Table 1. Oxfordshire Ischemic Stroke Subtypes and Clinical Features

Stroke subtype	Clinical features
Total anterior circulation infarct (TACI)	Combination of new higher cerebral dysfunction (e.g., dysphasia, dyscalculia, visual-spatial disorder); homonymous visual field defect and ipsilateral motor and/or sensory defect involving two areas of the face, arm, or leg
Lacunar infarct (LACI)	Pure motor or pure sensory symptoms, sensorimotor stroke, or ataxic hemiparesis; face-arm and arm-leg syndromes included
Partial anterior circulation infarct (PACI)	Patients with only two of three TACI components, with higher cerebral dysfunction alone, or with a motor/sensory deficit more restricted than those classified as LACI (e.g., confined to one limb or to face and hand, but not the whole arm)
Posterior circulation infarct (POCI)	Any one of the following: ipsilateral cranial nerve palsy with contralateral motor and/or sensory deficit; bilateral motor and/or sensory deficit; disorder of conjugate gaze; cerebellar dysfunction without ataxic hemiparesis; isolated homonymous visual field defect

Information from reference 6.

Table 2. Most Common Symptoms and Signs of Stroke and Their Reliability

Symptom or sign	Prevalence (%)[8]	Agreement among examiners (Kappa*)[4]
Symptoms		
Acute onset	96	Good (0.63)[4]
Subjective arm weakness†	63	Moderate (0.59)[4]
Subjective leg weakness†	54	Moderate (0.59)[4]
Self-reported speech disturbance	53	Good (0.64)[4]
Subjective facial weakness	23	—
Arm paresthesia‡	20	Good (0.62)[4]
Leg paresthesia‡	17	Good (0.62)[4]
Headache	14	Good (0.65)[4]
Nonorthostatic dizziness	13	—
Signs		
Arm paresis	69	Moderate to excellent (0.42 to 1.00)[4,9]
Leg paresis	61	Fair to excellent (0.40 to 0.84)[4,9]
Dysphasia or dysarthria	57	Moderate to excellent (0.54 to 0.84)[4,9] Fair to excellent (0.29 to 1.00)[4,9]
Hemiparetic or ataxic gait	53	Excellent (0.91)[9]
Facial paresis	45	Poor to excellent (0.13 to 1.00)[4,9]
Eye movement abnormality	27	Fair to excellent (0.33 to 1.00)[9]
Visual field defect	24	Poor to excellent (0.16 to 0.81)[4,9]

*—Kappa statistic: 0 to 0.20 = poor agreement; 0.21 to 0.40 = fair agreement; 0.41 to 0.60 = moderate agreement; 0.61 to 0.80 = good agreement; 0.81 to 1.00 = excellent agreement.
†—Noted as "loss of power."[4]
‡—Noted as "loss of sensation."[4]
Information from references 4, 8, and 9.

diagnostic accuracy.[10] The overall accuracy of a physician's diagnosis of stroke is moderate to good, with lower reliability in less experienced or less confident examiners.[4]

Physicians need to quickly assess persons with suspected acute ischemic stroke because acute therapies for stroke have a narrower time window of effectiveness than therapies for myocardial infarction. The National Institutes of Health Stroke Scale (NIHSS[11,12]; available at http://www.ninds.nih.gov/doctors/NIH_Stroke_Scale.pdf) was designed to be completed in five to eight minutes.

The exact time of symptom onset is critical for determining eligibility for thrombolysis. However, a community-based study found that examiners agreed to the minute less than 50 percent of the time,[4] suggesting the need to corroborate time of symptom onset with a witness or known event.

Reliably distinguishing between intracerebral hemorrhage and ischemic stroke can only be done through neuroimaging. Both entities are characterized by acute onset of focal symptoms. Persons with intracerebral hemorrhage may have gradual worsening of symptoms after the abrupt onset, reflecting an increasing size of the hematoma. Persons with hemorrhage also may have a decreased level of consciousness.

Subarachnoid hemorrhage presents differently from intracerebral hemorrhage and ischemic stroke. The most common

Table 3. Accuracy of Selected Stroke Screening Tools

Name	Components	Sensitivity (% [95% CI])	Specificity (% [95% CI])	LR (95% CI)
Cincinnati Prehospital Stroke Scale[15]	Facial paralysis; arm drift; abnormal speech	≥ 1 item: 85 (80 to 90)[8]	≥ 1 item: 79 (73 to 85)[8]	0 items: 0.39 (0.25 to 0.61)[9] ≥ 1 item: 5.5 (3.3 to 9.1)[9]
Face, Arm, Speech Test[16]	In patients with Glasgow Coma Scale > 6 and presence of at least one of the following: facial paralysis; arm weakness; speech impairment	82 (76 to 88)[8]	83 (77 to 89)[8]	—
Los Angeles Prehospital Stroke Screen	Presence of all six items is positive for stroke: age > 45 years; no seizure history; symptoms present < 24 hours; ambulatory at baseline; serum glucose > 60 mg per dL (3.35 mmol per L) and < 400 mg per dL (22.20 mmol per L); unilateral deficit of one of three items (facial paresis, arm drift, weak handgrip)	91 (76 to 98)[17]	97 (93 to 99)[17]	LR+ = 31 (13 to 75)[17] LR− = 0.09 (0.03 to 0.27)[17]
Melbourne Ambulance Stroke Screen[18]	Presence of all six items is positive for stroke: age > 45 years; no seizure history; symptoms present < 24 hours; ambulatory at baseline; serum glucose > 60 and < 400 mg per dL; presence of ≥ one of four items (facial droop, arm drift, weak handgrip, speech impairment)	90 (81 to 96)	74 (53 to 88)	LR+ = 3.49 (1.83 to 6.63) LR− = 0.13 (0.06 to 0.27)
Recognition of Stroke in the Emergency Room scale[8]	A score of 1 point or higher is positive for stroke*: History of syncope or loss of consciousness (−1 pt) History of seizure activity (−1 pt) New acute onset of: Asymmetric facial weakness (+1 pt) Asymmetric arm weakness (+1 pt) Asymmetric leg weakness (+1 pt) Speech disturbance (+1 pt) Visual field defect (+1 pt)	93 (89 to 97)	83 (77 to 89)	LR+ = 5.49 (3.11 to 9.68) LR− = 0.083 (0.04 to 0.17)
von Arbin[19]	All three positive for stroke: acute onset of focal neurologic deficit; onset < seven days prior; no recent head trauma	86 (81 to 91)	99 (98.5 to 99.4)	LR+ = 94 (59 to 152) LR− = 0.14 (0.095 to 0.20)

CI = confidence interval; LR = likelihood ratio; LR− = negative likelihood ratio; LR+ = positive likelihood ratio.

*—Assessment follows blood glucose check and treatment if < 63 mg per dL (3.50 mmol per L).

Information from references 8, 9, and 15 through 19.

symptom described by the patient is the "worst headache of my life." Symptoms may also include vomiting, seizures, meningismus, and a decreased level of consciousness.[13] Persons with subarachnoid hemorrhage may not exhibit focal signs because the bleeding occurs outside the brain, except when an aneurysm bleeds into a focal location, such as a posterior communication artery aneurysm compressing the third cranial nerve.

VALIDATED DECISION SUPPORT TOOLS

The NIHSS is one of the most common classifications of early stroke severity[12]; it provides a structured neurologic examination that has diagnostic[11,14] and prognostic value.[11] Current guidelines recommend the use of the NIHSS,[11] but no trial data exist to show its use improves outcomes. In general, combinations of signs and symptoms are more useful than single findings. *Table 3* describes operating characteristics for several validated stroke diagnostic tools.[8,9,15-19] Common signs across stroke diagnostic tools include acute onset of unilateral weakness or numbness and speech disturbance.[8,15-19] One of the validated instruments, the Recognition of Stroke in the Emergency Room (ROSIER) scale, adds a visual field defect on examination.[8] Most of these tools were designed for prehospital care, but emergency department physicians using the ROSIER scale correctly classified 90 percent of all patients in a community-based validation study of consecutive patients seen in the United Kingdom.[8] Physicians using ROSIER missed patients with ischemic posterior or lacunar lesions, which emphasizes the need

Table 4. Stroke Mimics and Distinguishing Features

Condition	Distinguishing features
Seizure	History of loss of consciousness, seizure activity, or postictal state[14,20]
Systemic infection	Chest most common source[14]; acute illness exacerbating an old deficit[21]
Syncope/presyncope or hypotension	Hypotension unusual in acute stroke; prevalence of blood pressure < 120/80 mm Hg at initial stroke presentation = 7.1 percent[22]; symptoms may be transient or respond to hydration
Toxic-metabolic disturbances	Hypoglycemia most common[14,21]
Tumor	Mass noted on neuroimaging
Acute confusional state	May be related to alcohol intoxication, medication adverse effect, or other encephalopathy
Vertigo or dizziness	Imbalance, but not vertigo, increases the likelihood of stroke[23]; prevalence of stroke or transient ischemic attack in adults older than 44 years with isolated dizziness symptoms in emergency setting = 0.7 percent[23]
Migraine	History of similar events, preceding aura and headache[11]
Functional or medically unexplained symptoms	Reported incidence is 0.2 to 4.3 percent of patients admitted for stroke[8,24]; only history of headache or pre- or post-presentation functional syndrome was associated with unexplained symptoms in a United Kingdom case-control study; dysarthria, vertigo, and/or ataxia were less likely to be unexplained; motor, sensory, and visual field symptoms, stroke risk factors, and history of pre- or post-presentation depression or anxiety equally likely in patients with stroke and unexplained symptoms[24]
Dementia	Presence of known cognitive impairment was one of two factors that independently predicted a stroke mimic in an Australian prospective study of patients admitted with suspected stroke[14]

NOTE: *Conditions in approximate order of likelihood as a stroke mimic.*

Information from references 8, 11, 14, and 20 through 24.

for an examination more thorough than a scale alone. No head-to-head trials have been performed to demonstrate improved patient outcomes using a validated stroke scale versus global clinical impression. In practice, because most of these tools demonstrate good clinical accuracy, a physician should become familiar with one of them to help confirm their overall clinical impression of stroke.

STROKE MIMICS AND DIFFERENTIAL DIAGNOSIS

Physicians need to consider a broad differential diagnosis when evaluating a patient presenting with a suspected stroke *(Table 4)*.[8,11,14,20-24] The two most common stroke mimics are hypoglycemia and seizure.[8,14,20,21]

One potential area of confusion is among patients presenting with a symptom of dizziness. In a population-based study of adults older than 44 years presenting to the emergency department or directly admitted to the hospital with a principal symptom of dizziness, only 0.7 percent of patients with isolated dizziness symptoms had an ultimate diagnosis of stroke or TIA.[23] Vertigo from a central cause such as stroke is normally associated with nystagmus or other cerebellar signs.

The rates of overdiagnosis of stroke in studies of consecutive patients vary from 19 to 31 percent.[14,20,21] Known history of cognitive impairment,[14] non-neurologic abnormal physical findings,[14] and decreased level of consciousness[20] are independent predictors of a stroke mimic in patients with suspected stroke. Patient factors such as confusion, aphasia, and presentation more than 48 hours after the event also make diagnostic information less reliable.[14]

Duration of symptoms distinguishes stroke from TIA, which has been traditionally defined as a focal ischemic neurologic event resolving within 24 hours. Subsequent observations have shown that a majority of TIAs resolve within one hour.[25] The National Institute of Neurological Disorders and Stroke trial found that patients treated with placebo who did not have resolution in one hour or improvement in three hours had only a 2 percent chance of resolving in 24 hours.[26]

Diagnostic Tests and Imaging

Figure 1 presents an algorithm for the diagnosis of acute stroke.[2,4,11] *Table 5* lists initial diagnostic studies recommended by current guidelines for patients with suspected stroke.[11] These studies help exclude stroke mimics, uncover critical comorbidities (e.g., myocardial ischemia), and establish the safety of thrombolytic therapy.

The primary purpose of neuroimaging in a patient with suspected ischemic stroke is to rule out the presence of other types of central nervous system lesions and to distinguish between ischemic and hemorrhagic stroke. *Figure 2* shows examples of intracerebral and subarachnoid hemorrhages on computed tomography (CT) scans. CT scans are considered sufficiently sensitive for detecting mass lesions, such as a brain mass or abscess, as well as detecting acute hemorrhage. However, CT scans may not be sensitive enough to detect an ischemic stroke, especially if it is small, acute, or in the posterior fossa (i.e., brainstem and cerebellum areas).[27] The purpose of a CT scan is to rule out certain stroke mimics and detect hemorrhage, not necessarily to rule in the diagnosis of

ischemic stroke. In other words, a normal CT scan does not rule out the diagnosis of ischemic stroke.

Multimodal magnetic resonance imaging (MRI) sequences, particularly diffusion-weighted imaging, have better resolution than CT; therefore, they have a greater sensitivity for detecting acute ischemic stroke and show ischemic lesions in about one half of all cases of TIA.[28] Recent studies also indicate that MRI sequences (particularly gradient-recalled echo and diffusion-weighted imaging sequences) are as sensitive as CT scans for detecting intracerebral hemorrhagic stroke.[29,30] *Figure 3* shows the head CT and diffusion-weighted MRI images of a patient with a prior stroke and a new acute stroke. *Figure 4* depicts the time course of resolution of ischemic changes on diffusion-weighted MRI.[28]

Although MRI scans have better resolution than CT scans, MRI scanners are less available and more expensive than CT scanners. Also, MRI sca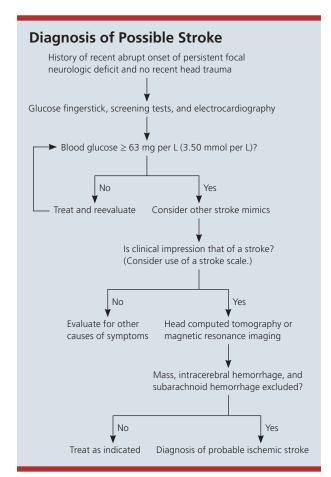ns cannot be performed on persons with certain types of implanted devices (e.g., pacemakers) or in persons with claustrophobia. If a patient is within the time window of acute stroke intervention, guidelines recommend that an MRI scan can be ordered if it can be obtained as quickly as a CT scan; if not, then CT is the recommended test because acute stroke treatments should not wait for detailed imaging when the history and physical are consistent for acute stroke.[11] *Online Table A* compares CT and MRI in the setting of acute stroke.[11,26] Guidelines recommend that whichever imaging modality is performed, it should be interpreted by a physician with expertise in reading brain imaging studies.[11]

Diagnosis of Possible Stroke

History of recent abrupt onset of persistent focal neurologic deficit and no recent head trauma

↓

Glucose fingerstick, screening tests, and electrocardiography

↓

Blood glucose ≥ 63 mg per L (3.50 mmol per L)?

- No → Treat and reevaluate
- Yes → Consider other stroke mimics

↓

Is clinical impression that of a stroke? (Consider use of a stroke scale.)

- No → Evaluate for other causes of symptoms
- Yes → Head computed tomography or magnetic resonance imaging

↓

Mass, intracerebral hemorrhage, and subarachnoid hemorrhage excluded?

- No → Treat as indicated
- Yes → Diagnosis of probable ischemic stroke

Figure 1. Algorithm for the diagnosis of acute stroke.

Information from references 2, 4, and 11.

Table 5. Immediate Diagnostic Studies: Evaluation of Suspected Acute Ischemic Stroke

All patients

Noncontrast brain CT or brain MRI

Blood glucose

Serum electrolytes and renal function tests

Electrocardiography

Markers of cardiac ischemia

Complete blood count, including platelet count*

Prothrombin time/International Normalized Ratio*

Activated partial thromboplastin time*

Oxygen saturation

Selected patients

Hepatic function tests

Toxicology screen

Blood alcohol level

Pregnancy test

Arterial blood gas (if hypoxemia suspected)

Chest radiography (if lung disease suspected)

Lumbar puncture (if subarachnoid hemorrhage suspected and head CT negative for blood)

Electroencephalography (if seizure suspected)

CT = computed tomography; MRI = magnetic resonance imaging.

—Although it is desirable to know the results of these tests before giving recombinant tissue plasminogen activator, thrombolytic therapy should not be delayed while awaiting the results unless there is clinical suspicion of a bleeding abnormality or thrombocytopenia, the patient has received heparin or warfarin (Coumadin), or the use of anticoagulants is not known.

Adapted with permission from Adams HP Jr, del Zoppo G, Alberts MJ, et al. Guidelines for the early management of adults with ischemic stroke: a guideline from the American Heart Association/American Stroke Association Stroke Council, Clinical Cardiology Council, Cardiovascular Radiology and Intervention Council, and the Atherosclerotic Peripheral Vascular Disease and Quality of Care Outcomes in Research Interdisciplinary Working Groups: the American Academy of Neurology affirms the value of this guideline as an educational tool for neurologists [published corrections appear in Stroke. *2007;38(6):e38, and* Stroke. *2007;38(9):e96].* Stroke. *2007;38(5):1665. http://lww.com.*
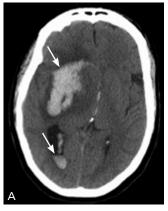
Figure 2. Head computed tomography (CT) scans showing *(A)* intracerebral hemorrhages *(arrows)* and *(B)* subarachnoid hemorrhages *(arrows)*. Note that acute hemorrhage appears hyperdense *(white)* on a CT scan.
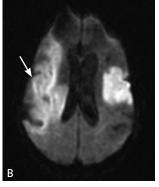

Figure 3. *(A)* Noncontrast computed tomography (CT) scan showing two hypodense regions indicating old infarctions in the distribution of the left-middle cerebral *(long arrow)* and posterior cerebral arteries *(short arrow)*. *(B)* Diffusion-weighted magnetic resonance imaging scan obtained shortly after the CT reveals a new extensive infarction *(arrow)* in the right-middle cerebral artery distribution not evident on the CT.

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Unlike ischemic stroke and intracerebral hemorrhage, diagnosing subarachnoid hemorrhage requires a different diagnostic algorithm. The frequency of misdiagnosis for subarachnoid hemorrhage can be as high as 50 percent on initial presentation.[13] Although MRI can detect subarachnoid hemorrhage, CT is still considered the imaging test of choice for persons suspected to have subarachnoid hemorrhage.[31] CT scans have a 95 to 100 percent sensitivity of detecting subarachnoid blood in the first 12 hours; however, unlike ischemic stroke, sensitivity greatly decreases over time as the subarachnoid blood is cleared.[13] The sensitivity of subarachnoid hemorrhage detection by CT drops to about 50 percent after one week, and is not detectable by CT after a period of about two to three weeks.[13,31]

Persons with suspected subarachnoid hemorrhage and a normal CT scan should undergo a lumbar puncture to detect bilirubin. Red blood cells can be found in a subarachnoid hemorrhage and a traumatic tap. Distinguishing between these two entities requires recognition that only within the human body do red blood cells break down into bilirubin. Red blood cells in cerebrospinal fluid collected from a traumatic tap will break down into oxyhemoglobin, but not into bilirubin. Because the breakdown of red blood cells can take up to 12 hours, guidelines recommend that the lumbar puncture should wait until 12 hours after the initial onset of symptoms.[32] Bilirubin will turn fluid yellow (xanthochromia), but visual inspection alone is not considered sufficiently reliable.[32] Therefore, all specimens should undergo spectrophotometry analysis to detect bilirubin, which can be detected as long as two weeks after the initial onset of symptoms. If subarachnoid hemorrhage is detected,
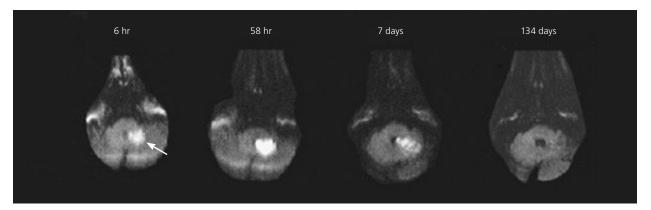

Figure 4. Time course of diffusion-weighted imaging abnormalities. Diffusion-weighted imaging can detect ischemic stroke *(arrow)* in the very early period *(six-hour image)*. Lesions typically reach maximum intensity three to five days after stroke onset *(as shown in the 58-hour image)*, then typically fade over one to four weeks *(abnormality still visible in seven-day image, but has disappeared in the 134-day image)*.

Adapted with permission from Schaefer PW, Grant PE, Gonzalez RG. Diffusion-weighted MR imaging of the brain. Radiology. *2000;217(2):336.*

SORT: KEY RECOMMENDATIONS FOR PRACTICE

Clinical recommendation	Evidence rating	References
Patients with an abrupt onset of a focal persistent neurologic deficit should be evaluated for stroke.	C	8, 9, 17, 19
Stroke mimics should be excluded by history and diagnostic testing.	C	8, 11, 14, 20, 21, 23
Diagnostic tools, such as the Recognition of Stroke in the Emergency Room scale, can aid in stroke diagnosis.	C	8, 9
All patients with stroke should have urgent neuroimaging with computed tomography or magnetic resonance imaging.	C	11
Patients and family members should be educated about stroke symptoms and the need for urgent evaluation.	C	11

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, go to http://www.aafp.org/afpsort.xml.

the patient should immediately undergo angiography (CT angiography, MRI angiography, or catheter angiography) to look for an aneurysm.

Teaching Patients to Recognize Stroke Symptoms

Guidelines recommend that persons having an acute stroke activate the emergency medical system by calling 9-1-1.[11] However, patients often do not activate the emergency medical system, or do not activate it immediately. Such behavior accounts for up to two thirds of the delay to hospital admission.[33] Therefore, patients commonly present outside the time window for thrombolytic therapy. Patient and family sense of urgency for stroke symptoms is associated with greater use of emergency medical systems,[34] which results in shorter times to evaluation and admission.[35,36]

Numerous surveys have shown that there is considerable room for improvement in knowledge of stroke in the general population.[37-39] When persons are asked to answer "yes" or "no" as to whether a described symptom can be a sign of a stroke, they are correct about 60 to 80 percent of the time.[37] However, when persons are asked open-ended questions to name stroke warning signs, most cannot name more than one warning sign.[37]

To improve public awareness of stroke warning signs, numerous organizations have embarked on public education campaigns.[40-41] Family physicians are well placed to emphasize these messages in their practices.

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