Diagnosis, Differential Diagnoses, and Classification of Diverticular Disease

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Societies of Gastroenterology (DGVS) and of Visceral Surgery (DGAV) has passed a new classification of diverticulitis displaying the different facets of diverticular disease. This classification addresses different types (not stages) of the condition, and includes symptomatic diverticular disease (SUDD), largely resembling irritable bowel syndrome, as well as diverticular bleeding.

Introduction

Diverticular disease, i.e. symptoms and/or complications due to diverticulosis of the colon, ranks 5th among the most costly gastroenterological diseases in the Western world [1]. Thus, and because the risk of diverticular disease increases with age, diagnosis will remain a pertinent challenge in the future. Even more challenging are i) the recognition of differential diagnoses particularly in relation to rather uncomplicated courses (e.g. segmental colitis associated with diverticular disease (SCAD) or symptomatic uncomplicated diverticular disease (SUDD) – the latter being probably not more than a variant of irritable bowel syndrome –, and ii) the accumulation of diagnostic information detailed enough to foster a classification that recognizes in detail any complicated course and thereby avoiding inappropriate undertreatment along with (antibiotic or surgical) overtreatment [2].

While the development of colonic diverticulosis per se is not considered a disease in itself, the condition may contribute to complications if other noxae are present, e.g. nonsteroidal anti-inflammatory drugs (NSAIDs; including increased bleeding risk under NSAIDs and aspirin), corticosteroids, opiates, or cigarette smoking.
Definition

Colonic diverticula are an acquired sacculation/outpouching of the mucosal and submucosal layers penetrating muscular leaks in the colonic wall at the site of the mucosa-supplying arteries. Among the prerequisites (i.e. functional such as increased luminal pressure, and morphological) for the development of colonic (pseudo-)diverticula, muscular hypertrophy is a diagnostic hallmark.

Diverticulitis is characterized by an inflammatory process starting usually within the diverticulum (occlusion by a fecalith and micropерforation) or at the neck of the diverticulum (ischemia or mechanical injury). Inflammation with micropерforation involves a peridiverticular mesenteric inflammatory reaction which may progress to pericolic and mural phlegmonous infiltration as well as fistulization, sealed perforation, abscess, free perforation, peritonitis, and a stenosing inflammatory sigmoid tumor. Another (distinct) complication of diverticular disease is bleeding [3].

The rather new term of SUDD must not be confused with uncomplicated diverticulitis because it does not rely on the criteria of diverticulitis (i.e. inflammation and imaging) but on local pain and, eventually, some inflammatory parameters (in the stool or serum) alone, thereby not distinguishing e.g. microbial infections of undetected etiology (such as SCAD) or irritable bowel symptoms from ‘true’ diverticulitis.

Smoldering diverticulitis on the other hand is a surgically coined phrase for those patients with symptomatic diverticulitis, in whom the diverticulitis remains obscure (in computed tomography (CT) scan, and sometimes also with barium enema and/or colonoscopy) until sigmoid resection is performed (histological diagnosis).

When Does Diverticulitis Occur – and What Does It Mean?

While the risk of diverticulosis and hospitalization for diverticulitis increases with age, the relative probability for developing diverticulitis decreases with age. Furthermore, the need for hospitalization and elective surgery showed highest growth rates in the younger (<45 years) population [4–6].

Non-elective hospitalization for ‘acute diverticulitis’ displays a remarkably stable seasonal cyclic undulation with highest incidence in the summer months [7]. Based on this pattern, it is tempting to speculate that undiagnosed ‘summer infections’ may contribute to the clinical picture, especially since recent first publications have questioned the necessity for antibiotic therapy in ‘uncomplicated diverticulitis’ [8].

The core question is whether or not uncomplicated and complicated diverticulitis are different intense expressions of the same disease or variants in a spectrum of various etiologies or pathogenetic mechanisms. The latter view is supported not only by the aforementioned cyclicity but also by the now established understanding that perforated diverticular disease usually occurs as the first manifestation and not as a complication of prior diverticulitis episodes with progressive age-dependent risk (as claimed by Parks’ theory). Furthermore, subtle differentiation of e.g. mesenteric inflammatory veno-occlusive disease (MIVOD), SCAD, NSAID lesions associated with diverticular disease (vs. NSAID-complicated diverticulitis), or prolapsing mucosal folds in diverticular disease requires more attention to properly understand diverticulitis with respect to diagnostic needs and therapeutic consequences [9–19].

The impact of careful differential diagnosis is underscored by the fact that approximately 80% of patients with acute diverticulitis treated at a surgical unit are discharged without operation [20, 21]. Thus, one could focus the current debate on i) mild episodes at the interface between ambulant medicine and hospitalization on the one hand, and ii) episodes at the interface between conservative clinical and operative treatment on the other hand.

As it is quite clear from a radiation exposure point of view (increasingly important with the decreasing age of the affected patients) that not every patient with suspected diverticulitis can and should undergo a CT scan, it has also become evident that not every patient with a minor perforation/small abscess must be operated on. As a consequence, however, without CT scan or operation there is no classification of diverticulitis for the vast majority of patients, because the hitherto used classifications (Hansen & Stock (HS), Hinchey) are based on either CT or operative criteria.

Classification of Diverticular Disease – Now and Then

Most classifications have been modified over time because weak points became evident when new aspects in diagnosis or therapy occurred. Hinchey’s classification (1978), based on that of Hughes (1963), focused on different expressions of abscess size and location. It was complemented by Sher (1997) and Wasvary (1999). The HS classification (Hansen & Stock, 1999) was used predominantly in Germany while other countries preferred Neff’s classification. Siewert (1995), Ambrosetti (2002), and Tursi (2008) suggested simplified classifications while a group from the Netherlands favored a more complex and sophisticated approach (Klarenbeck, 2012) [22–33].

The German guideline, recently (2014) passed by the German Society of Gastroenterology (DGVS) and of Visceral Surgery (DGAV), unanimously agreed on another classification (Classification of Diverticular Disease (CDD)), which takes practical algorithms (symptomatic, asymptomatic, complicated, uncomplicated, acute, recurrent), ongoing surgical aspects (purulent vs. fecal peritonitis), and contemporary diagnostic standards in clinical practice into account. As a result, this classification comprises the entire spectrum of diverticular disease; however, it is not tied to a specific diagnostic preference (such as CT vs. ultrasonography (US)) and it
Diagnosis and Classification of Diverticular Disease

Diagnosing Diverticulitis

The diagnosis of (sigmoid) diverticulitis in a patient with left lower quadrant pain requires both, proof of an inflammatory response (C-reactive protein (CRP) > white blood cell (WBC) count and erythrocyte sedimentation rate (ESR)) and localization of inflammation at the site of a diverticulum using an imaging method, i.e. US or CT [3].

According to the pathogenesis of diverticulitis, hypertrophy (and elastosis) with increasing width of the circular muscle layer is a constant trait with the diverticula developing at natural gaps associated with the penetration of arterioles supplying the (sub-) mucosal layers. Furthermore, diverticulitis starts in a single diverticulum only, which is usually the site of maximum pain under compression (although inflammation may secondarily involve further diverticula in the neighborhood, i.e. progression in longitudinal direction). Finally, inflammation usually starts in the outpouched mucosa in the diverticulum, i.e., it is endoscopically invisible unless inflammation spreads back to the mucosa from outside the wall, or a tear in the diverticular neck due to the passage ('birth') of a fecalith triggers diverticulitis (fig. 1a, b). Hence, the core question for cross-sectional imaging is not only whether an abscess/perforation is present or not, but also whether the aforementioned morphological criteria of diverticulitis are present or whether e.g. segmental colonic inflammation involves a diverticula-bearing segment only.

These considerations lend support to the view that clinical examination alone (signs and symptoms) is insufficient to diagnose diverticulitis.

**Table 1. Classification of diverticular disease (CDD)**

<table>
<thead>
<tr>
<th>Type</th>
<th>Definition</th>
<th>Symptoms</th>
</tr>
</thead>
<tbody>
<tr>
<td>Type 0</td>
<td>asymptomatic diverticulosis</td>
<td>random finding; asymptomatic; not a disease per se</td>
</tr>
<tr>
<td>Type 1</td>
<td>acute uncomplicated diverticulitis</td>
<td>symptoms attributable to diverticula; signs of inflammation (laboratory tests); optional; typical cross-sectional imaging</td>
</tr>
<tr>
<td>Type 1a</td>
<td>diverticulitis without peridiverticulitis</td>
<td>signs of inflammation (laboratory tests); optional; typical cross-sectional imaging</td>
</tr>
<tr>
<td>Type 1b</td>
<td>diverticulitis with phlegmonous peridiverticulitis</td>
<td>signs of inflammation (laboratory tests); mandatory; cross-sectional imaging; phlegmonous diverticulitis</td>
</tr>
<tr>
<td>Type 2</td>
<td>acute complicated diverticulitis</td>
<td>signs of inflammation (laboratory tests); mandatory; typical cross-sectional imaging</td>
</tr>
<tr>
<td>Type 2a</td>
<td>microabscess</td>
<td>concealed perforation, small abscess (&lt;1 cm); minimal paracolic air</td>
</tr>
<tr>
<td>Type 2b</td>
<td>macroabscess</td>
<td>paracolic or mesocolic abscess (&gt;1 cm)</td>
</tr>
<tr>
<td>Type 2c</td>
<td>free perforation</td>
<td>free perforation, free air/fluid; generalized peritonitis</td>
</tr>
<tr>
<td>Type 2c1</td>
<td>purulent peritonitis</td>
<td></td>
</tr>
<tr>
<td>Type 2c2</td>
<td>focal peritonitis</td>
<td></td>
</tr>
<tr>
<td>Type 3</td>
<td>chronic diverticular disease</td>
<td>relapsing or persistent symptomatic diverticular disease</td>
</tr>
<tr>
<td>Type 3a</td>
<td>symptomatic uncomplicated diverticulitis (SUDD)</td>
<td>localized symptoms; laboratory test (calprotectin); optional</td>
</tr>
<tr>
<td>Type 3b</td>
<td>relapsing diverticulitis without complications</td>
<td>signs of inflammation (laboratory tests); present; cross-sectional imaging: indicates inflammation</td>
</tr>
<tr>
<td>Type 3c</td>
<td>relapsing diverticulitis with complications</td>
<td>identification of stenoses, fistulas, conglomerate tumor</td>
</tr>
<tr>
<td>Type 4</td>
<td>diverticular bleeding</td>
<td>diverticula identified as the source of bleeding</td>
</tr>
</tbody>
</table>

**Fig. 1.** a Inflamed orifice of a diverticulum with occluding fecalith. This pattern is considered to reflect retrograde penetration of inflammation from the outpouched diverticulum. b Inflamed and torn mucosa in the orifice of an empty diverticulum. This pattern is considered to reflect the passage of a fecalith.
A position statement by the Association of Coloproctology of Great Britain and Ireland (ACPGBI) on elective resection for diverticulitis states that ‘patients should not be told that they have diverticulitis unless there is colonoscopic and/or radiological evidence of inflammation in the presence of diverticular disease’ [34]. Although there is no role for endoscopy in acute diverticulitis as long as cross-sectional imaging (US, CT) is diagnostic, this statement hints at the direction shared by the German guideline [3]: Imaging of the morphological substrate of diverticulitis (by either US or CT) is mandatory for the diagnosis of diverticulitis.

Signs of sigmoidal diverticulitis are also described by the term ‘left-sided appendicitis’ with the following symptoms: i) spontaneous pain in the left lower quadrant exaggerated by movement; ii) inflammatory reaction (CRP, WBC, temperature); and iii) local guarding upon palpation. This triad, however, is variable, time-dependent, and unspecific, and thus may raise the suspicion of diverticulitis without satisfying contemporary diagnostic requirements. Based on history and physical examination along with laboratory data, clinical judgment has a sensitivity of 65–70% only, a figure which is relatively consistent among different studies [35–38]. Therefore, a reliable diagnosis of diverticulitis requires cross-sectional imaging both in the ambulant setting and in hospital [3, 39].

### Differential Diagnoses

Clinically it ought to be emphasized that sigmoid diverticulitis may cause symptoms not only in the left lower quadrant but also in the right lower quadrant (elongated sigmoid loop) and anywhere in the lower abdomen. Right-sided diverticulitis even extends this spectrum towards the right upper abdomen. Thus, frequent differential diagnoses of diverticular disease/diverticulitis are inflammatory and non-inflammatory diseases of the gastrointestinal tract and the urogenital system, as well as vascular diseases (table 2).

### Diagnostic Steps

Clinical examination for diverticulitis comprises abdominal palpation, percussion and auscultation, examination of the inguinal rings, rectal examination, taking the temperature, urinalysis, and laboratory tests (with emphasis on CRP, WBC count, and ESR). Shifts in laboratory parameters may be delayed; therefore retesting after 2 days (‘48-h rule’) is prudent and improves safety for the benefit of the patient [3, 42, 43].
As already pointed out, imaging methods (US, CT) are deemed mandatory in addition to history taking and physical examination for an exact diagnosis, classification, and differential diagnosis. In contrast, there is currently no role for routine magnetic resonance imaging [3].

Because in Germany legal radiation protection applies according to § 23(1) of the German Radiation Control Regulation (Röntgenverordnung (RöV)) from 2011, radiology is only allowed ‘... if a justified indication applies. For such a balanced consideration other techniques with equivalent health benefit, which do not bear radiation hazards, must be taken into account’.

Thus, long in the shade of CT, it is time for US to take center stage for the following reasons: i) a meta-analysis certified ‘the best evidence for diagnosis of diverticulitis in the literature is on ultrasonography; only one small study of good quality was found on CT or MRI colonoscopy’ [44]; ii) US is applicable in all patients with suspected diverticulitis (e.g. outpatients and emergency cases); iii) it is cheap; iv) in addition to a reliable initial diagnosis, it allows close follow-up; and v) last but not least, it has higher resolution power than a CT scan.

**Ultrasonography – Why, How, Who, and Who not?**

US is applied directly at the point of pain and guarding, which usually reveals the inflamed diverticulum and/or associated complications (fig. 3a, b).

The inflamed diverticulum may or may not contain an echogenic and more or less crescent-shaped fecalith (see also fig. 1a, b), but once extruded, spontaneous drainage of pus into the colon is hypothesized to decrease pressure and the risk for perforation (fig. 4a, b) [45].

The core US finding of diverticulitis (figs. 3, 4) is ‘The diverticulum with different echogenicity in the centre of a pericolonic fatty tissue reaction’ (Hollerweger [46]), i.e. surrounded by an echogenic mesenteric cap, in conjunction with i) hypoechoic and initially asymmetrical wall thickening (>5 mm) with loss of wall layering, reduced wall compliance under pressure, and narrowing of the lumen, and ii) occasionally hypoechoic ‘inflammation lanes’ which are considered inflammatory exsudation (fig. 4b at 10:30 clockwise orientation).

Microperforations, fistulas, and abscesses are characterized by air bubbles in the mesentery, in a hypoechoic lane, or in an echopoor fluid retention, while free peritoneal air or air bubbles in the retroperitoneal space indicate free or retroperitoneal perforation.

In experienced hands, sensitivity and specificity of US are 98% [38]. In uncomplicated diverticulitis, direct visualization of the inflamed diverticulum amounts to 96%; however, it is more difficult if complications dominate (77%, specificity 99%) [46].

Deep abscesses, mesenteric air, and retroperitoneal perforation are the most difficult triad for US in this scenario, and the examiner should especially consider this if US produces an ‘unsatisfactory picture’ which is in contrast with an obviously ‘ill’ patient. This then becomes a case for CT; however, detection of diverticula or diverticular inflammation at CT has only insufficient sensitivity (30%) and interobserver agreement (despite an overall 99% sensitivity and specificity) [47, 48].

A comparative study with four experienced US examiners and CT scans performed at university level resulted in 100% sensitivity for US (vs. CT 98%) with 97% specificity for both procedures. Advanced peridiverticulitis and sealed perforations were overestimated by CT and rather underestimated at US, but free perforations and abscesses were missed by neither of the two techniques [49].

Often, it is argued that the outcome of US depends on the equipment and on the examiner. However, currently sold US devices, preferably with a ≥5 MHz probe, all meet the required standards and thus are throughout fully adequate for the diagnosis of diverticulitis and its differential diagnoses. With respect to the examiner, it must be clear that no medical technique – be it electrocardiogram, stethoscope, or CT – can ever be valid if the examiner
is not familiar with it. Adequate training in US for diverticulitis with production of valid results can be assumed after approximately 500 (targeted) US examinations [50]. US trainees with <500 examinations targeted to diseases of the gut should not issue US reports on diverticulitis on their own authority.

Computed Tomography – How, When, Why?

For the diagnosis of diverticulitis, abdominal CT scans are regularly performed with intravenous and oral contrast in the portal venous phase. Complementary rectal filling is recommended to provide better contrast at the rectosigmoid segment. Especially in obese patients, CT scans can at first also be performed satisfactorily without any contrast.

CT is a long-established and, at least from the surgeon’s perspective, valid technique for imaging acute diverticulitis and assessing its severity (complications) and differential diagnoses, thereby guiding e.g. the surgical strategy [21, 51]. Radiation exposure, however, precludes its use in all patients suspected to have an acute episode of diverticulitis.

In contrast to US, the key CT criteria do not include the inflamed diverticulum itself (which is detected in approximately 30% only), but rather perifocal ‘stranding’ (mesenteric inflammation), detection of free fluid, and a broadened colonic wall diameter of >3–5 mm [47].

A cornerstone study by Ambrosetti et al. [31] showed that age, abscess, and air are the triple-A clue to guide prognosis and surgical management. In detail, patients with severe CT findings (i.e. an abscess (median diameter ≥ 3 cm) or extracolonic air/contrast medium) had significantly more complications during their follow-up (36 vs. 17%) after conservative treatment of their first episode. Vice versa, it is noteworthy that two-thirds of patients with severe CT findings did not have such complications.

Currently, new publications may cast doubt on the unanimous faith in CT scans for acute diverticulitis. In patients operated on for perforation, accuracy of CT was found to be 71–92% (positive predictive value (PPV) 45–89%) in different stages. In 42% of patients at Hinchey stage III, CT led to understaging (Hinchey I and II), resulting in a PPV of CT of 61% in Hinchey stages I and II only [52].

Another comparison of preoperative CT findings with both the situs and the histology at surgery showed correct staging by CT at HS IIa in 52% (operation) and 56% (histology). For this stage, understaging occurred in 12 (11%), overstaging in 36 (33%). In this study, validity for stages with an abscess (HS IIb, Hinchey I and II) was much better (operation 92%, histology 90%), and free perforation (HS IIc, Hinchey III and IV) was detected in 100% resulting in a PPV for CT of 61% in Hinchey stages I and II only [52].

Such sizeable understaging (42%) in perforating diverticulitis [52] and overstaging (1/3) in phlegmonous diverticulitis [53] demonstrates that CT was probably the best technique available in the past but that it may be hard to accept it as gold standard. Nevertheless, in comparison with US, CT has definite advantages in the detection of distant mesenteric and pelvic abscesses.

Fig. 5. Abscess (type 2b) after colonoscopy in acute sigmoid diverticulitis (arrows).

Fig. 6. Diagnostic algorithm for suspected diverticulitis.
Colonic – When and Why?

Colonscopy is the diagnostic (and therapeutic) method of choice in patients with gastrointestinal bleeding and thus also in diverticular bleeding.

The current guideline [3], however, advises against the use of colonoscopy for diagnosing acute diverticulitis because it is i) inadequate and ii) potentially dangerous (increasing the risk or intensity of perforation). Figure 5 shows an abscess in such a patient who underwent colonoscopy during an episode of acute diverticulitis (in another hospital) and complained of more severe pain thereafter. This in conjunction with recovery failure prompted her family doctor to admit her for surgery.

In uncharacteristic cases, however, colonoscopy might be useful or mandatory after exclusion of perforation, because otherwise differential diagnoses such as SCAD, NSAID-induced lesions, ischemic colitis, right-sided diverticulitis, or foreign body-induced diverticular injury (fig. 2a, b) might not be detected or adequately assessed. In this scenario and gently performed by an experienced examiner, colonoscopy is regarded a safe procedure by most authorities [3] (see also the Interdisciplinary Discussion on this topic in this journal). It is noteworthy that inflammation at the orifice of diverticula is observed in approximately 0.8% of colonoscopies without evidence of diverticulitis [54].

After an episode of acute diverticulitis, colonoscopy is recommended in patients who are >50 years of age and who have not had a colonoscopy before, and also in patients who are scheduled to undergo surgery for diverticular disease. In these cases, a safety interval between acute diverticulitis and colonoscopy of 6 weeks appears advisable.

Algorithm for Diagnosing Diverticulitis

The above reassessed facts, considerations, and points of view form the basis for a summarizing diagnostic algorithm shown in figure 6, which allows tailored adaptation to the different prevailing clinical situations and facilities [3].

Disclosure Statement

The author has no conflict of interest to declare.

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4. Strate LL, Modi R, Cohen E, Spiegel BM: Diverticular diverticular injury (fig. 2a, b) might not be detected or adequately diagnosed. Optimum management requires accurate clinical situations and facilities [3].
5. Shahedi K, Fuller G, Bolus R, et al: Long-term risk of perforation. Figure 5 shows an abscess in such a patient who underwent colonoscopy during an episode of acute diverticulitis (in another hospital) and complained of more severe pain thereafter. This in conjunction with recovery failure prompted her family doctor to admit her for surgery.

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