

## Prescribing opioids

(MARCH 2016)

**TO THE EDITOR:** As a primary care physician who has practiced for 31 years, I have a few concerns about the article “Prescribing opioids in primary care: Safely starting, monitoring, and stopping” by Drs. Daniel Tobin, Rebecca Andrews, and William Becker in your March 2016 issue.

Prescribing narcotics and other controlled medications has become a hot topic with legal implications. Many physicians have lost their license, and some have lost their freedom as well.

The article notes that primary care physicians provide most of the care for chronic pain but implies that we don’t know what we are doing. Although we might not have completed a residency in the management of chronic pain, we were required to attend a graduate medical school and to complete residency programs. We are also required to read the medical literature and keep up with our CME requirements. We too strive to keep up with and practice the latest cutting-edge medicine. And we have experience. After seeing thousands of patients and writing millions of prescriptions, I think I do know something about whether or not a medicine is safe, effective, and cost-effective. I have encountered quite a few patients with pain, and not one of them has overdosed by taking the medications as prescribed.

Most of what is being said and published about pain medication focuses on the epidemic of young people who are getting pain medications (prescription narcotics) and street narcotics (heroin) illegally. Directly, it has nothing to do with conscientious physicians prescribing narcotics for pain, but indirectly, it has to do with all adults in society. We are allowing our young people to be destroyed by drugs and by a lack of proper discipline in our homes that spills over into society. Where are our children getting these drugs, and who is bringing them into our neighborhoods? The practice of medicine is not the primary problem. I am for scientific, conscientious regulations

concerning controlled substances. Medicine needs to inform all doctors about the changing laws surrounding the prescribing of controlled substances.

Cigarettes and alcohol are sold on every corner, and places selling marijuana are popping up everywhere. The former two drugs are harming and killing more young people than all the illegal drugs combined, and marijuana is fast approaching number three. I feel we need to stop picking on the medical profession and stop trying to blame it for all the woes that our young people are encountering every day. The reality is that legal and illegal drugs are not going to go away. We need to love our children more and better prepare them while in our homes to deal with the evils in our society.

**WILLIE J. PETTIWAY, MD**  
Lighthouse Medical Center  
Detroit, MI

doi:10.3949/ccjm.83c.08001

**IN REPLY:** We thank Dr. Pettiway for his remarks. The intent of our article was to identify common challenges when prescribing opioids for chronic pain and offer tips to the provider struggling with how to do so safely. We hope these comments will offer additional clarity.

First, as general internists who are essentially “self-trained” in the management of chronic pain, we fully acknowledge the importance of practical experience in learning how to prescribe opioids safely and effectively. Dr. Pettiway is correct that a dedicated physician who keeps up with the medical literature, attends relevant continuing medical education courses, and strives to provide deliberate, rational, and evidence-based care to his or her patients can do so effectively. However, the medical literature suggests that medical school training in the management of chronic pain is sparse; one review found that in 2011 only 5 out of 133 US medical schools required coursework on pain management, and only 13 offered it as an elective.<sup>1</sup> Many primary care providers do feel unprepared to handle this challenge.

Additionally, Dr. Pettway raises a good question about where misused prescription opioids originate and whether prescribers are responsible. The data show that the majority of misused prescription opioids are obtained from a family member or friend and not directly from a physician.<sup>2,3</sup> However, this supply does generally originate from a prescription. Providers need to educate their patients about the risk for diversion, the need to keep pills safely hidden and locked away, and the importance of safely discarding unused supplies. Responsible prescribers need to anticipate these concerns and educate patients about them.

In summary, we firmly believe that primary care providers are capable of safe, effective, and responsible opioid prescribing and hope that our paper provides additional guidance on how to do so.

DANIEL G. TOBIN, MD, FACP  
Assistant Professor, Yale University  
School of Medicine

REBECCA ANDREWS, MD, FACP  
Associate Professor, University of Connecticut

WILLIAM C. BECKER, MD  
Assistant Professor, VA Connecticut  
Healthcare System, Yale University  
School of Medicine

#### REFERENCES

1. Roehr B. US needs new strategy to help 116 million patients in chronic pain. *BMJ* 2011; 343:d4206.
2. Becker WC, Tobin DG, Fiellin DA. Nonmedical use of opioid analgesics obtained directly from physicians: prevalence and correlates. *Arch Intern Med* 2011; 171:1034–1036.
3. Substance Abuse and Mental Health Services Administration. Results from the 2013 National Survey on Drug Use and Health: summary of national findings. HHS Publication No. (SMA) 14-4863. Rockville, MD: Substance Abuse and Mental Health Services Administration; 2014. [www.samhsa.gov/data/sites/default/files/NSDUHresultsPDFHTML2013/Web/NSDUHresults2013.htm](http://www.samhsa.gov/data/sites/default/files/NSDUHresultsPDFHTML2013/Web/NSDUHresults2013.htm). Accessed June 29, 2016.

doi:10.3949/ccjm.83c.08002

## Measles: More than the rash

(MAY 2016)

**TO THE EDITOR:** I read the excellent overview article on measles by Drs. Kumar and Sabela.<sup>1</sup> However, there are additional important clinical points regarding measles diagnosis that deserve further comment. Prior to definitive diagnosis, measles is a clinical diagnosis. Properly, much attention is given to the rash, but there are important clinical clues besides the rash that are helpful diagnostically.

Some clinical findings are more characteristic of a disease than others, eg, eye findings in measles, and clinicians should specifically look for them. Other findings not characteristic but consistent with the diagnosis are less helpful, eg, measles with diarrhea.<sup>2</sup> Measles is a systemic infection with several extradermatologic manifestations. Characteristically, measles involves the respiratory tract, manifested by runny nose, dry cough, or shortness of breath, ie, measles pneumonia.<sup>2</sup>

Gastrointestinal involvement may be manifested as nausea, vomiting, diarrhea, or

abdominal pain. Abdominal pain, when located in the right lower quadrant, may mimic acute appendicitis, ie, pseudoappendicitis. In patients undergoing appendectomy, pathologically the appendix is normal but contains multinucleated giant cells (Warthin-Finkeldey cells). Measles pseudoappendicitis can be perplexing because it presents before the rash. Even without appendiceal involvement (pseudoappendicitis), Warthin-Finkeldey cells are also present in the nasal mucosa. If nasal swab cytology shows Warthin-Finkeldey cells, an early diagnosis of measles may be made days before IgM measles antibodies are reported.<sup>3</sup>

Other often-overlooked important clues are eye findings, eg, watery eyes, lower lid edema, conjunctival suffusion, and keratitis. No other disease in the differential diagnosis of measles presents with watery eyes with bilateral lower lid edema and conjunctival suffusion. Headache, mental confusion or oral ulcers may suggest alternate diagnoses. Typically, nonspecific laboratory abnormalities include leukopenia and thrombocytopenia, and importantly, the erythrocyte sedimentation rate is not elevated (Table 1).<sup>4,5</sup>

TABLE 1

### Nonrash clinical diagnostic findings in adults with measles

#### Symptoms

Head, eyes, ears, nose, and throat  
Headache  
Runny nose  
Sore throat

#### Systemic

Dry cough  
Abdominal pain (pseudoappendicitis if in the right lower quadrant<sup>a</sup>)

#### Signs

##### Eyes

Conjunctival suffusion  
Watery eyes  
Lower lid swelling  
Keratitis

##### Throat

Koplik spots  
Dark red buccal mucosa  
Oral ulcers

#### Laboratory tests

Complete blood cell count  
Leukopenia  
Relative lymphopenia  
Thrombocytopenia  
Unelevated erythrocyte sedimentation rate

#### Other

Elevated lactate dehydrogenase  
Elevated ferritin  
Elevated aminotransferases  
Warthin-Finkeldey cells on nasal swab cytology (early only)

<sup>a</sup>If appendectomy is performed, cytology shows Warthin-Finkeldey multinucleated giant cells.

Based on information in reference 5.

In measles, much is made of Koplik spots, which are found early on dark red buccal mucosa (vs Fordyce spots).<sup>1</sup> However, if these are missed or not present, clinicians can use other characteristic findings to make a presumptive diagnosis of measles.

BURKE A. CUNHA, MD  
State University of New York  
School of Medicine  
Stony Brook, New York

#### REFERENCES

1. Kumar D, Sabella C. Measles: back again. *Cleve Clin J Med* 2016; 83:340–344.
2. Leibovici L, Sharir T, Kalter-Leibovici O, Alpert G, Epstein LM. An outbreak of measles among young adults. Clinical and laboratory features in 461 patients. *J Adolesc Health Care* 1988; 9:203–207.
3. Cunha BA, Hage JE, Turi GK. A 45-year-old male with rash, fever and diarrhea. *J Clin Microbiol* 2012; 50:1835,2184.
4. Cunha BA, Mickail N. Fever and rash in an adult traveler: the many masquerades of measles. *Travel Med Inf Dis* 2011; 9:255–257.
5. Cunha CB, Cunha BA. Rash and fever in the intensive care unit. In: Fink MP, Abraham E, Vincent JL, Kochanek P, editors. *Textbook of Critical Care Medicine* (7th Ed). Elsevier, Philadelphia, 2016:97–115.

doi:10.3949/ccjm.83c.08003

**IN REPLY:** We thank Dr. Cunha for his comments and appreciate the opportunity to emphasize important points that he highlights.

We agree that measles is a systemic illness with important extradermatologic manifestations that are critical to the diagnosis, and that the nondermatologic manifestations often precede the rash and serve to distinguish measles from other systemic illnesses. As discussed in our review, the respiratory prodrome of cough, coryza, and conjunctivitis is very distinctive and serves as an important clue to the diagnosis. Likewise, we acknowledge the importance of gastrointestinal findings in measles and note appendicitis as an important complication that is well described. Although Koplik spots are pathognomonic, we do stress that these often are not present at the time of presentation.

Finally, we agree that measles is a clinical diagnosis, and that the clinical manifestations beyond the dermatologic manifestations noted in our review and highlighted by Dr. Cunha are extremely helpful to the clinician in considering the diagnosis.

CAMILLE SABELLA, MD  
Cleveland Clinic  
Cleveland Clinic Lerner College of Medicine of Case Western Reserve University

DHEERAJ KUMAR, MD  
Cleveland Clinic

doi:10.3949/ccjm.83c.08004

## Acute liver failure

(JUNE 2016)

**TO THE EDITOR:** In the article by Drs. Singh et al in your June issue, I was surprised that the role of hepatitis delta wasn't mentioned as a potential cause of acute liver failure. My understanding is that this peculiar virus can only infect those with hepatitis B surface antigenemia, but when it does, it results in far more serious liver injury, including acute liver failure in some.

HOWARD HOMLER MD, FACP  
Carmichael, CA

doi:10.3949/ccjm.83c.08005

**IN REPLY:** We thank Dr. Homler for bringing hepatitis D as a potential cause of acute liver failure to our attention.

Hepatitis D virus, first described in the 1970s, requires the hepatitis B surface antigen (HBsAg) capsid to enter the hepatocyte and, thus, can only cause liver injury when the patient is also infected simultaneously with hepatitis B virus.<sup>1</sup> Hepatitis D virus can cause either coinfection (presence of immunoglobulin M anti-HB core antibody with or without HBsAg) or superinfection (presence of HBsAg without immunoglobulin M anti-HB core antibody) with hepatitis B virus. In India, coinfection has been reported to be the cause of acute liver failure in about 4% of all

patients, and superinfection in 4.5%.<sup>2</sup>

While simultaneous treatment for hepatitis D and B viruses with pegylated interferon and any of the agents used for treatment of hepatitis B has been successful in treating chronic hepatitis, it has not been proven useful in patients with acute liver failure, and liver transplant remains the only treatment option.<sup>3</sup>

TAVANKIT SINGH, MD  
Cleveland Clinic

NANCY GUPTA, MD  
Valhalla, NY

NAIM ALKHOURI, MD  
Cleveland Clinic

WILLIAM D. CAREY MD  
Cleveland Clinic

IBRAHIM HANOUNEH, MD  
Minneapolis, MN

### ■ REFERENCES

1. **Rizzetto M.** The adventure of delta. *Liver Int* 2016; 36(suppl 1):135–140.
2. **Irshad M, Acharya SK.** Hepatitis D virus (HDV) infection in severe forms of liver diseases in North India. *Eur J Gastroenterol Hepatol* 1996; 8:995–998.
3. **Noureddin M, Gish R.** Hepatitis delta: epidemiology, diagnosis and management 36 years after discovery. *Curr Gastroenterol Rep* 2014; 16:365.

doi:10.3949/ccjm.83c.08006