# What's the best diagnostic evaluation of night sweats?

#### **Evidence-based answer**

There is no single best evidence-based approach to the diagnostic evaluation of night sweats, given the limited number of studies on the subject. A detailed history, however, does appear to be the most important initial diagnostic tool (strength of recommendation [SOR]: **C**, based on usual practice and clinical opinion).

No clinical trials have directly studied symptomatic relief of night sweats alone.

Among menopausal women with hot flashes associated with night sweats, oral hormone therapy is highly effective in reducing their frequency (SOR: **A**, based on a Cochrane review with a clear recommendation). Antireflux therapy may also be effective (SOR: **B**, based on a cohort study). Therapy aimed at decreasing perspiration has also been suggested (SOR: **C**, based on clinical opinion.)

#### Cindy W. Su, MD and Sean Gaskie, MD

Sutter Santa Rosa Family Medicine Residency Program, Santa Rose, Calif

Kristin Hitchcock, MSI Department of Preventive Medicine, Northwestern University, Chicago, III

#### **Clinical commentary**

## Night sweats are an increasingly common complaint

Complaints of night sweats among my menopausal patients have become very common with the declining use of hormone replacement therapy. Both women and their bed partners are affected, and sleep deprivation is a significant side effect, so the problem must be taken seriously.

Though venlafaxine can cause night sweats, it is also a reasonable treatment

strategy for menopause-related night sweats. Gabapentin may hold promise for hormonal symptoms if reflux is not the issue. Other sinister causes of night sweats are uncommon, but are always in the back of my mind when the issue is raised, so the history and review of systems help focus the work-up. The pretest probability of unusual diagnoses guides specific laboratory testing.

Lisa Johnson, MD Providence Health Care Systems, University of Washington, Seattle

#### Evidence summary

Night sweats are a common complaint in the ambulatory primary care setting: Of 2267 patients in 1 cross-sectional study, 41% reported night sweats, defined as "sweating at night even when it isn't excessively hot in your bedroom" within the previous month.¹ Because the peak prevalence in both men and women occurred in the group ages 41 to 55 years, there was concern

that menopausal hot flashes were a confounding factor, at least for women. In a subsequent study of 795 patients older than 64 years, 10% still reported being bothered by night sweats.<sup>2</sup>

## The more common causes are not widely studied

Few studies look at the causes of night sweats. Although they have been associated with tuberculosis, lymphoma, and

#### FAST TRACK

If history
and exam
are unrevealing,
a trial of antireflux
medication
is recommended

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Medications that may cause sweating or flushing	
ANTIDEPRESSANTS	HYPOGLYCEMIC AGENTS
Bupropion (Wellbutrin) SSRIs Tricyclic antidepressants Venlafaxine (Effexor)	Insulin Sulfonylureas SYMPATHOMIMETIC AGENTS  Beta-agonists Phenylephrine (Sudafed)
ANTIMIGRAINE DRUGS	
Naratriptan (Amerge)	OTHER AGENTS
Rizatriptan (Maxalt) Sumatriptan (Imitrex) Zolmitriptan (Zomig)	Alcohol Beta-blockers
ANTIPYRETICS	Bromocriptine (Parodel) Calcium channel blockers Clozapine (Clozaril) Cyclosporine Hydralazine (Hydra-Zide) Niacin Nitroglycerin Omeprazole (Prilosec) Opioids Sildenafil (Viagra) Tamoxifen (Nolvadex) Theophylline Tramadol (Ultram, Ultracet)
Acetaminophen Aspirin Nonsteroidal anti-inflammatory drugs (NSAIDs) CHOLINERGIC AGONISTS	
Bethanechol (Urecholine) Pilocarpine	
GNRH AGONISTS	
Gonadorelin Goserelin (Zoladex) Histrelin (Vantas) Leuprolide (Lupron) Nafarelin (Synarel)	
Source: UpToDate.4	

#### FAST TRACK

For menopausal women, oral hormone therapy effectively reduces night sweats

HIV infection, these are not common causes of night sweats in outpatient care.

In the only study that specifically addressed the causes of night sweats in an ambulatory population, Reynolds<sup>3</sup> interviewed 200 consecutive patients, 70% from a primary care practice and 30% from a gastroenterology practice. Of the 81 patients who reported having an episode of night sweats at least once a week, esophageal reflux and menopause were the most frequent causes.

Several authors agree that certain medications are frequently associated with night sweats, although the exact incidence is unknown due to a lack of published epidemiologic data.<sup>4-6</sup> Anti-depressants and antipyretics are among the more commonly cited offenders (TABLE 1).<sup>4</sup>

## Finding the right diagnosis requires thorough history & exam

With such a long differential diagnosis (TABLE 2),<sup>4-6</sup> night sweats should initially be evaluated with a thorough history and physical examination (according to a consensus opinion of various authors). If these don't elicit possible causes, the appropriate next step in the work-up can vary. Some authors recommend multiple laboratory and imaging studies, while others advise against any routine tests. None of these approaches is evidence-based.

One reasonable algorithm recommends an initial work-up including a complete blood count, thyroid-stimulating hormone (TSH) and erythrocyte sedimentation rate (ESR) level, a purified protein derivative (PPD) and HIV test, and a chest x-ray.<sup>5</sup> If the results are un-

revealing, a trial of antireflux medication is recommended. If the patient does not improve, consider a diary of nocturnal temperatures to help discern the presence or absence of febrile pulses and further evaluate for suspected endocarditis or lymphoma.

#### **Evidence** is scant for symptom relief

Very few clinical trials have directly studied symptomatic relief of night sweats. A large Cochrane meta-analysis found that oral hormone therapy—estrogens alone or estrogens with progesterone-reduced the frequency of night sweats associated with hot flashes among menopausal women by 75% when compared with placebo alone.<sup>7</sup> Neither primrose oil nor foot reflexology proved effective.8

A cohort study found that 80% of the patients with frequent night sweats responded to antireflux therapy.<sup>3</sup> One author suggests using therapies aimed at relieving hyperhydrosis.<sup>6</sup> These include local treatment with aluminum chloride hexahydrate (Drysol), antiperspirants, scopolamine, or phenoxybenzamine hydrochloride (Dibenzyline).

#### **Recommendations from others**

A thorough literature search through Cochrane Database Systematic Reviews, AHRO, National Guideline Clearinghouse, and Medline did not yield any guidelines or consensus statements from other organizations or specialty groups on the evaluation or treatment of night sweats. ■

#### References

1. Mold JW, Mathew MK, Belgore S, Dehaven M. Prevalence of night sweats in primary care patients: an OKPRN and TAFP-Net collaborative study. J Fam Pract 2002; 51:452-456.

### TABLE 2

#### Differential diagnosis for night sweats

Pinoronala diagnosis for highe officato		
ENDOCRINE	NEUROLOGIC DISORDERS	
Carcinoid syndrome Diabetes insipidus Hyperthyroidism	Autonomic dysreflexia Autonomic neuropathy Stroke	
Hypoglycemia Pheochromocytoma	SUBSTANCE WITHDRAWAL	
Post-orchiectomy	Alcohol Cocaine Opioids	
INFECTIONS		
Coccidioidomycosis Endocarditis Histoplasmosis Human immunodeficiency virus Infectious mononucleosis Lung abscess Mycobacterium avium complex Osteomyelitis Tuberculosis	MISCELLANEOUS	
	Chronic fatigue syndrome Gastroesophageal reflux disease Menopause Obstructive sleep disorder Panic disorder Pregnancy Prinzmetal's angina Takayasu's arteritis Temporal arteritis	
MALIGNANCY		
Leukemia Lymphoma Prostate cancer Renal cell carcinoma Other neoplasms		

Source: UpToDate; Viera et al, Am Fam Physician 2003; Chambliss, Arch Fam Med 1999.

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- 3. Reynolds WA. Are night sweats a sign of esophageal reflux? J Clin Gastroenenterol 1989; 11:590-
- 4. Smetana GW. Approach to the patient with night sweats. UpToDate [database online]. Updated October 3, 2006. Available at: www.uptodate.com.
- 5. Viera AJ, Bond MM, Yates SW. Diagnosing night sweats. Am Fam Physician 2003; 67:1019-1024.
- 6. Chambliss ML. What is the appropriate diagnostic approach for patients who complain of night sweats? Arch Fam Med 1999; 8:168-169.
- 7. MacLennan AH, Broadbent JL, Lester S, Moore V. Oral oestrogen and combined oestrogen/progestogen therapy versus placebo for hot flushes. Cochrane Datab Syst Rev 2004; CD002978.
- 8. Williamson J, White A, Hart A, Ernst E. Randomised controlled trial of reflexology for menopausal symptoms. BJOG 2002; 109:1050-1055.

#### **FAST** TRACK

Local treatmentssuch as antiperspirants and scopolaminehave limited evidence

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