

Five Things Physicians and Patients Should Question

1

Do not routinely send urine for *Chlamydia trachomatis* and *Neisseria gonorrhoeae* (CT/NG) testing from females if vaginal swab collection is possible.

Nucleic acid amplification testing (NAAT) for CT/NG is standard of care for testing adults and has largely replaced culture. Vaginal swabs are the preferred single specimen for screening and diagnosis of CT/NG by NAAT providing 5% to >10% increased sensitivity compared to urine for females; testing multiple specimens (vaginal, endocervical, and urine) can further increase sensitivity. When vaginal collections are not possible due to the setting, test or collection device shortages, or very strong patient preference, *first-void* urine can be considered as a non-invasive alternative.

2

Do not perform heterophile antibody (monospot) testing to diagnose acute EBV infection in children less than 5 years of age.

Approximately 40% of children under 5 years do not develop heterophile antibodies following primary EBV infection. If the heterophile is the only test ordered, the diagnosis may be missed. The U.S. Centers for Disease Control and Prevention has advised against heterophile testing in this age group due to lack of specificity and potential for false negative results. Testing in this age group should be a panel of EBV-specific serologic antibody immunoassays for viral capsid antigen (VCA) IgM and IgG and Epstein-Barr nuclear antigen (EBNA).

3

Do not test for influenza unless the patient is symptomatic and the result will influence clinical management and decision making.

The United States Centers for Disease Control and Prevention (CDC) National Center for Immunization and Respiratory Diseases (NCIRD) recommends when influenza is circulating in the community that hospitalized patients with influenza indications undergo influenza testing, and the Infectious Disease Society of America (IDSA) recommends rapid influenza molecular assays or reverse transcriptase polymerase chain reaction [RT-PCR] testing for greatest diagnostic certainty. The NCIRD indications include typical influenza symptoms, atypical presentations, complications, and admission status.

Three scenarios to consider:

1. Perform molecular testing if a patient has signs and symptoms suggestive of influenza, including an atypical clinical presentation, or findings suggestive of complications associated with influenza and is being admitted to the hospital.
2. Perform molecular testing if a patient has signs and symptoms suggestive of influenza, and is not being admitted to the hospital, but the result will influence clinical management.
3. If a patient is not symptomatic, then influenza testing is probably not indicated.

4

Do not prescribe immune suppressive agents for suspected autoimmune hepatitis (AIH) without first excluding hepatotropic virus infections (e.g., viral hepatitis A, B, and C). Viral hepatitis may mimic AIH, both serologically and histologically, features that may resolve with direct-acting antiviral (DAA) treatment.

Viral hepatitis, including that caused by hepatitis A, B, and C, can lead to development of autoantibodies (e.g., antinuclear, anti-smooth muscle actin (SMA)/F-actin, liver–kidney microsomal type 1 (LKM-1), soluble liver antigen (SLA), and immunoglobulin G (IgG)) in approximately 50% of cases. The autoantibody profile in patients with chronic HCV may cause clinical suspicion of concurrent AIH and may prompt unnecessary liver biopsies and/or immune suppressive treatments. HAV, HBV, and HCV, may have similar histologic and clinical features including frequent plasma cells, elevated transaminases and may lead to cirrhosis. However, treatment is markedly different. Treatment of HCV is with interferon (IFN)-free or DAA therapy, achieving sustained viral responses (SVR) in most cases and has been shown to eliminate both clinical and histologic features of AIH. Therefore, patients with chronic hepatitis C who have serum markers and/or histologic features of AIH should first be treated with DAA in most cases.

5

Don't order tissue transglutaminase IgG antibody or Deamidated Gliadin Peptide (DGP) antibodies (IgG or IgA) in the initial screening for Celiac Disease.

Tissue transglutaminase IgA antibody (anti-tTG IgA) is the recommended first-line screening test for celiac disease because it provides the best diagnostic sensitivity and specificity. Serum IgA should also be included to detect IgA deficiency. Tissue transglutaminase IgG antibody (anti-tTG IgG) or deamidated gliadin peptide antibodies (IgG or IgA) could be appropriate as reflex tests in specific situations based on initial findings although they have less specificity than the tissue transglutaminase IgA antibody. In particular, Deamidated Gliadin Peptides result in a higher false positive rate that can lead to further unnecessary testing and/or endoscopy.

How This List Was Created (1–2)

The American Society for Microbiology's (ASM) list was developed under the leadership of the ASM's Clinical and Public Health Microbiology Committee. The subject matter experts who identified the list and formulated the recommendations are laboratory directors at academic, commercial and public health laboratories and test utilization experts across the fields of microbiology and laboratory medicine. They worked together to identify a list of diagnostic and management decisions that have resulted in misuse of laboratory studies and resources.

In this submission, two statements were written to address the most common clinical microbiology laboratory test misconceptions. They consist of diagnostic tests or treatments that are commonly ordered, expensive and have no evidence to illustrate its value and in some cases, may be potentially harmful to the patient. The recommendations, if instituted, would result in higher quality care, lower costs, and more effective use of our laboratory resources and personnel. The experts involved in the new 2022 recommendations are James Dunn, Laura Filkins, Omai Garner, Elizabeth Palavecino and Preeti Pancholi.

How This List Was Created (3–4)

These ASCLS recommendations were developed under the leadership of ASCLS's Choosing Wisely Committee and the ASCLS Board of Directors. The Committee examined numerous options based on evidence available. Subject matter experts from the ASCLS Scientific Assemblies reviewed, edited, and recommended approval of these recommendations, which were subsequently reviewed and approved by the ASCLS Board of Directors.

How This List Was Created (5)

The American Society for Clinical Pathology (ASCP) recommendation was developed under the leadership of the ASCP Effective Test Utilization Steering Committee. This committee is chaired by an ASCP Past President and is comprised of subject matter and test utilization experts across the fields of pathology and laboratory medicine. The committee considered a list of possible recommendations compiled as the result of a survey administered to Society members serving on ASCP's many commissions, committees and councils. In addition, an announcement was made to ASCP's Advisory Board seeking suggestions for possible recommendations to promote member involvement. The laboratory tests targeted in our recommendations were selected because they are tests that are performed frequently; there is evidence that the test either offers no benefit or is harmful; use of the test is costly and it does not provide higher quality care; and eliminating it or changing to another test is within the control of the clinician. Implementation of these recommendations will result in higher quality care, lower costs and a more effective use of our laboratory resources and personnel.

Sources

1

1. Papp JR, Schachter J, Gaydos CA, Van Der Pol B. 2014. Recommendations for the laboratory-based detection of *Chlamydia trachomatis* and *Neisseria gonorrhoeae* - 2014. MMWR Recomm Reports 63:1–19.
2. Shafer MA, Moncada J, Boyer CB, Betsinger K, Flinn SD, Schachter J. 2003. Comparing first-void urine specimens, self-collected vaginal swabs, and endocervical specimens to detect *Chlamydia trachomatis* and *Neisseria gonorrhoeae* by a nucleic acid amplification test. J Clin Microbiol 41:4395–4399.
3. Van Der Pol B, Taylor SN, Liesenfeld O, Williams JA, Hook EW. 2013. Vaginal swabs are the optimal specimen for detection of genital *Chlamydia trachomatis* or *Neisseria gonorrhoeae* using the cobas 4800 CT/NG test. Sex Transm Dis 40:247–250.

2

1. C.A. Horwitz, W. Henle, G. Henle, et al., Clinical and laboratory evaluation of infants and children with *Epstein-Barr virus-induced infectious mononucleosis*: report of 32 patients (aged 10–48 months), Blood 57 (1981) 933–938.
2. Centers for Disease Control and Prevention. Epstein-Barr virus and infectious mononucleosis. Centers for Disease Control and Prevention; 2014. [Accessed 08 Aug 2014 2014]. <http://www.cdc.gov/epstein-barr/laboratory-testing.html>.
3. American Academy of Pediatrics. Kimberlin DW, Barnett ED, Lynfield R, Sawyer MH, Epstein-Barr Virus Infections. Red Book: 2021 Report of the Committee on Infectious Diseases. 32nd ed. Itasca, IL: American Academy of Pediatrics; 2021:318–322.

3

1. Centers for Disease Control and Prevention. Guide for considering influenza testing when influenza viruses are circulating in the community. Website accessed December 2020. <https://www.cdc.gov/flu/professionals/diagnosis/consider-influenza-testing.htm>.
2. Centers for Disease Control and Prevention. Overview of Influenza Testing Methods. Website accessed December 2020. <https://www.cdc.gov/flu/professionals/diagnosis/overview-testing-methods.htm>.
3. Centers for Disease Control and Prevention. Algorithm to assist in the interpretation of influenza testing results and clinical decision-making during periods when influenza viruses are circulating in the community. Website accessed December 2020. <https://www.cdc.gov/flu/professionals/diagnosis/algorithm-results-circulating.htm>.
4. Merckx J, Wali R, Schiller I, Caya C, Gore GC, Chartrand C, Dendukuri N, Papenburg J. Diagnostic accuracy of novel and traditional rapid tests for influenza infection compared with reverse transcriptase polymerase chain reaction: a systematic review and meta-analysis. Ann Intern Med. 2017;167:394–409.

4

1. Simoes CC, Saldarriaga OA, Utay NS, Stueck AE, Merwat SK, Merwat SN, Schiano TD, Fiel MI, Stevenson HL. Direct-acting antiviral treatment of patients with hepatitis C resolves serologic and histopathologic features of autoimmune hepatitis. Hepatol Commun. 2019; 3:1113–23. doi: 10.1002/hep4.1388. PMID: 31388631; PMCID: PMC6671831.
2. Himoto T, Masaki T. Extrahepatic manifestations and autoantibodies in patients with hepatitis C virus infection. Clin Dev Immunol 2012; 2012:871401.
3. Cassani F, Cataleta M, Valentini P, Muratori P, Giostra F, Francesconi R, et al. Serum autoantibodies in chronic hepatitis C: comparison with autoimmune hepatitis and impact on the disease profile. Hepatology 1997; 26:561-566.
4. Pavic S, Simonovic J, Boricic I, Svrtlih N. Autoantibodies characteristic for autoimmune hepatitis found in chronic hepatitis C. [In Serbian] Srp Arh Celok Lek 2003; 131:437-442.
5. Clifford BD, Donahue D, Smith L, Cable E, Luttig B, Manns M, et al. High prevalence of serological markers of autoimmunity in patients with chronic hepatitis C. Hepatology 1995;21:613-619.
6. Sugiura A, Wada S, Mori H, Kimura T, Matsuda Y, Tanaka N, et al. Successful treatment for chronic hepatitis C-autoimmune hepatitis overlap syndrome due to daclatasvir and asunaprevir. Case Rep Gastroenterol 2017; 11:305-311.

1. Rubio-Tapia A, Hill ID, Kelly CP, Calderwood AH, Murray JA; American College of Gastroenterology. ACG clinical guidelines: diagnosis and management of celiac disease. *Am J Gastroenterol*. 2013 May;108(5):656-76; quiz 677. doi: 10.1038/ajg.2013.79. Epub 2013 Apr 23. PMID: 23609613; PMCID: PMC3706994.
2. Husby S, Koletzko S, Korponay-Szabó I, Kurppa K, Mearin ML, Ribes-Koninckx C, Shamir R, Troncone R, Castillejo G, Christensen R, Dolinsek J, Gillett P, Hróbjartsson A, Koltai T, Maki M, Nielsen SM, Popp A, Størdal K, Werkstetter K, Wessels M. European Society Paediatric Gastroenterology, Hepatology and Nutrition Guidelines for Diagnosing Coeliac Disease 2020. *J Pediatr Gastroenterol Nutr*. 2020 Jan;70(1):141-156. doi: 10.1097/MPG.0000000000002497. PMID: 31568151.
3. Husby S, Murray JA, Katzka DA. AGA Clinical Practice Update on Diagnosis and Monitoring of Celiac Disease-Changing Utility of Serology and Histologic Measures: Expert Review. *Gastroenterology*. 2019 Mar;156(4):885-889. doi: 10.1053/j.gastro.2018.12.010. Epub 2018 Dec 19. PMID: 30578783; PMCID: PMC6409202.

About the ABIM Foundation

The mission of the ABIM Foundation is to advance medical professionalism to improve the health care system. We achieve this by collaborating with physicians and physician leaders, medical trainees, health care delivery systems, payers, policymakers, consumer organizations and patients to foster a shared understanding of professionalism and how they can adopt the tenets of professionalism in practice.

To learn more about the ABIM Foundation, visit www.abimfoundation.org.



About the American Society for Microbiology

The American Society for Microbiology is the largest single life science society, composed of 30,000 scientists and health professionals. ASM's mission is to promote and advance the microbial sciences.

ASM advances the microbial sciences through conferences, publications, certifications and educational opportunities. It enhances laboratory capacity around the globe through training and resources. It provides a network for scientists in academia, industry and clinical settings. Additionally, ASM promotes a deeper understanding of the microbial sciences to diverse audiences.

To learn more about ASM, visit asm.org



About the American Society for Clinical Laboratory Science

The American Society for Clinical Laboratory Science (ASCLS) and its 9,000 clinical laboratory professional, student, and educator members in more than 50 state and regional constituent societies work to advance the expertise of clinical laboratory professionals who, as integral members of interprofessional healthcare teams, deliver quality, consumer-focused, outcomes-oriented clinical laboratory services through all phases of the testing process to prevent, diagnose, monitor and treat disease. The Society promotes high standards of practice by holding the profession accountable to a Code of Ethics, through dissemination of knowledge at educational programs and through publications; maintains a supportive community to advocate on behalf of current and future laboratory professionals; and provides laboratory professionals a voice to legislators and regulators through collective, grassroots efforts.

To learn more about ASCLS, visit ascls.org



About the American Society for Clinical Pathology

Founded in 1922 in Chicago, ASCP is the world's largest professional membership organization for pathologists and laboratory professionals. ASCP provides excellence in education, certification, and advocacy on behalf of patients, anatomic and clinical pathologists, and medical laboratory professionals.

To learn more about ASCP, visit www.ascp.org.

