Cervical Cancer Screening Visualization Technologies

Overview

This Coverage Policy addresses visualization technologies used for cervical cancer detection and identification.

Coverage Policy

The following visualization technologies are considered experimental, investigational or unproven for any indication including, but not limited to, cervical cancer screening because they are considered experimental, investigational or unproven:

- cervicography
- spectroscopy/optical detection systems
- speculoscopy

General Background
Several technologies have been proposed for the screening or identification of cervical cancer. They are proposed as an adjunct to or a replacement for the standard techniques of Papanicolaou (Pap) smear, human papillomavirus (HPV)-deoxyribonucleic acid (DNA) (HPV-DNA) testing, and colposcopy.

**Cervicography**
Cervicography is a visual screening method introduced in the 1970s that uses a specially designed 35-mm camera to take photographs of the cervix after the application of a 3–5% acetic acid wash. The film is then sent to a laboratory for processing and evaluation. The theory behind cervicography is that when an expert evaluates the cervical photographs there will be an improvement in identification of cervical lesions and improved ability to discriminate between high grade and more trivial lesions than the mid-level clinicians who perform direct visual inspection (Wright, et al., 2002).

**Spectroscopy**
A spectroscopy system may be referred to as an optical detection system. Spectroscopy emits light from a probe onto the cervix, allowing the examiner to objectively categorize tissues as either normal or diseased. Spectroscopy is based on the principle that epithelial tissues that are abnormal have different optical properties than normal tissues and that these optical differences can be used to determine whether a tissue is normal or abnormal. Devices that are currently under various stages of research and development for diagnostic purposes use various approaches, including: fluorescence spectroscopy, white light elastic backscatter spectroscopy, infrared spectroscopy, Raman spectroscopy, image analysis of visible images, or combinations of the different methods (Wright, et al., 2002).

One optical detection system is the LUMA™ Cervical Imaging System (MediSpectra, Inc., Lexington, MA). This device received premarket approval from the FDA in March 2006. The LUMA system uses three different optical measurements to document cervical abnormalities: native evoked fluorescence, diffuse reflectance backscatter, and video imaging. The FDA indicated use is as an adjunct to colposcopy for identification of high-grade disease (cervical intraepithelial neoplasia [CIN] 2, 3+) in women referred for colposcopy with a Pap test result of atypical squamous cells, low-grade squamous intraepithelial lesion, high-grade squamous intraepithelial lesion or cancer.

**Speculoscopy**
The speculoscopy exam includes the same principle that is used in a colposcopy of applying acetic acid to the cervix, but then utilizes magnified vision and a chemiluminescent light to detect abnormalities. The vaginal vault and cervix are mostly illuminated by light that reflects off of the surface of the cervix, thus producing a nonspecific glare and obscuring the clear-cut definition of any acetowhite lesions that may exist. Lesions that are below the surface of the epithelium may not be detected, thus leading to decreased efficacy of this exam process.

Speculite produces a low-energy, diffuse, blue-white light. Due to chemiluminescence, the light that is emitted is "cold light" and independent of temperature. This light is affixed to the top of the speculum, and through special spectral frequencies it is theorized that early dysplastic lesions that reside on or below the epithelial surface can be discovered. The dysplastic tissue will appear white and have sharp borders between the normal and abnormal epithelium that can be visualized.

In 2002, PapSure® was granted 401(k) approval from the FDA. PapSure combines the results of a traditional Pap smear and speculoscopy using Speculite, a disposable, chemiluminescent light for vaginal illumination. Both of these devices are manufactured by Watson Diagnostics, Corona, CA.

**Literature Review for Cervicography:** The early studies evaluating this technology are small and use heterogeneous populations and testing protocols. Based on results in large screening studies, cervicography does not appear to have an adequate sensitivity, even when the performance of the test is highly optimized to be used as a primary screening method for cervical cancer screening (Wright, et al., 2002).

Song et al. (2019) conducted a study that compared the screening capacities and cost-effectiveness of the human papillomavirus (HPV) test versus cervicography as an adjunctive test to Papanicolaou (Pap) cytology to detect high-grade cervical neoplasia in Korea, a country with a high prevalence of cervical cancer. The study included 33,531 women who underwent cervicography as a screening test for cervical cancer, with a
retrospective analysis of the records of 4117 women who simultaneously or subsequently underwent Pap cytology, an HPV test, cervicography, and colposcopically directed biopsy. At a threshold of cervical intraepithelial neoplasia grade 2 or worse (CIN2+), based on colposcopic biopsy, the diagnostic capacities and cost-effectiveness of these screening tools were compared. The CIN2+ prevalence was 10.8% (446 of 4117 women) and the positive rate of high-risk HPV was 61.0% (2511 of 4117 women). Cervicography as an adjunctive to Pap cytology was a more sensitive test (97.5% vs 93.7%) with a higher odds ratio (15.65 vs 5.86) than the HPV test for detection of CIN2+ ( P -value = 0.003). The cost of cervicography co-testing was 23% less than that of HPV co-testing. The authors concluded that cervicography and Pap co-testing had superior screening capacity and cost-effectiveness for detection of preinvasive cervical lesions than HPV and Pap co-testing and may be an effective and cost-saving screening strategy in clinical practice in country with a high prevalence of cervical cancer; however, further large, randomized controlled trials comparing the screening capacities of HPV test versus cervicography as an adjunctive test to Pap cytology to detect high-grade cervical neoplasia in general population are needed to obtain more conclusive data. The study was limited by the lack of randomization and retrospective nature of the study.

de Castro Hillmann et al. (2019) conducted a cross sectional study of 212 patients in a colposcopy referral center to evaluate the performance of cervical digital photography (CDP) as an alternative to colposcopy. CDP is also referred to as digital cervicography or digital camera assessment for the detection of cervical intraepithelial neoplasia. Colposcopy and CDP with the cervical digital photographs evaluated through the Internet by three colposcopy experts. The agreement between methods was calculated with kappa and percentages of agreement. Then the sensitivity, specificity, positive predictive value, negative predictive value, and diagnostic accuracy were calculated for colposcopy and cervical digital photography. Histology was used as the gold standard (Canadian Task Force Classification II-2). CDP and colposcopy were in agreement in 89.9% of the cases (κ = 0.588). CDP had higher sensitivity (52.5%) and positive predictive value (60%) than colposcopy (35% and 48.28%, respectively). There were no other significant differences between CDP and colposcopy: specificity was 91.86% and 91.28%, negative predictive value was 89.3% and 85.8%, and diagnostic accuracy was 84.4% and 80.7%, respectively. The authors concluded that CDP is a promising alternative method to colposcopy and additional research could better determine the overall utility of CDP in clinical practice.

Singhakum et al. (2018) conducted a non-inferiority study of new portable device as an alternative method for cervical cancer screening. The performance of device was tested on the assessment of cervical lesions using cervicograph score with the cervical cytology. The study included 325 non-pregnant women. The cervical and vaginal cells from the sample were collected for cytology, then all of them received the digital cervicography conducted with the new device and scored using cervicograph score. Small pieces of cervical tissues were also collected for histologic examination. Cytology results and cervicograph scores were grouped to two subgroups, two subgroups, ≤ ASC-US and ≥ LSIL, and 0-3 points and 4-6 points, respectively. The data was then correlated with histology results which sub-grouped to ≤ CIN 1 and ≥ CIN 2. The accuracy, sensitivity, specificity, and positive predictive value (PPV) of cervicograph scores, 4-6 points, to detect CIN 2+ were 92%, 72.41%, 97%, and 84%, respectively which were not inferior to Pap smear. The authors concluded that the digital cervicography device provides similar accuracy to Pap cytology screening and is suitable to use in the area that lacks cytoscreeners, however large scale use and generalization are required for this new device.

Kim et al. (2013) conducted a study of 261 patients that compared the sensitivities and false-positive rates of cervical cytology (Pap smear), human papilloma virus (HPV) DNA test, cervicography, first double-combined testing (cervical cytology and HPV DNA test), second double-combined testing (cervical cytology and cervicography) and triple-combined testing (cervical cytology, HPV DNA test and cervicography). All women simultaneously underwent cervical cytology, HPV DNA test and cervicography for uterine cervical cancer screening and colposcopy-directed biopsy for diagnostic evaluation. Twenty-eight cases classified as atypical squamous cells of undetermined significance (ASCUS) on cervical cytology were excluded from the statistical analysis due to the ambiguity in classification. The sensitivity of cervical cytology was 87.5%; specificity 93.5%; positive predictive value (PPV) 77.8%; and negative predictive value (NPV) 96.7%. The sensitivity of the HPV DNA test was 72.7%; specificity 91.7%; positive predictive value 70.2%, and negative predictive value 92.7%. The sensitivity of the cervicography was 94.3%; specificity 89.8%, PPV 71.4%; and NPV 98.3%. The sensitivity of this double-combined testing was 92.3%; specificity 86.6%; PPV 65.8%; and NPV 97.6%. Triple- combined testing the sensitivity was 100%; specificity 82.2%; PPV 62.8%; NPV 100%. The authors note that results of this study cannot be applied directly for uterine cervical cancer screening since it was conducted in patients showing
a high incidence—further group studies should be carried out using mass screening. In addition, further problems that remain to be resolved include regional biases, objectivity in reading, accuracy in diagnostic criteria, economic feasibility, excessive treatment due to high sensitivity, and the inconvenient nature of the tests themselves.

The results of a nested study conducted during a large multicenter, randomized, prospective analysis were reported on by Guido[a] et al. (2005). This nested study was designed to address the issue of the topographic distribution of lesions, particularly CIN 3 lesions. The researchers felt that using a population that was already enrolled in the prospective study provided a well-studied and documented source of cervical intraepithelial neoplasia (CIN) of different grades from four diverse clinical centers. During this study, all women were randomized to three treatment arms at the time of their enrollment: 1) immediate colposcopy; 2) HPV triage to colposcopy using Hybrid Capture 2; and 3) conservative management based on repeat cytology with colposcopic referral at an HSIL threshold. All participants had liquid-based cytology sampling for Pap and HPV typing, and cervigrams were taken at enrollment and follow-up visits. Those participants enrolled in the immediate colposcopy arm had both cytology and colposcopic exams conducted on the same day. All participants were followed every six months by cytology and underwent exit colposcopy at two years. Guido and colleagues wanted to study the possible relationship between the outcomes of cervical biopsies, the biopsy's cervical location based on an o'clock position, and the quality of the biopsy based on cervigram acetowhiteness. Acetowhite areas were more common on the anterior and posterior lips of the cervix; however, this variance did not correlate to an increase in the number of CIN or HPV positive cytology results. The presence of acetowhiteness may have indicated a resolving HPV infection, although acetowhiteness appeared when HPV results were negative as well. These findings raised concern that in the absence of disease, the anterior and posterior lips of the cervix still reacted to acetowhiteness, causing an increase in the numbers of biopsies taken, but the biopsies did confirm the presence of CIN. The researchers therefore concluded that the use of this technique requires additional research.

**Literature Review for Spectroscopy:**

Hermens et al. (2017) reported on the diagnostic value of alternative (digital) colposcopy techniques for detection of cervical intraepithelial neoplasia (CIN) 2 or worse in a colposcopy population. The inclusion criteria included: an alternative (digital) colposcopy technique was used in a colposcopy population; a histologic outcome was reported, classified as CIN, differentiating between mild dysplasia or less (CIN 1 or less), and moderate dysplasia or worse (CIN 2 or greater); the entire cervix was scanned at once or a per-woman analysis was performed; no other topical application than acetic acid and Lugol's solution was used; and at least three eligible studies had to be available within a single technique. Thirteen studies met the inclusion criteria. With six studies on fluorescence and reflectance spectroscopy (2,530 women) with a pooled sensitivity of 93% (95% confidence interval [CI] 89-95%) and specificity of 62% (95% CI 47-76%). Four studies on dynamic spectral imaging were found including 1,173 women with a pooled sensitivity of 69% (95% CI 48-85%) and specificity of 83% (95% CI 76-88%). Previously published conventional colposcopy results showed a sensitivity of 61% (95% CI 58-63%) and a specificity of 85% (95% CI 83-86%). The authors concluded that alternative digital colposcopy techniques may result in increased or similar sensitivity and specificity compared with conventional colposcopy; however, the techniques are still in development, randomized controlled trials comparing alternative techniques with conventional colposcopy are still lacking, and therefore no recommendation for introduction in clinical practice can yet be made.

El-Tawail et al (2008) reported on a comparative study between Pap smear cytology and Fourier transform infrared (FTIR) spectroscopy. Eight hundred cervical scrapings were taken by cytobrush and placed in ThinPrep medium. The samples were dried over infrared transparent matrix. Beams of infrared light were directed at the dried samples at frequency of 4000 to 400 cm-1. The absorption data were produced using a Spectrum BX II FTIR spectrometer. Data was then compared with the reference absorption data of known samples using FTIR spectroscopy software. FTIR spectroscopy was compared with cytology (gold standard). It was noted that FTIR spectroscopy could differentiate normal from abnormal cervical cells in the samples examined—the sensitivity was found to be 85%, specificity 91%, positive predictive value 19.5% and negative predictive value of 99.5%.

Alvarez et al. (2007a) conducted multicenter, two-arm, randomized trial to assess whether the use of an optical detection system as an adjunct to colposcopy increases the detection of biopsy confirmed CIN 2, 3. The trial compared colposcopy alone with colposcopy plus a pre-commercial optical detection system that utilized
Alvarez et al. (2007b) conducted a multicenter internally controlled trial to evaluate the impact of using an optical detection system as an adjunct to colposcopy. The trial was designed to evaluate the performance of a pre-commercial optical detection system (LUMA) used as an adjunct to colposcopy among women referred for the evaluation of an abnormal cervical cytology that were randomized with stratification by cytology. The main study outcomes were differences in TP rates (CIN 2, 3 and cancer identified) and FP rates between the study arms. The TP rates were 14.4% versus 11.4% (p=0.035, one-sided) for the combined colposcopy and optical detection system arm compared to the colposcopy-only arm, respectively, in women with either an atypical squamous cell (ASC) or low-grade squamous intraepithelial lesion (LSIL) cytology result. The TP rates were similar between the two arms among women referred for the evaluation of high-grade squamous intraepithelial lesion (HSIL) in the combined colposcopy and ODS arm, among women with ASC or LSIL, the PPV of biopsies indicated by optical detection system was 15.0% and the PPV of biopsies indicated by colposcopy was 15.2%.

DeSantis et al. (2006) conducted a prospective multicenter study to evaluate the potential safety and effectiveness of tissue spectroscopy for the diagnosis of cervical cancer. The study involved 572 women who were scheduled to undergo colposcopy on the basis of an abnormal Pap test or other risk factor. The spectroscopy measurements were taken over a scan period of four minutes and 30 seconds. The measurements were integrated by a cross-validated pattern recognition model and then compared with biopsy results to yield sensitivity and specificity of cervical spectroscopy. The sensitivity of cervical spectroscopy was 95.1% with a corresponding 55.2% specificity for benign lesions. There were several potential confounding factors (e.g., mucus, blood, patient motion, ambient light) examined to determine their potential impact on the accuracy of the test. Ambien light appeared to have the greatest effect, but no single factor contributed significantly to the results.

**Literature Review for Speculoscopy:** To evaluate the efficacy of Pap smear, speculoscopy, and the combination of Pap smear and speculoscopy (PapSure examination) as screening tests in pre- and postmenopausal women, Twu and colleagues conducted a multicenter study in 2006. Based on records within a nationwide government database of Pap smear registration, 1813 women were assessed for possible inclusion in this study and of these, 1701 were eligible (873 premenopausal and 828 postmenopausal). The patients underwent successive Pap smears, speculoscopy, and the first 40 patients each day received simultaneous colposcopic examinations. The remaining patients were referred for colposcopy if their Pap smear or speculoscopy revealed abnormal results. A positive Pap smear was defined as ASUS/AGUS or worse. Positive speculoscopy was defined as a marked acetowhite lesion with sharp margins. Abnormal colposcopic findings were defined as acetowhite lesions with sharp margins, irregular surface, or atypical vessel patterns (coarse punctuations, mosaic, etc.). Punch biopsies and endocervical curettage (ECC) was performed on all patients with unsatisfactory examinations. For premenopausal women, speculoscopy and PapSure had significantly higher sensitivity (p<0.005) and lower specificity (p<0.001) than did the Pap smear. The PapSure examination showed a higher sensitivity than the Pap smear (85.7% versus 57.1%), but the results were not statistically significant. Speculoscopy and PapSure had significantly lower specificity than did Pap smear (96.8%, 96.6% and 99.6%, respectively, [p<0.001]). The authors concluded that based on their data, combining Pap smear with speculoscopy improved sensitivity with minimal reduction in specificity within premenopausal women; however, in postmenopausal women this lower specificity could lead to unnecessary colposcopic examinations or possible conizations. It is unclear if the cytologist was blinded to the findings of other test outcomes as the first 40
individuals were referred on a daily basis for colposcopy examination. Patients were referred for cervical biopsy based on the presence or absence of acetowhite lesions; this referral for additional testing may have led to an increase in false-positive readings that were observed in the premenopausal group versus the post-menopausal women. Although the researchers set the minimum inclusion age at 30, participants younger than this were allowed to be a part of the study, which may have also led to an increase in false-positive outcomes.

Parham [a] et al. (2000) reviewed the outcomes of using immediate (i.e., within four weeks) colposcopy on women who had a positive Pap smear versus delaying colposcopy for six months if the speculoscopy exam alone was positive. Of the 800 women Parham screened, 124 had negative Pap smears but positive speculoscopy. Of the 124 women, 57 were offered immediate colposcopy and 67 were offered colposcopy in six months. More than 80% of the women in the immediate group had positive colposcopy results, with 64.9% histologically-proven neoplasms. Thirteen (29%) lesions in the deferred group showed speculoscopy-negative results on repeat testing. Of the lesions that remained positive at the six-month mark, 90.6% were confirmed neoplasms on biopsy; this provided a sensitivity yield of 65–90%. The researchers concluded that this sensitivity yield was due to the combination of the Pap smear, colposcopy and additional biopsy of tissues. The population set that was studied by delayed colposcopy was small in size (14 of 67 lost to follow-up). Individuals were subjected to additional diagnostic tests due to false-positive speculoscopy readings. Individuals with positive Pap smear results that were read as low-grade SIL or ASCUS were not detected as having cervical neoplasia by speculoscopy alone, but required additional biopsies to confirm the presence of neoplasia. After six months, 29% (13 of 45) positive speculoscopy readings converted from positive to negative; these individuals would not have required a colposcopy.

Cervical Cancer Screening

Several professional organizations have published guidelines for cervical cancer screening, including American Cancer Society (ACS), American College of Obstetricians and Gynecologists (ACOG), National Comprehensive Cancer Network® (NCCN®) and the U.S. Preventive Services Task Force (USPSTF). These screening guidelines include criteria for tests that should be used and frequency according to age. The following standard techniques are included in the guidelines:

- **Pap smear**: The Pap smear or Pap test is the gold standard of cervical cancer screening tests. While the clinical utility of this test has never been examined in a randomized controlled trial, there is a large body of consistent observational data that supports its effectiveness in reducing mortality from cervical cancer (NCI-a, 2019). The Pap smear remains the standard of screening for cervical cancer, being recognized by numerous guidelines from professional organizations and medical/specialty associations and included in medical textbooks.

- **Liquid-Based Cytology**: In an attempt to improve the Pap smear’s sensitivity and specificity, researchers have developed, studied and implemented the use of liquid-based cytology collection systems and automated screening computerized programs. The use of liquid-based cytology has become a standard of care in screening protocols for cervical cancer, according to statements from professional and specialty medical organizations, published literature and textbooks (Eifel, et al., 2011; Niederhuber, et al., 2013; Smith, et al., 2008).

- **Human Papillomavirus Deoxyribonucleic Acid (HPV-DNA) Testing**: The major risk factor for cervical cancer is infection by the human papilloma virus (HPV). HPV is a group of more than 100 related viruses. There are certain types of HPV that are considered high-risk or carcinogenic since they often lead to cancer of the cervix. These types include HPV 16, HPV 18, HPV 31, HPV 33, HPV 45—approximately 70% of all cervical cancers are caused by HPV 16 and 18 (American Cancer Society [ACS], 2016). HPV-DNA testing allows for the detection of primary etiologic infectious carcinogenic carriers. The HPV-DNA test, like the Pap test, is performed by collecting cells from the cervix and then sending them to a laboratory for analysis. The test detects high-risk types of HPV in cell DNA even before there are any conclusive visible changes to the cervical cells.

- **Colposcopy**: colposcopy along with colposcopically directed biopsies is the primary method for evaluating women with abnormal cervical cytologies. During a colposcopy, the cervix is visualized and excess mucus is gently removed with a dry cotton ball, the cervix is treated with 3% to 5% acetic acid. Flat condylomata or dysplastic areas turn white or develop a vascular pattern with a mosaic appearance or punctuation. The squamocolumnar junction and transformation zone are then inspected thoroughly, and biopsy of suspicious areas is performed (Damewood, et al., 2008).
**Professional Societies/Organizations**
The use of cervicography, speculoscopy or spectroscopy/optical detection systems as a primary screening technique of the cervix for the early detection of cervical cancer, as an alternative or adjunct to pap smear or colposcopy is not included in professional society/organization guidelines for cervical cancer screening.

**Use Outside of the US**
**NHS Cervical Screening Programme (United Kingdom):** the NHS includes the following recommendations (NHS, 2015; 2019):
- Women age less than 24: not invited to screen
- Women age 25-49: every three years with cytology
- Women age 50–64 years: every 5 years with cytology
- Women aged ≥ 65 years: to undergo screening only if they have not had screening since age 50, or they have had recent abnormal test results
- HPV testing: additional (triage) HPV testing is recommended for women ≥ 25 years with abnormal test results in some circumstances

**Singapore Ministry of Health:** The Singapore Ministry of Health published guidelines for cancer screening. Regarding cervical cancer screening, the guidelines include (2010):
- All women who have ever had sexual intercourse should undergo screening for cervical cancer from the age of 25 years. Grade C, Level 2*
- Pap smear screening should be performed at least once every 3 years. Grade B, Level 2**
- Screening should be performed using the Papanicolaou (Pap) smear. Grade B, Level 2**

1**: High quality meta-analyses, systematic reviews of randomized controlled trials (RCTs), or RCTs with a very low risk of bias
1*: Well conducted meta-analyses, systematic reviews of RCTs, or RCTs with a low risk of bias.
2**: High quality systematic reviews of case control or cohort studies. High quality case control or cohort studies with a very low risk of confounding or bias and a high probability that the relationship is causal
2*: Well conducted case control or cohort studies with a low risk of confounding or bias and a moderate probability that the relationship is causal

Grade B: A body of evidence including studies rated as 2**, directly applicable to the target population, and demonstrating overall consistency of results; or extrapolated evidence from studies rated as 1** or 1*
Grade C: A body of evidence including studies rated as 2*, directly applicable to the target population and demonstrating overall consistency of results; or extrapolated evidence from studies rated as 2**

**Medicare Coverage Determinations**

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Note: Please review the current Medicare Policy for the most up-to-date information.

**Coding/Billing Information**

**Note:**
1) This list of codes may not be all-inclusive.
2) Deleted codes and codes which are not effective at the time the service is rendered may not be eligible for reimbursement.

Considered Experimental/Investigational/Unproven when used to report cervicography, spectroscopy or speculoscopy:
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<th>Description</th>
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<td>58999</td>
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**References**

combined testing, including Pap test, HPV DNA test and cervicography, as screening methods for the

2013 Jul 1;18(7):76013.


Guidelines Conference. 2012 updated consensus guidelines for the management of abnormal cervical

overview on applications of optical spectroscopy in cervical cancers. J Cancer Res Ther. 2008 Jan-

19. Nam K, Kim SN, Sim SH, Han S. The performance of tele-cervicography for detection of preinvasive and
invasive disease of the uterine cervix as an adjunctive test to Pap smears. Contemp Oncol (Pozn).

Accessed July 20, 2020. Available at URL address:

http://www.cancer.gov/cancertopics/factsheet/Risk/HPV

22. National Comprehensive Cancer Network® (NCCN). NCCN GUIDELINES™ Clinical Guidelines in

http://www.nccn.org/clinical.asp

at URL address: http://www.cancerscreening.nhs.uk/cervical/

6;8:1844-1862.


28. QIAGEN Inc. HC2 HPV DNA Test Hybrid Capture ® 2 High-Risk HPV DNA Test. Accessed July 20,
2020. Available at URL address: https://www.qiagen.com/us/products/diagnostics-and-clinical-
research/sexual-reproductive-health/cervical-cancer-screening/digene-hc2-high-risk-hpv-dna-test/#orderinginformation


