# Paediatric Gastrointestinal Endoscopy: **European Society for Paediatric** Gastroenterology Hepatology and Nutrition and European Society of Gastrointestinal Endoscopy Guidelines

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#### ABSTRACT

This guideline refers to infants, children, and adolescents ages 0 to 18 years. The areas covered include indications for diagnostic and therapeutic esophagogastroduodenoscopy and ileocolonoscopy; endoscopy for foreign body ingestion; corrosive ingestion and stricture/stenosis endoscopic management; upper and lower gastrointestinal bleeding; endoscopic retrograde cholangiopancreatography; and endoscopic ultrasonography. Percutaneous endoscopic gastrostomy and endoscopy specific to inflammatory bowel disease has been dealt with in other guidelines and are therefore not mentioned in this guideline. Training and ongoing skill maintenance are to be dealt with in an imminent sister publication to this.

Keywords pediatric, esophagogastroduodenoscoy, ileocolonoscopy, colonoscopy, ESPGHAN guidelines, ESGE guidelines

(JPGN 2017;64: 133-153)

## INTRODUCTION

astrointestinal (GI) endoscopy in the pediatric population has evolved during the last 30 years with an increasing number of diagnostic and therapeutic applications. Technological improvements in endoscope design and endoscopic devices have contributed to the evolution of pediatric endoscopy.

Endoscopy in the pediatric population has generally, to date, been performed by both nonpediatric endoscopists and pediatric endoscopists.

The aim of this evidence-based and consensus-based guideline, commissioned by the European Society for Paediatric Gastroenterology Hepatology and Nutrition (ESPGHAN) and the European Society of Gastrointestinal Endoscopy (ESGE) is to provide a comprehensive review of the clinical indications and timing of diagnostic and therapeutic endoscopy in pediatric

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The authors report no conflicts of interest.

Copyright © 2016 by European Society for Pediatric Gastroenterology, Hepatology, and Nutrition and North American Society for Pediatric Gastroenterology, Hepatology, and Nutrition DOI: 10.1097/MPG.000000000001408

Received May 20, 2016; accepted September 2, 2016.

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patients. It is not meant to be a comprehensive overview of a patient's care and investigation/therapy for each area will, of course, involve the clinician's discretion in terms of the place of endoscopy in overall management, encompassing, as it must, complementary nonendoscopic approaches. The role of endoscopy in the overall management will depend on a number of factors including but not limited to the specific clinical features, the availability/appropriateness of nonendoscopic approaches, and the available skills of the endoscopist. This Guideline tries to address this issue of endoscopist skills, and certainly the upcoming ESPGHAN/ESGE Guideline on training in pediatric endoscopy will help in this respect. How, where, and when endoscopy may be employed in pediatric management is particularly important in the areas of GI bleeding and endoscopic retrograde cholangiopancreatography (ERCP)/endoscopic ultrasonography (EUS).

This undertaking is the first joint endoscopy review between pediatric and adult endoscopy representative groups in Europe. Our aspiration is that the guideline may lead to a degree of standardization in the utility and practice of endoscopic approaches for children, thereby contributing to excellence and appropriateness of care.

Percutaneous endoscopic gastrostomy and endoscopy specific to inflammatory bowel disease (IBD) have been dealt with in other Guidelines (1-3), and are therefore not mentioned in the pediatric GI endoscopy (4) guideline. Training and ongoing skill maintenance will be addressed in an imminent sister publication.

#### **METHODS**

ESPGHAN and ESGE agreed to develop a joint guideline. Two guideline leaders (M.T. for ESPGHAN and A.T. for ESGE) invited the listed authors to participate in the project. The key questions were prepared by the coordinating team (A.T., M.T., M.M.T., R.F., Y.V., J.M.D.) and then approved by the other members. The coordinating team established task force subgroups, each with its own leader, and assigned the following key topics among these task forces: esophagogastroduodenoscopy (EGD) and ileocolonoscopy (IC), foreign bodies (FBs), corrosive ingestion, corrosive ingestion and esophageal strictures/stenoses, GI bleeding, ERCP, and EUS. Each task force performed a systematic literature search to prepare evidence-based and well-balanced statements on their assigned key questions. Searches were performed in PubMed and/or EMBASE and/or Cochrane (publication year from 2000 to May 2015 or before if strictly needed) including as a minimum the key words "pediatric" and "endoscopy." All articles studying the application of diagnostic and therapeutic endoscopy in the pediatric age range were selected by title or abstract. The results of the relevant publications were summarized in literature tables and graded by the level of evidence and strength of recommendation according to the GRADE system (Grading of Recommendations Assessment, Development and Evaluation) (3,5). Each task force proposed statements on their assigned key questions which were discussed and voted on during the plenary meeting held in February 2015 in Munich. In November 2015, a draft prepared by A.T., C.H., and M.T. was sent to all group members. After agreement from all the authors on a final version, the manuscript was reviewed by 2 members of the ESGE Governing Board, ESGE individual members, and the ESPGHAN Council. The manuscript was then submitted to the Journal of Pediatric Gastroenterology and Nutrition for publication in full length, and to Endoscopy for publication of an executive summary. Both the Guideline and Executive summary were issued in 2016/2017 and will be considered for review and update in 2021/

2022 or sooner if new and relevant evidence becomes available. Any updates to the guideline in the interim will be noted on the ESPGHAN and ESGE and Web sites: http://www.espghan.org/guidelines/; http://www.esge.com/esge-guidelines.html.

# ESOPHAGOGASTRODUODENOSCOPY

ESGE/ESPGHAN suggests diagnostic and therapeutic EGD for the indications listed in Tables 1 and 2, respectively.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN does not suggest EGD in the case of uncomplicated gastroesophageal reflux, functional gastrointestinal disorders, or for diagnosing perforation.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests routine tissue sampling even in the absence of visible endoscopic abnormalities in all children undergoing EGD.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests using ESPGHAN guidelines (eosinophilic esophagitis [EE], eosinophilic esophagitis, Helicobacter pylori [H pylori], celiac disease, and IBD) for precise indications and preferred sites for biopsy during EGD in children suspected of a specific disease (Table 3).

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests performing EGD in children under general anesthesia (GA) or, only if GA is not available, deep sedation in a carefully monitored environment.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests performing EGD in a childfriendly setting with appropriate equipment and by an endoscopist trained in pediatric gastroenterology.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that when adult endoscopists perform pediatric procedures, collaboration between adult gastroenterologists and pediatricians is always warranted.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that the choice of the gastroscope type should depend on the child's weight and age (Table 4).

(Weak recommendation, low quality of evidence)

EGD is a useful diagnostic and therapeutic tool in children (6), from which information can be obtained from visualization and biopsy of the mucosal surfaces of the esophagus, stomach, and duodenum. Although one third of children have a sore throat or hoarseness after EGD, EGD is generally considered to be safe for all ages (7). In a pediatric cohort including 345 procedures (231 EGD alone, 26 colonoscopy alone, 44 combined EGD and colonoscopy) in 301 children with a median age of 7 years, 20 (5.8%) adverse events were reported (12 secondary bleeding following variceal banding/sclerotherapy, 2 colonoscopy-related perforations, 6 anesthesia related) (8). Fourteen events were procedure-related (12 secondary bleeding after banding or sclerotherapy, 2 bowel perforations during colonoscopy) and 6 were anesthesia/sedation related. None of the adverse events were fatal. It is, however, important to minimize risk of complications, that EGD only should be performed for appropriate indications and by well-trained endoscopists (6). A diagnostic EGD is indicated in the presence of symptoms listed in Table 1 to confirm an underlying disease. A selection of therapeutic indications is also listed in Table 2. Nonindications are uncomplicated gastroesophageal reflux and functional GI disorders. Contraindications include diagnosis of perforation (Table 1).

Routine tissue sampling according to the indication, even in the absence of visible endoscopic abnormalities, is of major

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Diagnostic indications	Therapeutic indications	Nonindications	Contraindications
Weight loss, failure to thrive	Percutaneous endoscopic gastrostomy (re)placement	Uncomplicated GERD	To diagnose perforation
Unexplained anemia	Duodenal tube placement	Functional GI disorders	
Abdominal pain with suspicion of an organic disease	Foreign body removal		
Dysphagia or odynophagia	Food impaction		
Caustic ingestion	Hemostasis		
Recurrent vomiting with unknown cause	Percutaneous jejunostomy placement		
Hematemesis	Esophageal varices		
Hematochezia	Dilatation of esophageal or upper GI strictures		
Unexplained chronic diarrhea	Perforation closure if this occurs during an endoscopy itself		
Suspicion of graft-versus-host disease	Achalasia pneumodilation or occasionally botulinum injection		
GI allergy	PEGJ tube insertion		
	Cystogastrostomy for drainage of pancreatic pseudocyst (preferably with endoultrasound guidance)		
Chronic GERD to exclude other diseases or surveillance of Barrett esophagus	Polypectomy, endomucosal resection		

#### TABLE 1. Typical diagnostic and therapeutic indications, nonindications and contraindications for EGD in pediatric patients

EGD = esophagogastroduodenoscopy; GERD = gastroesophageal reflux disease; GI = gastrointestinal; PEGJ = percutaneous endoscopic gastro-jejunostomy.

importance in all children undergoing EGD. Two studies in children assessed the value of routine esophageal, gastric, and duodenal biopsies and new diagnoses based on biopsy samples alone were identified in 17% and 11% (9,10). A study including 823 infants younger than 1 year of age, a group in which both symptoms and signs are notoriously difficult to interpret, the histological findings during EGD and/or colonoscopy were

helpful in diagnosis in 63.8% of the cases (11). One pediatric study showed that biopsies from the first and third part of the duodenum were important when assessing a patient for suspected celiac disease: biopsies from the duodenal bulb had an incremental diagnostic yield of 10.6% compared with biopsies only from the third part of the duodenum (12). Table 3 sets out the ideal location for biopsies to allow the greatest diagnostic yield

TABLE 2. Diagnostic indications for EGD in pediatric patients and symptoms/signs according to suspected disease

Symptoms/signs	Suspicion of:			
Weight loss, failure to thrive, chronic diarrhea, malabsorption, anemia, abdominal pain with suspicion of an organic disease	Celiac disease, IBD, giardia, allergic enteritis/enteropathy, bleeding lesions, graft-versus-host disease, peptic ulcer disease			
Dysphagia, odynophagia, chest pain, feeding difficulty	Foreign-body ingestion, food impaction, postcaustic ingestion, eosinophilic esophagitis, achalasia, aberrant vasculature affecting the esophagus, congenital webs, or other abnormalities such as Schatzki ring, stricture postsurgical, eg, postrepair of tracheoesophageal fistula			
Hematemesis, hematochezia, melena	Polyps, angiodysplasia, arteriovenous malformations, peptic ulcer with or without <i>H pylori</i> infection, less common conditions such as duplication cysts			
Family history of polyposis syndromes	Polyps (diagnostic and surveillance)			
Vomiting	Any obstructive or partially obstructive pathology involving the upper GI tract, for example, pyloric stricturing, duodenal webs and strictures, bezoars, superior mesenteric artery syndrome, and so on; allergic GI pathology; peptic ulceration; assessment of esophagitis associated with reflux.			

EGD = esophagogastroduodenoscopy; GI = gastrointestinal; IBD = inflammatory bowel disease.

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TABLE	3.	Indicat	ion	and	site	of	tissue	sampling	during	upper	and
lower of	enc	loscopy	in	pedia	atric	pat	tients				

Indication	"Tissue samples: sites and numbers"		
Eosinophilic esophagitis	At least 3 biopsy sites should be targeted with 1–2 biopsies from proximal, middle, and distal esophagus, regardless of the endoscopic appearance of the esophagus		
Helicobacter pylori infection	Six biopsies (2 from antrim and 2 from corpus for Sydney classification; 2 for specific <i>H pylori</i> diagnosis: CLO and culture)		
Celiac disease	At least 1 biopsy from the duodenal bulb and at least 4 biopsies from the second or third portion of the duodenum		
IBD	Multiple biopsies (2 or more per section) from all sections of the visualized GI tract, even in the absence of macroscopic lesions		

GI = gastrointestinal; IBD = inflammatory bowel disease.

with respect to suspected diagnosis (1,13-15). In contrast to adults, in children EGD should be performed with GA or, if not available, under deep sedation with a specifically trained pediatrician in charge only of the sedation leaving the endoscopist to concentrate on the procedure alone. Propofol-based sedation is likely to be the safest and most convenient way of sedation; however, this remains the subject of debate (16). Furthermore, endoscopy should be performed in a child-friendly setting. This is an important point and pertains to not only the child but also the family. The "journey" that a child and their parents/carers take should involve wherever possible a previsit to the unit, a play specialist to allow an age-specific approach to prepare the child, a nonthreatening environment with age-appropriate wall decorations and toys, an anesthetist with the requisite human skills to allay the fears of the child and their family, and a recovery area that is child specific with parents/carers invited to be present before their child fully wakes. A multidisciplinary team consisting of a pediatric anesthetist, pediatric gastroenterologist, endoscopic nurse, and specialized pediatric nurses should be available to take care of the specific needs of pediatric patients. In some hospitals, adult endoscopists are needed to perform advanced therapeutic procedures, which are not routinely performed by a pediatric gastroenterologist and these are dealt with further on in this guideline; however, the advanced training of pediatric endoscopists is occurring and it is envisaged that upper GI endotherapeutic procedures in the near future in children will and should be performed routinely by such individuals, as already occurs in a small number of supraregional centers at present. A retrospective study suggested that "adult" endoscopists when supported by pediatricians are able to safely conduct EGD and IC in children, but this Guideline Group suggest that this should not be the ideal arrangement and that pediatric endoscopists should manage the child and perform the procedure thereby providing a seamless service (17). The choice of the equipment including the actual endoscope depends on a child's weight and age. Table 4 details the different endoscope sizes as adapted from American Society for Gastrointestinal Endoscopy (18).

#### ILEOCOLONOSCOPY

ESGE/ESPGHAN suggests IC for the diagnostic and therapeutic indications listed in Table 5.

(Weak recommendation, low quality of evidence) ESGE/ESPGHAN suggests against IC in the case of toxic megacolon, recent colonic perforation (<28 days), recent intestinal resection (<7 days) or functional GI disorders.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests performing IC in children under GA or, only if GA is not available, deep sedation in a carefully monitored environment.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that IC should be performed in a child-friendly setting with appropriate equipment and by an endoscopist trained in pediatric gastroenterology.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that when nonpediatric endoscopists perform pediatric procedures in older children, collaboration with a pediatrician is always warranted.

(Weak recommendation, low quality of evidence) ESGE/ESPGHAN suggests that the choice of the colonoscope type should depend on child's weight and age (Table 4). (Weak recommendation, low quality of evidence)

In children, IC is primarily indicated for suspected IBD, in cases of per-rectal bleeding and/or unexplained anemia (6) and for genetic polyposis syndromes including familial adenomatous polyposis (19-21) (Table 5). The recommendations concerning environment, endoscopists, and type of endoscopes are similar to those formulated for EGD. There are no published data to support specific colonoscope choice in children but recommendations based on experience state that the lower weight limit for use of a standard adult colonoscope is approximately 10 kg (Table 4). The largest safety report was a 5-year retrospective database study of 7792 procedures with an overall complication rate of 1.1% of which approximately 50% were GI related, most commonly bleeding, 30% were cardiopulmonary complications, and 10% were miscellaneous, which included allergic drug rash reactions amongst others. Perforation was uncommon (0.01%) (22) and as in adults, perforations in children are mostly due to advancement of the endoscope itself or are related to polypectomy, not to biopsy.

# BOWEL PREPARATION FOR ILEOCOLONOSCOPY IN CHILDREN

ESGE/ESPGHAN recommend low-volume preparation for bowel cleansing in children, using either polyethylene glycol (PEG) along with ascorbate or picosulfate magnesium citrate/ Senokot.

(Strong recommendation, high quality of evidence) ESGE/ESPGHAN recommend against the use of sodium phosphate for bowel cleansing.

(Strong recommendation, high quality of evidence)

Success and safety of IC relies very much on the quality of bowel preparation. In adults, ESGE Guidelines recommend a lowfiber diet on the day preceding IC and a split regimen of 4 L PEG solution; alternatives include a split regimen of 2 L PEG along with ascorbate or of sodium picosulfate and magnesium citrate (23). The ESGE advised against the use of oral sodium phosphate due to the risk of renal insufficiency. Furthermore, PEG is the only recommended regimen in patients with renal failure (23). In children, a national working group performed a systematic review and a national-based survey of all endoscopy units performing IC in Israel (24); 6 different protocols were compared, but none of these showed significant advantages. Another publication concluded that

Weight or age	EGD	Colonoscopy	ERCP	EUS
<10 kg or <1 year	≤6 mm gastroscope preferred. Consider standard adult gastroscope if endotherapy required.	≤6 mm gastroscope, standard adult gastroscope, or pediatric colonoscope (<5-8 kg or <6 months then an upper GI endoscope can be used and the size of the scope would depend on the size of the child—neonatal colonoscopy is rare but may require a pediatric upper GI endoscope)	7.5 mm duodenoscope	Miniprobe or 7.4 mm EBUS-scope
$\geq 10 \text{ kg or } \geq 1 \text{ year}$	Standard adult gastroscope. Therapeutic gastroscope if needed.	Pediatric or adult colonoscope	Therapeutic duodenoscope. (4.2 mm operative channel)	Miniprobe or 7.4 mm EBUS scope
$\geq$ 15 kg or $\geq$ 3 years	-	-	-	Adult radial/linear echoendoscope

#### TABLE 4. Type/size of gastrointestinal endoscope in pediatric patients according to body weight

EBUS = endoscopic retrograde cholangiopancreatography; EUS = endoscopic retrograde cholangiopancreatography; EUS = endoscopic ultrasonography; GI = gastrointestinal.

the most used agents in children include PEG-3350 solutions, picolax, senna, bisacodyl, and magnesium salts (25); their efficacy was found to be similar. Recently a randomized controlled trial (RCT) including 299 children evaluated 4 different regimens (26): the 3 low-volume regimens were noninferior in terms of bowel cleansing compared with the "high-volume" regimen (PEG at a dose of 100 mL/kg, maximum 4 L). Of note, low-volume regimens were better tolerated and were associated with a less frequent need for nasogastric tube placement compared with the high-volume

regimen. The authors suggested that the most suitable low-volume preparation was sodium picosulfate along with magnesium citrate. Regimens using sodium picosulfate with magnesium citrate (sodium picosulfate 0.01 g, magnesium oxide 3.5 g, citric acid 12.0 g per sachet) are used as follows: 2 doses 5–10 hours apart (0.25 sachet/dose for <6 year, 0.5 sachet/dose for 6 to 12 years, 1 sachet/dose for >12 years) with liberal drinking of clear fluids such as cold tea/sport drinks and approximately 40 mL/kg after each dose.

TABLE 5. Typical diagnostic and therapeutic indications, nonindications, and contraindications for ileocolonoscopy in pediatric patients					
Diagnostic indications	Therapeutic indications	Nonindications	Contraindications		
Unexplained anemia	Polypectomy, endomucosal resection or extended submucosal dissection, and removal of sessile polyps	Functional GI disorders	Toxic megacolon		
		Constipation with normal fecal calprotectin			
Unexplained chronic diarrhea	Dilatation of ileocolonic or colonic stenosis		Recent colonic perforation		
Perianal lesions (fistula, abscess)	Treatment of bleeding lesions		Recent intestinal resection (<7 days)		
Rectal blood loss	Foreign body removal				
Unexplained failure to thrive	Reduction of sigmoidal volvulus				
Initial work up for IBD	Cecostomy or sigmoidostomy				
Suspicion of graft-versus-host disease					
Rejection or complications after intestinal transplantation					
Radiological suspicion of ileocolonic stenosis/stricture					
Polyposis syndromes					
Orofacial granulomatosis when Crohn disease is suspected					

GI = gastrointestinal; IBD = inflammatory bowel disease.

# ILEOCOLONOSCOPY IN CHILDREN: BIOPSY, CARBON DIOXIDE INSUFFLATION, ILEAL INTUBATION, POLYPECTOMY TECHNIQUE

ESGE/ESPGHAN suggests routine biopsy even in the absence of visible endoscopic abnormalities in all children with suspected IBD undergoing IC.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests using ESPGHAN guidelines relating to ulcerative colitis and the revised Porto criteria for diagnosis of IBD for precise indications and preferred sites to biopsy.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN did not find any evidence to recommend against or for the use of routine CO2 insufflation during IC in children. Pain seems to be rare and mild after IC in children.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that ileal intubation should be attempted in symptomatic children with abdominal pain, intestinal bleeding, diarrhea, or with any suspicion of IBD.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests removal of small polyps (<3 mm) by cold biopsy forceps and 3 to 8 mm polyps by hot or cold snaring. Cold snaring is advisable in the right colon where the perforation risk is higher. For polyps >8 mm, hot snaring is suggested.

(Weak recommendation, low quality of evidence)

A study on 390 pediatric ileocolonoscopies reported 84% agreement between endoscopists and pathologists, especially when an endoscopist reports normal-appearing colonic mucosa and if histology was considered the criterion standard, endoscopy was found to have a 90% sensitivity and 78% specificity (27). When children present with diarrhea, abdominal pain, weight loss, or other symptoms in which initial investigations are normal and in the absence of macroscopic lesions of the colon, biopsies should be taken from different colonic segments to exclude conditions such as collagenous or microscopic colitis (28). Collagenous colitis is rare in children. One small study found that 5 of 26 children with chronic diarrhea and macroscopically normal colonic mucosa had histological abnormalities (3 lymphocytic colitis and 2 collagenous colitis) (29). In suspected pediatric IBD, recent ESPGHAN Guidelines highlighted the importance of biopsies in all segments of the lower digestive tract to differentiate Crohn disease from ulcerative colitis and to determine the extent of the inflammatory process (2,15). At the initial diagnostic stage, the presence of a granuloma allows differentiation between Crohn disease and ulcerative colitis and when combined with EGD can make this distinction in up to 15% to 20% of cases over and above that which is made by IC alone (30). Granulomas are more frequently observed when the biopsies are performed at the edges of ulcerative lesions (31,32). In severe acute colitis, a careful examination limited to the rectum and sigmoid may be performed initially because of the risk of perforation when the risk/benefit is managed by the possibility of performing a subsequent IC (2). Other pathologies such as acute graft-versus-host disease (GVHD) can be safely investigated using endoscopy and in 1 retrospective study of 48 children a sensitivity of rectosigmoid biopsies of 77% for GVHD diagnosis was reported (33). Biopsies taken proximal to the rectosigmoid only contributed to the GVHD diagnosis in 2 of 48, however. This compares to a study of adults with acute intestinal GVHD in the lower GI tract, of whom 20% had lesions only in the ileum (34). To increase the sensitivity of endoscopic exploration for suspected intestinal GVHD, a lower GI endoscopy may be accompanied by an EGD.

Two meta-analyses studied the usefulness of  $CO_2$  insufflation during colonoscopy in adults (35,36). Both found that  $CO_2$  insufflation significantly reduced pain during and after colonoscopy (35,36). In a retrospective pediatric population, post-IC pain was reported using CO<sub>2</sub> only by 2 of 68 (3%) children (37). A recent RCT (38) compared insufflation of CO<sub>2</sub> versus air during IC in children 7 to 18 years of age: CO<sub>2</sub> insufflation significantly decreases postprocedural discomfort. CO<sub>2</sub> has been used in large series of double balloon enteroscopy (DBE) in children as the rapidity of gas reabsorption is particularly useful during this procedure—no adverse events such as clinically significant rise in blood CO<sub>2</sub> were identified (39,40). Caution should be taken in small children because the amount of insufflated CO<sub>2</sub> could induce adverse effects due to the smaller blood volume of young children.

A registry of newly diagnosed IBD cases in children in 44 centers in 18 countries reported a successful intubation of the ileum in 75% of 1995 colonoscopies (41). In an adult study of 500 consecutive colonoscopies in which the ileum intubation rate was 99%, time and probability of ileal intubation were significantly correlated to the quality of bowel cleansing and to the experience of the endoscopist (42). In a pediatric cohort of 44 complete colonoscopies with an ileal intubation rate of 61%, ileal examination did not modify the patient-reported symptoms after endoscopy but this is essentially irrelevant as the diagnosis clearly needs to be established histologically to inform subsequent management (37). The aim of 100% terminal ileum intubation is to be highlighted in children especially in the IBD situation (43).

## Polyps

In a study including 11,637 children, polypectomy was performed in 6.1% of procedures and this rose to 12% in which lower intestinal bleeding was a symptom (44). Performing a polypectomy, the endoscopist has 3 goals: to remove the lesion; to retrieve it for histological examination; and to avoid adverse events. The 3 main adverse events of polypectomy are bleeding, perforation, and postpolypectomy syndrome, also known as transmural burn syndrome. The specific technique of polypectomy is generally chosen based on polyp localization, morphology, and size. In an RCT adult study, the use of a cold snare versus cold forceps polypectomy to remove diminutive (≤5 mm) polyps was significantly correlated with a shorter procedure time and a more complete polyp eradication rate (93% vs 76%, respectively) (45). Cold snaring is a safe technique with no adverse events reported in large series in adults (46). Hot biopsy forceps on the other hand induced a larger histological lesion compared with conventional snare procedure in a porcine model, especially when the snare diameter is large (5 vs 2.5 mm) (47) and necrosis depth is increased after hot forceps polypectomy with a more frequent inflammatory reaction under the submucosa due to the smaller area for electrical current diffusion compared with polyp snares. Cytological artifacts are more frequent in polyps removed by hot forceps technique compared with cold forceps (48). For both of these reasons it is suggested that hot forceps polyp removal should be avoided in children.

## FOREIGN BODY INGESTION

ESGE/ESPGHAN recommended an early referral to the emergency room and x-ray evaluation in all patients with suspected FB ingestion, even if asymptomatic. Biplane radiographs should be obtained of the neck, chest, abdomen, and pelvis if indicated. Computed tomography (CT) scan can be considered for radiolucent FBs.

(Strong recommendation, moderate quality of evidence) ESGE/ESPGHAN suggests early EGD if the FB is in the esophagus.

#### (Weak recommendation, low quality of evidence)

The approach to the endoscopic management of FBs should take in to consideration the type (food, batteries, magnets, sharp, blunt, drug packets, and size), the symptoms, the time since probable ingestion, the probable GI location, any suspected impaction, and so on. In the case of batteries symptoms are immaterial especially if the battery is impacted in the esophagus. Indeed any patient who is symptomatic with an ingestion of a sharp or a blunt FB should have endoscopic removal attempted. The standard approach to resuscitation of Airway, Breathing and Circulation is pertinent in this clinical context. If drooling is present and the patient is not able to swallow their secretions, there is a risk of aspiration. In cases of proximal esophageal FB ingestion, it will be necessary to ensure airway protection and endoscopy for FB removal should be performed under GA (49).

FB ingestion leading to impaction and food bolus impaction are quite common and the majority occur in the younger child with a peak incidence between the ages of 6 months and 6 years (49).

Pre-endoscopic series have shown that 80% or more of FBs will pass without the need for any intervention. Mortality due to FB ingestion was not reported in a large pediatric series (50,51). A case of death has been reported in a 2-year-old boy due to an aortoe-sophageal conduit caused by an impacted sharp FB in the esophagus (52). The patient's age and size, the type and form of the ingested object, its location and the clinical symptoms, and duration since ingestion will all contribute to the decision whether to intervene endoscopically and to the timing of any intervention.

Symptoms associated with FB ingestion varied among studies from vomiting, drooling, dysphagia, odynophagia, globus sensation, and also included respiratory symptoms of coughing, stridor, and choking. Some children are completely asymptomatic. In 9 of 12 studies (53) in which coins were most the most frequent or only FB ingested, vomiting and drooling were the predominant symptoms. It should also be remembered that evaluation for peritonitis or small-bowel obstruction should occur in any case of FB ingestion and in such situations endoscopy should not delay surgical consultation but simultaneous endoscopy can complement the surgical approach (49,54,55).

For the purpose of initial diagnosis, radiographs can confirm the location, size, shape, and number of ingested FB and can help to exclude aspirated objects (49,56). Radiographs identify most radio-opaque FBs but radiolucent FBs are common, limiting the reliability of radiographs in this initial evaluation (56). Fish bones, wood, plastic, and thin metal objects are some of the most common radiolucent objects (49,56). Thin fragments of aluminum, such as pull-tabs or pop-tabs of beverages, present a relatively high radiolucency (57). Biplane radiographs should be obtained of the neck, chest, abdomen, and pelvis if indicated. In addition to localization of radio-opaque objects, the presence of free mediastinal or peritoneal air should be assessed. A contrast examination should not be performed routinely in the patient with suspected proximal esophageal obstruction because of the risk of bronchoaspiration. Furthermore, opaque contrast agents, such as barium, coat the FB and esophageal mucosa, compromising subsequent endoscopy and Gastrografin (amidotrizoeacid), a hypertonic nonopaque contrast agent, which can produce a severe chemical pneumonitis if aspirated, should not be used. There are no pediatric studies evaluating CT scan in the diagnosis of FB ingestion in the digestive tract. A pediatric study (58) showed a 93% positive predictive value and 100% negative predictive value using spiral and cine CT scan in the diagnosis of radiolucent FBs in the airways. CT scanning can be considered in the diagnosis of radiolucent ingested FBs in selected cases considering also the risk of x-ray exposure in children. There is not enough evidence for use of metal detectors or ultrasonography in localizing

ingested coins in children (59,60). Magnetic resonance imaging is not helpful in detecting FBs (61).

## **BLUNT FOREIGN BODIES AND COINS**

ESGE/ESPGHAN recommends removal of blunt FBs and coins or impacted food from the esophagus urgently (<24 hours), even in asymptomatic children. If the child is symptomatic an emergent (<2 hours) removal is indicated especially for button batteries (BBs).

(Strong recommendation, moderate quality of evidence)

ESGE/ESPGHAN suggests removal of blunt FBs from the stomach or duodenum if the child is symptomatic or if the object is wider than 2.5 cm in diameter or >6 cm in length. Otherwise blunt FBs in the stomach can be followed and retrieved only if they produce symptoms or do not pass spontaneously after 4 weeks.

(Weak recommendation, low quality of evidence)

As noted above the timing of endoscopy depends on a number of factors including age, the patient's clinical status, the time of the patient's last oral intake, type of FB ingested, location within the GI tract, and the time that has elapsed since ingestion. In addition, an assessment of the relative risk of aspiration, obstruction, or perforation may determine the timing of any endoscopy (49). Generally speaking, timing can be divided into emergent (<2 hours from presentation, regardless of nil by mouth status), urgent (<24 hours from time of ingestion) and elective (>24 hours from ingestion). Patients who are clinically stable without symptoms of proximal esophageal obstruction do not require emergent endoscopy because the ingested FB will usually pass spontaneously (49). Even in asymptomatic children esophageal FBs and food that has, however, impacted in the esophagus should be removed urgently (<24 hours from presentation) as any delay decreases the likelihood of successful removal and increases the risk of adverse events, including the risk of perforation. These data are based on adult studies as those in children are not available (49,62,63). If the FB is located in the stomach and there is no risk of impaction distally (eg, due to strictures) then most FBs will pass in 4 to 6 days. Therefore conservative outpatient management is appropriate for most asymptomatic gastric FBs. If a child with an FB ingestion is being followed on an outpatient basis he/she should continue a regular diet and the child and their parents should be instructed to observe the stools for evidence of having passed the object and they should be advised that small blunt objects (including coins) may take as long as 4 weeks to pass spontaneously. Coins are the most common ingested FB objects among children but radiologically can be mistaken for BBs and therefore a careful history is mandatory (64,65). A child with witnessed or suspected ingestion of a coin or another blunt FB should undergo radiography as mentioned above. One should not mistake a coin for a BB therefore it is essential to closely examine the edges of the image of the coin on the x-ray to exclude the double halo sign of a possible BB. Lateral films also can be helpful to distinguish one from the other.

Localization of the ingested coin in the GI tract, age of the child, and coin size all are factors that influence the likelihood of spontaneous passage. Depending on the localization in the child's esophagus spontaneous clearance occurs in approximately 30% to 60% of children and is more likely if the coin is stuck in the distal esophagus at the time of diagnosis (66,67).

Blunt FBs and coins stuck in the esophagus should be removed urgently (within 24 hours) to avoid significant esophageal injury or erosion into the mediastinum and as detailed below BBs require emergency (<2 hours) removal from the esophagus—this also applies to other FBs if the patient is symptomatic, unable to manage secretions, or with respiratory or other concerning symptoms. A radiograph should be repeated immediately before GA to avoid an unnecessary procedure in case the FB has spontaneously passed through the esophagus but this should not delay emergency endoscopy in the case of a disc/BB (68).

Large or long objects that do not pass the pylorus and are trapped in the stomach should be removed electively or urgently in the case of a symptomatic child. There exists only expert opinion regarding the definition of "large" and "long" FBs. If the diameter of the FB is >2.5 cm it is unlikely to pass the pylorus, especially in younger children. In 1 adult study (49), 80% of FBs longer than 6 cm were unable to pass the pylorus in the 48 hours following presentation. Furthermore it is unlikely that FBs longer than 6 cm in length will pass from the first to the second part of the duodenum and are equally unlikely to pass through the ileocecal valve if the duodenum is traversed (49,68). After each extraction one should examine the mucosa to exclude significant injury.

# SHARP-POINTED OBJECTS

ESGE/ESPGHAN recommends emergent (<2 hours) removal of sharp-pointed objects located in the esophagus (all cases).

(Strong recommendation, moderate quality of evidence) ESGE/ESPGHAN recommends emergent (<2 hours) removal of sharp-pointed objects in the stomach or proximal duodenum even in asymptomatic children.

(Strong recommendation, moderate quality of evidence)

There are many reports of ingested sharp objects in children (69-71). The frequency and type of ingested sharp objects are highly dependent on cultural and environmental factors. One can see more young children with fish bone ingestions in Asian and Mediterranean families, where fish is a main food and introduced early in life (72). Symptoms of ingestions are quite common if the FB is lodged in the upper-mid esophagus (pain, dysphagia, odynophagia, drooling). A significant percentage of patients, however, remain asymptomatic for weeks and delayed intestinal perforation, extraluminal migration, abscess, peritonitis, fistula formation (68,73–75), appendicitis, liver, bladder, heart, and lung penetration (76-78) and rupture of the common carotid artery (79) have been described. The ileocecal region is the most common site for intestinal perforation but perforations have been reported in the esophagus, pylorus, at the junction between the first and second parts of the duodenum and in the colon (80). Rates of complications are higher in patients who are symptomatic, have a delay in diagnosis beyond 48 hours (81) or have swallowed a radiolucent FB (82,83). Toothpick and bone ingestions present a high risk of perforation (76,82) and are the most common FB that require surgical removal (82). Patients suspected of having swallowed sharp-pointed objects must be evaluated to define the location of the object. Many sharp-pointed objects are not visible by radiographs, so endoscopy should still follow a radiologic examination with negative findings when a high index of suspicion is present. Sharp-pointed objects lodged in the esophagus are a medical emergency due to the potentially high risk of perforation and migration. Direct laryngoscopy is an option to remove objects lodged at or above the cricopharyngeus. Otherwise, flexible endoscopy may be performed if laryngoscopy is unsuccessful and for treatment of objects lodged below this area. Sharp-pointed objects in the stomach or proximal duodenum should also be removed emergently but if these pass through the duodenum then enteroscopy, if available, or surgery, in a symptomatic patient must be considered. If observation rather than removal is chosen in the asymptomatic patient, then monitoring in a hospital setting with daily abdominal x-ray may be considered. Patients should be instructed to immediately report abdominal pain, vomiting, persistent temperature elevations, hematemesis, or melena (61,71). The average transit time for a foreign object ingested by children has

been reported as 3.6 days (70) and the mean time from ingestion of a sharp object to perforation has been reported as 10.4 days (84). If the FB has not progressed on imaging in 3 days or the patient becomes symptomatic, surgical removal may be considered (70,84).

## BATTERIES

ESGE/ESPGHAN recommends to emergently (<2 hours) remove BBs impacted in the esophagus.

(Strong recommendation, low quality of evidence)

ESGE/ESPGHAN suggests to remove BBs in the stomach emergently (<2 hours) if the child is symptomatic and/or has a known or suspected anatomical pathology in the GI tract (eg, Meckel diverticulum), and/or has simultaneously swallowed a magnet.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests that BBs >20 mm present in the stomach should be checked by radiograph and removed if still in place after > 48 hours.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN recommends an urgent endoscopic removal (<24 hours) for single cylindrical battery ingestion when impacted in the esophagus and as soon as possible elsewhere in the GI tract when the child is symptomatic (strong recommendation, moderate quality of evidence)

ESGE/ESPGHAN suggests that a single cylindrical battery in the stomach can be observed and the child monitored as an outpatient and followed by x-ray 7 to 14 days after ingestion if the battery is not passed in the stool.

(Weak recommendation, low quality of evidence)

## **Button Batteries and Cylindrical Batteries**

The US Poison Centers, from 1985 to 2009, reported an incidence of 6.3 to 15.1 cases of batteries ingested per million population (85,86).

An observed increase in poor outcome was attributed to the emergence of the larger, 20-mm diameter, lithium coin cells as an increasingly popular battery type. Thirteen deaths related to tissue damage in the esophagus or airway and 73 major outcomes (with debilitating and prolonged compromise of feeding and/or breathing that required multiple surgical procedures, enteral tube feeding and/or tracheostomies) were described (85,86). These devastating cases occurred predominantly in children who were younger than 4 years (85,86). Almost all of these major outcomes involved esophageal BB injuries and therefore impaction at this site represents the highest risk for injury and this leads to our recommendation to emergently (<2 hours) remove BB impacted in the esophagus (87). The removal of BBs in the stomach is controversial. The largest retrospective study of 8648 cases is reassuring with no reported significant gastric injuries from BB ingestion (85). There are, however, case reports of severe gastric injury (88) and also fatalities reported from aortoesophageal fistulae due to gastric BBs that had caused esophageal injury before reaching the stomach (89). The recommendation with respect to removal of BBs from the stomach is based on expert opinion and in the knowledge that only 2 cases of BB-induced gastric lesions have been reported during the last 30 years. Consistent with other guidelines (49) BBs >20 mm in the stomach should be checked by radiography and removed if still in place after >48 hours. Less evidence exists regarding cylindrical battery ingestion and although these batteries do not typically discharge electrical current in the way that BBs do they nevertheless have the potential to leak caustic fluid if the outer casing is compromised. In the largest published series identified of 62 children with cylindrical battery ingestions, approximately 82% were unaffected and no patient had major complications or death (90). Of

particular interest to the practitioner caring for adolescents, a suicide attempt was the reason for ingestion in only 1.3% of the 2382 total battery ingestions in this series, which is lower compared with other objects or poisons sought out for the same purposes (91). For single cylindrical battery ingestions we suggest urgent endoscopic removal (<24 hours) when impacted in the esophagus but if located in the stomach the patient can be monitored as an outpatient and followed by x-ray if the battery is not observed to pass in the stool. Once these batteries pass the pylorus they almost universally pass the remainder of the GI tract without incident. For the adolescent with multiple gastric cylindrical batteries as the result of intentional ingestion 1 article advocates endoscopic removal of the batteries at the time of presentation (92).

#### MAGNETS

ESGE/ESPGHAN recommends urgent (<24 hours) removal of all magnets within endoscopic reach. For those beyond endoscopic reach, close observation and surgical consultation for nonprogression through the GI tract is advised.

(Strong recommendation, moderate quality of evidence)

Ingestion of a single magnet is typically innocuous and would be expected to behave much like another blunt FB. In contrast, multiple magnets or a magnet and another metallic FB can lead to bowel wall necrosis with fistula formation, perforation, obstruction, volvulus, or peritonitis (93) by attracting themselves and trapping a portion of the bowel wall.

## FOOD BOLUS IMPACTION

ESGE/ESPGHAN recommends removal of impacted food from the esophagus as an emergency 2 hours from the time of presentation (and ideally from the time of ingestion) in case of symptoms (drooling, neck pain). If the child is asymptomatic an urgent (<24 hours) removal is indicated.

(Strong recommendation, moderate quality of evidence) ESGE/ESPGHAN suggests investigation for underlying pathology of the esophagus in all cases of food impaction.

(Weak recommendation, low quality of evidence)

Food bolus impaction in the esophagus is the most common type of impaction in adults (94). Data in children are rare but several studies show that underlying esophageal pathology, such as eosinophilic esophagitis, peptic or other strictures, achalasia and other motility disorders often are the cause of food bolus impaction (94–98). Esophageal food bolus impaction in a symptomatic patient with drooling or neck pain is an indication for emergent endoscopic removal. If the child tolerates their secretions, endoscopic removal may be postponed and an urgent (<24 hours) endoscopic removal may be considered, allowing an elective procedure and providing additional time for spontaneous clearance. The technique of removal can include piece-meal extraction, suction, and/or gentle pushing of the bolus down into the stomach, although visualization of the distal esophagus is necessary to ensure that there is no stricture distal to the bolus.

Use of glucagon to relax the lower esophageal sphincter to hasten spontaneous clearance has been studied with equivocal results and has not generally been recommended in this setting (99). Recent data suggest that it may be particularly ineffective in cases with underlying eosinophilic esophagitis (68,99,100).

## DRUG PACKETS

ESGE/ESPGHAN recommends against endoscopic removal of drug-containing packets.

(Strong recommendation, low quality of evidence)

In regions of high drug trafficking, so called "body packing" can also involve teenagers. Illegal drugs are packed into latex condoms, balloons, or plastic and swallowed for transportation (101). Leakage or rupture of these packets can be fatal; therefore, endoscopic removal should not be attempted. Surgical removal is indicated when packets fail to progress or if signs of intestinal obstruction are present. If packet rupture is suspected, surgery and urgent medical consultations for drug toxicity are indicated.

## EQUIPMENT FOR FOREIGN BODY REMOVAL

ESGE/ESPGHAN suggests that flexible endoscopy is an effective and safe procedure for removing FB from the GI tract, with a high success rate using retrieval nets, polypectomy snares, and rat-tooth forceps.

(Weak recommendation, very low quality of evidence)

## Flexible Endoscopy

Most ingested FBs are best removed using flexible endoscopes and have a high success rate. Chaves et al (102) performed a prospective mixed child and adult study showing that flexible endoscopy is an effective and safe procedure to remove FBs from the GI tract, with a high success rate using only polypectomy snares and rat-tooth forceps.

# **Rigid Endoscopy**

Some studies have shown that rigid esophagoscopy carries a higher complication rate than flexible endoscopy in performing esophageal FB extraction and its routine use is not recommended (103). Rigid endoscopy can be considered only for proximally located blunt objects, since the rigid tube provides protection (103,104).

## Magill Forceps

In a retrospective study, coins located in the proximal third of the esophagus in which successfully removed using Magill forceps. Using direct laryngoscopy under anesthesia coins were visualized and grasped with Magill forceps. The coin was removed in 96% of 165 children with a proximal esophageal coin, 82% at the first attempt (105).

## **Retrieval Devices/Overtubes**

Devices used for retrieving FBs include alligator and rattooth forceps, retrieval nets, polypectomy snares, tripod forceps, and baskets. The availability of latex cone and overtubes to protect the cardia and esophagus when removing sharp FBs is important for procedure safety. There are no studies performed in children on the use of different retrieval devices or overtubes.

## **Pharmacological Agents**

One RCT did not find an advantage in spontaneous passage of a coin from the esophagus to the stomach when using glucagon compared with placebo (106).

## **CORROSIVE INGESTION**

ESGE/ESPGHAN suggests that every child that has ingested a corrosive substance should have a thorough followup, with endoscopy dictated by symptoms and dependent on the symptoms/signs, the timing should be within 24 hours.

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(Strong Recommendation, high quality of evidence)

ESGE/ESPGHAN recommends that every child with a suspected caustic ingestion and symptoms/signs (eg, any oral lesions, vomiting, drooling, dysphagia, hematemesis, dyspnea, abdominal pain, etc) should have an EGD to identify all consequent digestive tract lesions.

#### (Strong recommendation, high quality of evidence)

ESGE/ESPGHAN suggests that in the case of suspected corrosive ingestion, EGD is withheld if the child is asymptomatic (no drooling of saliva/other symptoms and no mouth lesions) and that adequate follow-up is assured.

#### (Weak recommendation, moderate quality of evidence

Ingestion of corrosive substances can cause serious injuries to the digestive tract and occurs most commonly in children (approximately 80% of the cases) (107,108). Corrosive ingestion is mostly accidental in children (but intentional ingestion has been described in teenagers (109,110)) and although reported at any age, it is more frequent in children younger than 5 years with a maximum incidence at 2 years of age (107,109). In the developed world with the advent of child-unfriendly packaging, corrosive ingestion has become quite rare (111). Household, industrial, and farm products, especially if stored in nonoriginal containers, represent the most frequently ingested caustic agents.

The ingested substance varies between rural and urban living areas and also with the level of economic development. In developing world countries the most frequent accidental poisoning are medications (48.3%), followed by corrosive acidic substances (23.1%), carbon monoxide intoxication (12.5%), and batteries (112,113). A variety of substances have been reported that were ingested leading to caustic injuries ranging from alkaline bases with pH up to 12 (eg, sodium hypochlorite and sodium hydroxide), to acidic substances with a pH as low as 2 (eg, hydrochloric acid and salicylic acid) and also bleaching substances in which the pH is around 7 (114,115). More recently the so-called hair relaxers and liquid tabs (pods) containing detergents are a new addition to the long list of ingested products, but fortunately it seems that the upper digestive tract is not as severely damaged by these substances (116,117).

The extent of the esophageal damage is related to the nature and the concentration of the caustic substance, the duration of contact with the mucosa, and quantity ingested (118). Strong alkalis produce liquefaction necrosis with deep ulcerations and risk of esophageal stricture and/or perforation. Acids usually cause coagulative necrosis with limited tissue penetration and superficial scar formation (119). Upon swallowing, acids cause severe oropharyngeal pain and therefore they are usually consumed in small volumes compared with alkaline substances (107), resulting in a lower incidence of stricture formation and/or esophageal perforation. Other substances that may result in stricture formation are bleaching agents, nonphosphoric detergents, ammonia, and sodium bicarbonate.

Gastric lesions, with or without outlet obstruction syndrome, are almost always related to acidic ingestions, because alkalis are neutralized by gastric acid (120).

If not actually observed then corrosive ingestion may be inferred from oral burns; however their absence does not exclude ingestion and esophageal/gastric damage (121) and the consequent need for EGD. Although there are some discrepancies between studies, it is known that up to 70% of corrosive ingestions may be asymptomatic at presentation (120,122,123). It has been proposed that routine EGD is not performed in asymptomatic patients in the absence of oral lesions (119,124–126). Drooling saliva and oral lesions have been noted significantly more frequently in high-grade injury and symptoms such as pain, hypersalivation, swallowing difficulties, and bleeding are other suggestive symptoms of esophageal injury (113,121). Some children may even develop dyspnea and other respiratory symptoms (cough) and in severe cases hemodynamic instability and/or circulatory collapse; however none of these presenting symptoms is completely predictive of esophageal injury (114), although hematemesis or dyspnea as single symptoms have a high positive predictive value for esophageal lesions after caustic ingestion (122,124,126). Both in retrospective and prospective studies, the presence of 3 or more symptoms that occurred after caustic ingestion was positively associated with severe lesions seen at EGD (124,126). Young children presenting with suggestive symptoms in the absence of an observed ingestion of corrosives require EGD to exclude esophageal lesions (120,122–124,127,128).

Esophageal lesions after corrosive ingestion are described according to the Zargar Classification (129):

Grade 0 Normal.

Grade I Edema and hyperemia of mucosa.

Grade **IIa** Friability, hemorrhage, erosion blisters, exudates or whitish membranes, superficial ulcers.

Grade **IIb** Grade IIa and deep discrete or circumferential ulceration.

Grade **IIIa** Small scattered areas of necrosis, areas of brownish-black or grey discoloration.

Grade **IIIb** Extensive necrosis.

Images of the lesions are important for an accurate follow-up. Patients with low-grade lesions at endoscopy (grade 0 to IIa) who have in addition a normal physical examination and who can eat and drink normally can be discharged (126,127) but if not then admission to a hospital environment for observation should occur.

ESGE/ESPGHAN recommends to have the same grade of suspicion for both acidic and alkali ingestion regarding potential mucosal injury. (Alkali ingestion, especially lye, is associated with more severe esophageal lesions and severe gastric lesions can occur in acidic ingestion.) Stricture development has been associated with both acidic and alkali ingestion.

(Strong recommendation, high quality of evidence)

ESGE/ESPGHAN recommends high doses of intravenous dexamethasone (1 g/1.73  $m^2$  per day) administration for a short period (3 days) in IIb esophagitis after corrosive ingestion as a method of preventing esophageal stricture development. There is no evidence of benefit from the use of corticosteroids in other grades of esophagitis (I, IIa, III)

(strong recommendation, moderate quality of evidence)

Efforts should be undertaken to prevent vomiting after corrosive ingestion. Small amounts of water can be allowed if the child asks for it or even stimulated to rinse the mouth and esophagus. If the child has severe pain and if perforation is suspected, nothing should, however, be given by mouth.

Experimental studies showed decreased incidence of grade III burns and stricture formation with early corticosteroid and antibiotic use compared with controls (119,130,131). Their efficacy and safety in children with esophageal burns is, however, under discussion (132,133) because of many inconsistent variables including the type of corticosteroids used, the dosage, and duration. A recent RCT has concluded that corticosteroids are beneficial for stricture prevention in grade IIb esophageal burns (131). As yet unstudied is the possibility of the antifibrotic mitomycin-C (MMC) used topically to prevent postingestion fibrosis.

# BENIGN ESOPHAGEAL STRICTURES

ESGE/ESPGHAN recommends esophageal dilation using balloon or bougies for benign esophageal strictures only when symptoms occur.

(Strong recommendation, low-quality evidence)

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Esophageal strictures in children may have multiple etiologies including congenital or inflammatory disorders, caustic ingestion, eosinophilic esophagitis (EE), and gastroesophageal reflux disease (134). The relative proportions of etiologies vary between countries (eg, higher proportion of caustic strictures in developing countries) (135,136).

EE will not be discussed as this topic is covered by another recent guideline (14). Safety and long-term efficacy of esophageal dilation for benign esophageal strictures has been confirmed in children (137,138).

Data on the ideal timing of esophageal dilation are scarce. Two retrospective studies including 100 and 76 esophageal atresia (EA) patients compared routine esophageal dilation every 3 weeks starting 3 weeks postsurgery versus when symptoms developed. No difference in outcome and complications were found between the 2 groups after 2 and 3 years of follow-up but significantly fewer dilations were needed in the on-demand dilation group (139,140).

Through-the-scope balloons and wire-guided polyvinyl bougie dilators (Savary Gilliard) are most frequently used to dilate benign esophageal strictures in children and similar results are reported—a retrospective study (141) compared 125 balloon dilations versus 88 bougie dilations in children with benign esophageal strictures and reported 4 failures in the bougie group related to unsuccessful passage of the stricture.

Expert opinion suggests that both anesthesiology and surgical assistance should be available during esophageal dilation procedures in children—the latter in case of complications (141).

## **Balloon Dilation**

Balloon dilation can be performed under direct endoscopic view or fluoroscopic view. The size of the balloon catheter can vary from 4 to 22 mm and the balloon inflation duration varies between studies from 20 to 120 seconds (142). A study (137) on 77 children (median age 1.8 years) who underwent 260 balloon dilations of benign esophageal strictures under endoscopic view (mean 3.4 dilations/patient), reported 4 (1.5%) esophageal perforations, with 1 requiring surgery—the remaining patients were all asymptomatic after a median follow-up of 6.6 years. Strictures shorter than 5 cm in length appeared to have a significantly better outcome (143,144). In a retrospective study of 34 patients with EA and symptomatic esophageal strictures, balloon dilation appeared to be more effective and less traumatic than bougienage (145).

## Bougies

Bougie dilation is a safe and effective dilation technique for esophageal strictures. In a study of 107 children, dilation was performed at 2 to 3 week intervals using Savary-Gilliard bougies and was considered adequate if the esophageal lumen could be dilated to 15 mm diameter (12 mm in children <5 years of age) with complete relief of symptoms (146). Subsequently, dilation was performed on an "as needed" basis dependent on symptoms. Dilation was successful in all but 3 cases. Dilation was also successful in patients with strictures 5 cm or more in length and/or in patients with multiple corrosive strictures, although these required a higher number of subsequent sessions for recurrence. Six esophageal perforations occurred during 648 dilation sessions (0.9%) with 1 requiring surgical repair.

## Size, Number, and Interval Between Dilations

There are no data on the "optimal" increase in size that should be aimed for at each dilation session. The "rule of three" is often invoked: no more than 3 times the diameter of the stricture so a 3-mm stricture should not be dilated to >9 mm and so on. Balloons are to be preferred over bougies if financially possible. There exists no consensus in regard to the interval between dilations and the frequency of this intervention is often individualized according to relief of dysphagia and the severity of the stricture observed during repeat endoscopy. Most studies used a minimal period of 3 weeks between dilation sessions (144,146,147) and for balloon dilation an average of 3 dilations appeared to be required (142).

ESGE/ESPGHAN suggests the following definition of a benign refractory or recurrent stricture in children: "An anatomic restriction because of cicatricial luminal compromise or fibrosis that results in dysphagia in the absence of endoscopic evidence of inflammation. This may occur as the result of either an inability to successfully remediate the anatomic problem to obtain age-appropriate feeding possibilities after a maximum of 5 dilation sessions (refractory) with maximal 4-week intervals, or as a result of an inability to maintain a satisfactory luminal diameter for 4 weeks once the age-appropriate feeding diameter has been achieved (recurrent)."

#### (Weak recommendation, very low level of evidence)

To define refractory and recurrent strictures in children, we adopted the definition used in adults based on Kochman criteria (148). In an online survey of the working group members, the number of sessions, intervals, and target diameter were assessed. Seventeen of 18 members (94%) supported the definition stated above, with 39% and 61% of respondents mentioning a maximum of 3 and 5 sessions, respectively. Proposed intervals between sessions were 2 or 3 weeks (39% of members each) or 4 weeks (22%). Refractory and recurrent stenosis is reported in approximately 30% of the cases (142).

ESGE/ESPGHAN suggests temporary stent placement or application of topical MMC following dilation for refractory esophageal stenosis in children. ESGE/ESPGHAN does not suggest the routine use of intralesional steroids for refractory esophageal stenosis in children.

## (Weak recommendation, low quality of evidence).

There is no standard treatment for refractory stenosis.

Local application of MMC is a therapeutic option for the treatment of refractory esophageal strictures in children (149,150). A systematic review identified 11 publications including 31 cases (151) of various etiologies. In 1 study, cotton pledgets soaked in a solution (0.1 mg/mL) of MMC were applied endoscopically directly onto the mucosa postdilation with a frequency of 1 to 12 times during 12 weeks. After a mean follow-up of 22 (6–60) months complete relief of symptoms was reported for 21 of 31 children (67.7%), and 6 of 31 (19.4%) had a partial relief. In 4 children (12.9%) MMC application failed. No direct or indirect adverse effects were reported. Two double blind RCTs showed that MMC application significantly reduced the number of esophageal dilations sessions needed to alleviate dysphagia following EA and corrosive strictures (152,153).

MMC is a cytostatic agent; therefore, dysplasia of healthy tissues after application should be considered as a theoretical risk (151). This complication was not observed in a study performing esophageal biopsies during a 24-month mean follow-up (149). Future studies with long-term follow-up are required to evaluate the potential adverse effects (149,151).

With the advent of removable, fully covered, self-expandable metal stents (FCSEMS), the use of esophageal stents in children has expanded in particular for the treatment of refractory stenoses. In 3 studies that included a total of 25 children, complete clinical response following stent removal with no recurrence of dysphagia or need for subsequent dilations was reported in 50% to 85% of

patients (154). Most patients experienced nausea or chest pain in the days following stent placement, in some cases leading to premature removal of the stent. Duration of stenting varied from 1 to 24 weeks. Stent migration was the most commonly cited complication and occurred in 0% to 29% of patients. In a recent retrospective study in children with perforations and refractory strictures after EA repair a total of 41 plastic and FCSEMS were placed in 24 patients, including 14 patients who had developed esophageal leaks. Success in the treatment of refractory strictures was limited due to a high stricture recurrence rate after stent removal (39% at 30 days and 26% at 90 days). Stent-related adverse events included migration (21% of plastic and 7% of FCSEMS), granulation tissue (37% of FCSEMS and none of plastic), and deep ulceration (22% of FCSEMS and none of plastic) (155).

A recent uncontrolled study in 10 children with intractable esophageal strictures due to caustic ingestion reported symptom resolution using stricture dilation preceded by intralesional triamcinolone injection (156). In a recent double-blind RCT in adults with benign anastomotic strictures, no benefit of intralesional triamcinolone could, however, be demonstrated (157).

In patients operated for EA, ESGE/ESPGHAN suggests long-term endoscopic surveillance for Barrett esophagus and cancer. Frequency would be dictated by the presence or not of dysplasia and should follow standard guidelines already published in the literature.

#### (Weak recommendation, low quality of evidence).

A systematic review and meta-analysis to define the prevalence of chronic long-term problems in 907 EA operated patients showed, compared with controls, a 40.3 relative risk for dysphagia during adolescence and adulthood due to altered peristalsis (158). Gastroesophageal reflux is a known risk factor for subsequent development of esophageal intestinal metaplasia. The overall estimated prevalence of Barrett esophagus was 6.4% in patients with EA, which is 4 times and 26 times higher than its prevalence in adults and pediatric general population, respectively. In a systematic review the prevalence of esophageal carcinoma was low (1.4%) and only squamous cell carcinoma was described. Cases of adenocarcinoma in EA patients have, however, been reported (159). In view of the high incidence of Barrett esophagus in patients with EA at a young age (160), endoscopic surveillance is warranted in adolescence and adulthood (161). The question of whether endoscopic surveillance should occur for metaplastic change following corrosive ingestion is not one that can be adequately answered with the present medical evidence.

# UPPER AND LOWER GASTROINTESTINAL BLEEDING

ESGE/ESPGHAN suggests that having employed all necessary medical interventions as standard, EGD be performed early (<12 hours) in acute upper GI bleeding (AUGIB) cases which require ongoing circulatory support or where a large hematemesis or melena occurs.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN recommends that having employed all necessary medical interventions as standard, EGD be performed early (<12 hours) in AUGIB in cases with known esophageal varices.

(Strong recommendation, moderate quality of evidence)

ESGE/ESPGHAN suggests that having employed all necessary medical interventions as standard, EGD be performed within 24 hours in AUGIB cases which require transfusion due to hemoglobin drop <8 g/dL, where an acute drop of 2 g/dL is identified, and in those who are stable but whose bleeding score is above a recognized threshold/validated score for probable endoscopic intervention requirement.

(Weak recommendation, moderate quality of evidence)

ESGE/ESPGHAN suggests that EGD be performed before hospital discharge in children with AUGIB and pre-existing liver disease or portal hypertension.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN does not suggest routine use of wireless capsule endoscopy/enteroscopy in AUGIB in children.

(Weak recommendation, moderate quality of evidence) ESGE/ESPGHAN suggests that urgent therapeutic IC is not usually necessary in lower GI bleeding unless severe enough to cause circulatory compromise but diagnostic IC is needed as soon as is practical and safe.

## (Weak recommendation, weak quality of evidence)

Adult studies are the primary guides for evaluation of pediatric practice, but are not entirely applicable to children. Distinction is drawn between the speed of intervention required for AUGIB and acute lower GI bleeding in children. It is rare to require intervention for lower GI bleeding as the majority of massive hemorrhage (fresh red blood/melena) originates in the upper GI tract with the occasional exception of a Meckel diverticulum or severe colitis.

Scoring systems of intervention in children are emerging but require prospective evaluation of predictive accuracy and reliability.

In adults with AUGIB validated scoring systems have been published (162–164) while, to date, only 1 such scoring system exists in pediatrics ("Sheffield Scoring System"), which may predict the requirement or otherwise for endoscopic hemostatic therapy, is undergoing prospective multicenter validation at present (165). This retrospective case-control study reliably predicted which children most likely require endoscopic intervention. Particular weighting was given on modeling to such factors as: transfusion requirement; signs of hypovolemia (raised HR>20 bpm above age-appropriate median and prolonged capillary refill time); large hematemesis; melena; and drop of Hb >2 g/dL. The timing of such intervention is clearly dependent on the circulatory stability of the child. For uncontrolled bleeding requiring volume support immediate intervention is suggested. For children in whom the threshold score is reached but who are stable then endoscopic intervention within 12 hours is suggested. Finally, for children whose clinical bleeding risk score does not reach the intervention threshold and in whom AUGIB would appear to have ceased then elective or no endoscopy is suggested (165).

The matter is further complicated by the wide variability of the following important practical factors in the provision of such life-saving techniques for children: availability of appropriately trained pediatric therapeutic endoscopists; availability of units with adequate and appropriate equipment/skills within geographical proximity; and agreed guidelines/algorithms of care for this clinical emergency with, to date, no universal view of when and how to intervene endoscopically.

This is further compounded by an absence of knowledge of the size of the clinical problem in pediatrics. Many pediatric endoscopists would not encounter an AUGIB case more than a handful of times each year. A case, then, may be made for centralization of such units and skills, but the caveat to this is the need then for safe transport of a child who may be actively bleeding to such a center.

There seems to be little requirement for urgent or early use of the wireless capsule endoscope or enteroscopy in acute bleeding in children (40,166,167).

Endoscopy has been advocated for the management of AUGIB, but the optimal timing is still uncertain. Ideally, endoscopy should occur after the stabilization of the patient and various studies have been conducted comparing various timing of endoscopy performed within 6, 12, and 24 hours of presentation (168). Adult literature recommends that endoscopy in AUGIB should be performed within 24 hours of presentation (169,170) or within 12 hours when bleeding continues at a rate considered potentially life threatening (170–174). A clinical benefit of endoscopy performed of presentation is reported in acute variceal bleeding in children (175). In the pediatric population endoscopy in AUGIB is recommended within 24 hours (165,170,172).

# Lower Gastrointestinal Bleeding

Most cases of acute colonic bleeding (or lower GI bleeding) in children presenting either as hematochezia (bright red blood, clots) or melena will stop spontaneously, and thus not needing urgent evaluation (176), but IC following adequate bowel preparation need to be planned before discharge from the hospital. For children with severe hematochezia, defined as continued bleeding within the first 24 hours of hospitalization with a drop in the hemoglobin of at least 2 g/dL and/or a transfusion requirement, urgent diagnosis and intervention are required to control bleeding (165,177). When hematochezia is not severe, elective IC need to be scheduled.

# ENDOSCOPIC HEMOSTASIS TECHNIQUE FOR GASTROINTESTINAL BLEEDING IN CHILDREN

ESGE/ESPGHAN recommends hemostasis of esophageal variceal bleeding in children using band ligation, if feasible, or sclerotherapy as an alternative.

(Strong recommendation, moderate quality of evidence)

ESGE/ESPGHAN suggests that the treatment of peptic ulcers and Dieulafoy lesion should not be carried out with epinephrine injection alone but in combination with thermal or mechanical techniques.

(Weak recommendation, low quality of evidence)

Minimal data exist comparing endoscopic equipment and techniques in children. Adult studies are the primary guides for evaluation of pediatric equipment. Working channel size is the major factor limiting the choice of accessories (18). In children weighing >10 kg, endoscopes for therapeutic endoscopy are generally identical to those used in adults (Table 4).

Standard pediatric gastroscopes have a 4.9- to 6.0-mm outer diameter and a 2.0-mm working channel. They will accommodate needles for injection therapy (4–6 mm length), bipolar, and argon plasma coagulation probes but not heater or multipolar probes, or ligating or mechanical devices (178). Removing the Teflon sheath from a hemostatic clip allows use with pediatric endoscopes. Patient electrodes and grounding pads are available in neonatal (<3 kg) and pediatric (<15 kg) sizes.

# Nonvariceal Bleeding

## Dieulafoy Lesion

In a review of 24 pediatric cases, half were treated surgically, the others were managed endoscopically by injection therapy, band ligation, and thermocoagulation (179). Epinephrine combined with either mechanical treatment or heater probe is preferable to epinephrine alone for hemostasis (180). One should consider tattooing the bleeding site to aid location in the event of rebleeding (181).

## **Bleeding Ulcer**

A report describes the successful treatment of a newborn by heater probe thermocoagulation (182). Argon plasma coagulation with a 1.5 or 2.3 mm probe was used in 12 children (183). Generally, for older children standard adult GI practice should apply to AUGIB (184).

## Variceal Bleeding

A randomized prospective study in 49 children showed that band ligation is safe and effective, superior to sclerotherapy in terms of variceal eradication and was associated with a lower rebleeding rate (185). Most studies support band ligation but if that is impossible due to patient size then sclerotherapy can be used (185-187). The use of band ligation sets for gastroscopes with a diameter of 8.5 to 9.2 mm is limited primarily by the narrowness of the pharynx, and not only by body weight. Recently band ligation sets are available for pediatric gastroscopes and although previously sclerotherapy was the method of choice for children weighing < 8 kg(188), this may change soon. Evidence is limited concerning the management of gastric varices in children. In case reports, Nbuthyl-2 cyanoacrylate "glue" injection has been successful (186,189). Small cohort studies in children using variceal banding as prophylaxis exist (190). It has been found that variceal grading can be a subjective assessment. There is no evidence that PPI use postbanding is beneficial (191). A retrospective review of the safety and efficacy of expanded polytetrafluoroethylene-covered transjugular portosystemic shunt in 12 children showed a satisfactory result and therefore this may be a useful alternative in acute or recurrent medically or endoscopically uncontrollable variceal bleeding (192).

ESGE/ESPGHAN suggests adopting GA in children undergoing endoscopy for GI bleeding. GA is recommended when there is variceal bleeding. Deep sedation may be used in less severe bleeding in older children.

(Weak recommendation, low quality of evidence)

ESGE/ESPGHAN suggests using video capsule endoscopy (VCE) in children when there is suspected small intestinal bleeding and in addition balloon enteroscopy for therapeutic purposes.

(Weak recommendation, moderate quality of evidence)

The majority of studies are retrospective analyses focusing on the diagnostic yield and therapeutic success of endoscopy in children with GI bleeding. The type of sedation/anesthesia used when performing upper GI endoscopy in children for AUGIB is not always reported, but most of the procedures were performed under GA with endotracheal intubation, whereas conscious or deep sedation should *not* be preferred (193).

VCE in children finds a main indication in the study of obscure GI bleeding and suspected or known Crohn disease. The youngest VCE patient investigated was 8 months (194). If the capsule cannot be swallowed it is placed endoscopically using various devices (166). In a meta-analysis including 723 VCE examinations in children, the diagnostic yield of VCE was 65.4% with retention rates comparable to those of adults (195). Interventional studies on small bowel endoscopy mostly reported on double-balloon enterosocopy using an endoscope with either 9.4 or 8.3 mm diameter (39,40,196).

Angiodysplastic lesions, polyps, Meckel diveritculi, chronic mucosal erosive/inflammatory diseases such as diaphragm disease, and congenital lesions such as duplication cysts are all noted in the literature as causes of bleeding either acutely or in a more chronic fashion. Diagnostic approaches include VCE, DBE, CT scan, CT angiogram, intravascular angiography, and isotope-labeled bleeding scans.

# ENDOSCOPIC RETROGRADE CHOLANGIOPANCREATOGRAPHY

ESGE/ESPGHAN suggests ERCP in pediatric patients (>1 year old) for therapeutic purposes following diagnostic information from noninvasive diagnostic modalities such as magnetic resonance cholangiopancreatography (MRCP). Diagnostic ERCP can be considered in selected cases in which advanced noninvasive imaging is inconclusive.

(Weak recommendation, low quality of evidence).

ESGE/ESPGHAN recommends that therapeutic ERCP in pediatric patients (>1 year old) is considered for diseases listed in Table 6 following diagnostic information from noninvasive modalities such as MRCP. Results and complication rates of ERCP in children are similar to those reported in adults.

(Weak recommendation, low quality of evidence).

Published series on pediatric ERCP are retrospective data collections. Indications for ERCP in children are more frequently related to congenital abnormalities or trauma than to malignancy, which is more frequent in adults (197). The division between biliary and pancreatic indications is age dependent: pancreatic and biliary indications prevail in children of 7 to 12 years and 13 to 17 years, respectively, whereas they are similarly distributed in children younger than 6 years (198). Indications for ERCP in pediatric patients are summarized in Table 6.

Currently, the majority of ERCPs in children are therapeutic as MRCP has mostly replaced diagnostic ERCP but this is not reflected in the literature. (199). MRCP accurately depicts pancreaticobiliary anatomy and related diseases in children and secretin stimulation can enhance the visualization of nondilated pancreatic ducts thereby improving diagnostic sensitivity (200). Two large retrospective series in children with a mean age of 10 years (199,200) reported an 11% to 13% nondiagnostic rate of MRCP and in such cases diagnostic ERCP can be considered.

Technical success for diagnostic and therapeutic ERCP in children (>1 year old) is high, with adverse event rates similar to those in adults (201,202). The success rate reported in pediatric ERCP case series is >90%, with a complication rate of 2.3% to 9.7%, and no procedure-related mortality (197,198,201–213). A retrospective case-controlled study (201) compared results of ERCP in children and adults performed at the same center. Pediatric and adult patients were matched according to indications, diagnostic findings, and technical complexity. ERCP success rate was 97.5% in children compared with 98% in the adult cohort and complications rates were similar (3.4% in adults vs 2.5% in pediatric patients). The risk of post-ERCP pancreatitis (PEP) is increased in therapeutic compared with diagnostic ERCP, in the case of

pancreatic duct injection and in more (vs less) complex procedures in children (202,211,214).

Chronic pancreatitis (CP) is a frequent indication for therapeutic ERCP in children. ESGE recommend endotherapy as a firstline therapy for CP in children starting at 8 years in the same conditions as in adults (215). Two recently published large series (216,217) confirmed good results of endoscopic treatment of CP in children with complete pain resolution in 63.6% of the cases and improvement in 21.6% (216), whereas pancreatic duct stenting significantly decreased the number of pancreatitis after a mean follow-up of 4.5 years (217).

ESGE/ESPGHAN suggests that diagnostic ERCP in neonates and infants ( $\leq 1$  year old) with cholestatic hepatobiliary disease is considered if noninvasive investigations are not conclusive to allow timely referral to surgery for suspected biliary atresia (BA) or to avoid unnecessary surgery if BA is excluded. (Weak recommendation, low quality of evidence).

The first-line imaging modalities in neonatal cholestasis are abdominal ultrasound (triangular cord sign) and cholescintigraphy; ERCP and MRCP are not routinely recommended for the diagnosis of cholestatic jaundice in infants (218).

In the setting of neonatal anatomy, and in particular, the minute structures of biliary hypoplasia or BA, MRCP still appears to have unsatisfactory diagnostic accuracy (70% in a recent series on 190 infants) (219). Because the diagnosis of BA at MRCP is based on the absence of visualization of the extrahepatic bile ducts and a prospective evaluation (220) of normal infants by MRCP visualized extrahepatic bile ducts in 62.5% of the cases, the authors concluded that MRCP led to a high level of false positivity in the setting of neonatal cholestasis.

In this particular indication, retrospective series (221-227) report that ERCP has a >85% success rate but also report a complication rate of up to 10%, although this included cases of increased pancreatic enzymes. All complications resolved by conservative treatment. Procedure-related mortality was not reported. Keil et al (221), in a series of 104 infants, reported 86% sensitivity and 94% specificity of ERCP for BA and 100% sensitivity and 90% specificity for choledochal cyst. According to published data (221,222,224-227) ERCP avoided unnecessary laparotomies in 18% to 42% of the infants. Liver biopsy is indicated as a complimentary investigation especially if ERCP is inconclusive (218).

ERCP currently offers superior diagnostic visualization of the biliary tree in infants and neonates (221). ERCP, however, remains an invasive procedure; thus, its indication in infants needs to be carefully evaluated in a multidisciplinary setting, balancing risks and benefits.

TABLE 6. Typical endoscopic retrograde cholangiopancreatography indications in pediatric patients						
B1I	lary	Pancreatic				
Diagnostic	Therapeutic	Diagnostic	Therapeutic			
Cholestasis in neonates and infants	Common bile duct stones	Evaluation of anomalous pancreaticobiliary junction	Chronic pancreatitis			
Choledochal cyst	Bile leak (postsurgical/post- traumatic)		Recurrent acute pancreatitis			
Primary sclerosing cholangitis (brush cytology)	Benign biliary strictures		Pancreas divisum			
	Primary sclerosing cholangitis		Pancreatic duct leak (postsurgical/ post-traumatic)			
	Malignant biliary strictures		Pancreatic pseudocyst			
	Parasitosis (ascariasis, Fasciola)		Injection of botulinum toxin for sphincter of Oddi dysfunction			

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#### ESGE/ESPGHAN recommends that ERCP in children is performed by an experienced endoscopist, in a high-volume tertiary care center and with pediatric involvement.

#### (Strong recommendation, moderate quality of evidence).

The annual number of pediatric ERCPs performed even in a referral endoscopy unit is usually low. The 2 largest published series (206,210) report 24 and 36 pediatric ERCPs/year; in neonates and infants this figure is lower with a minimum of 2.7 ERCP/year and a maximum of 20 (224). Most pediatric endoscopy training programs offer limited exposure of their trainees to ERCP. Training in ERCP requires performance numbers that often exceed the number of patients an average pediatric gastroenterologist will encounter in their training. Pediatric gastroenterologists undoubtedly perform a lower volume of ERCP compared with adult-trained endoscopists at expert centers and it could be argued that initial numbers for competency should be the same (228,229). North American Society for Pediatric Gastroenterology, Hepatology and Nutrition (230) suggests a minimum of 200 diagnostic and therapeutic ERCPs to achieve competence in pediatric patients, although this is a suggestion rather than based on evaluation of prospective competence. The combination of a pediatric gastroenterologist, who is knowledgeable about the pediatric GI disease, with an experienced ERCP endoscopist is perhaps an ideal alternative.

#### ESGE/ESPGHAN suggests GA for ERCP in children. Deep/conscious sedation can be considered for teenagers (age 12–17 years) although GA is the preferred choice.

#### (Weak recommendation, low quality of evidence).

Given the long duration and degree of difficulty of ERCP in small children and neonates, and the softness of their tracheal wall, it is recommended to perform ERCP in children under GA with endotracheal intubation. Some series report the use of conscious/ deep sedation in 70% of cases but this should be considered to be historical and age dependent and should not be considered for children younger than 12 years (207,208,211).

#### Prophylaxis of PEP with Nonsteroidal Anti-inflammatory Drug(s) (diclofenac/indomethacin suppository) is recommended in children older than 14 years.

## (Strong recommendation, high quality of evidence).

No RCTs on the prophylaxis of PEP in children have been published. In a series of 423 ERCPs, prophylactic pancreatic stenting was associated with higher rates of PEP in high-risk patients and did not eliminate severe PEP (214). Pharmacologic prophylaxis with diclofenac/indomethacin suppositories are recommended in adults (231) and may be used in children, although evidence is lacking to date.

Protection of radiosensitive organs (thyroid gland, breasts, gonads, and eyes) is recommended together with adjustment of collimation to the smaller size of children.

# (Strong recommendation, high quality of evidence).

Children are more sensitive than adults to radiation exposure and the life-time risk of cancer induction is possibly 3 to 5 times higher. ESGE guidelines on radiation exposure (232) recommends to adjust collimation to the smaller size of the patient and to protect with radiation protection shields the most radiosensitive organs (thyroid gland, breasts, gonads, and eyes) and to keep these organs out with the main x-ray beam.

#### ESGE/ESPGHAN recommends the pediatric 7.5 mm duodenoscope for children weighing <10 kg and that a therapeutic duodenoscope can be used in those weighing $\geq$ 10 kg.

# (Strong recommendation, low quality of evidence).

ERCP in infants and neonates ( $\leq 1$  year old) is feasible with a 7.5 mm pediatric duodenoscope. This endoscope has a 2 mm working channel limiting the array of devices that can be used; however, double-lumen sphincterotomes, extraction baskets, and retrieval balloons are commercially available. Previously specific

pediatric ERCP scopes with a standard 3.2 mm working channel or a therapeutic 4.2 mm working channels were available but are no longer commercially available. Commercially available therapeutic duodenoscopes have an insertion tube diameter of 11.3 to 11.6 mm and a distal end of 13 to 13.5 mm (Table 4).

## ENDOSCOPIC ULTRASONOGRAPHY

The endobronchial ultrasound (EBUS) can be adapted for EUS in children with a weight <15 kg. A standard linear echoendoscope should only be employed in children under GA, considering the stiff and potentially traumatic distal part.

(Weak recommendation, low quality of evidence).

ESGE-ESPGHAN suggests the use of EUS in children only in tertiary referral centers with experience in therapeutic endoscopy. Strict collaboration between adult and pediatric gastroenterologists is required in the case of EUS with standard echoendoscopes.

#### (Weak recommendation, low quality of evidence).

Experience of EUS in pediatric patients is limited partly because commercially available echoendoscopes have a distal end diameter of 11 to 14 mm for radial probes and 14 mm for linear probes which cannot traverse from D1 to D2 small children. GA and careful insertion of the rigid tip of the linear echoendoscope is needed. The use of adult echoendoscopes was recently described (233) in children ages >3 years with weight  $\geq$ 15 kg. In smaller children the EBUS endoscope can be considered (Table 4).

Reported experiences with standard EUS scopes and miniprobes in children are limited to small series (233,234-251), with only 2 articles including >50 cases (233,243). Many articles are from adult endoscopy centers, which routinely perform EUS with standard echoendoscopes.

Miniprobes can be used with standard endoscopes in small children (234–240) and allow EUS in special anatomic situations such as stenoses through which standard EUS scopes may not feasibly be passed.

ESGE-ESPGHAN suggests the use of radial EUS with miniprobes to diagnose congenital esophageal strictures (tracheobronchial remnants vs fibromuscular stenosis subtypes).

(Weak recommendation, very low quality of evidence).

Congenital esophageal stenosis (CES) is an esophageal malformation with a stenosis generally located in the middle or more often in the lower esophagus. Three CES subtypes have been described: fibromuscular, tracheal cartilaginous remnants, and the membranous web. In approximately 10% CES is associated with EA (247) and differentiation between the CES subtypes is possible by histopathology after surgical resection (234,235,239,240,247).

A systematic review (247) from 144 CES cases confirmed the importance of EUS as the main diagnostic tool to distinguish CES subtypes and modify patient management. In tracheal cartilaginous remnant CES, some authors suggest stenosis resection and anastomosis to avoid the risk of post-dilation esophageal perforation (235,247). A recent series reported CES dilations effective in all the different CES subtypes in 96% of the cases and suggests that surgery is reserved for cases of endoscopic dilation failure (252).

ESGE/ESPGHAN suggests consideration of EUS for the diagnosis of pancreaticobiliary diseases in children in whom noninvasive imaging modalities (ultrasonography, MRCP) are inconclusive (Table 7).

(Weak recommendation, very low quality of evidence).

ESGE/ESPGHAN suggests EUS-guided drainage of pancreatic pseudocysts in children should be performed in large EUS centers with specific experience and expertise.

(Weak recommendation, low quality of evidence).

TABLE 7. Typical endoscopic ultrasonography indications in pediatric patients					
Esophagus	Stomach	Duodenum	Pancreaticobiliary		
Congenital esophageal stenosis Eosinophilic esophagitis Esophageal duplication	Gastric duplication Gastric varices	Duodenal duplication	Bile duct stone Pancreatic pseudocyst (diagnosis and treatment) Pancreatic disease (±FNA)		

FNA = fine-needle aspiration.

EUS and EUS-guided fine-needle aspiration have been reported as feasible in small series of children for assessing pancreaticobiliary diseases (233,241–246) in which noninvasive imaging modalities (eg, MRCP) are inconclusive. Therapeutic EUS with the drainage of pancreatic pseudocysts can be performed with the same technique as that described in adult patients, although in large cysts EUS guidance may not be necessary (233,250,251).

Indications for EUS in children are summarized in Table 7.

ESGE and ESPGHAN guidelines represent a consensus of best practice based on the available evidence at the time of preparation. They may not apply in all situations and should be interpreted in the light of specific clinical situations and resource availability. Further controlled clinical studies may be needed to clarify aspects of these statements, and revision may be necessary as new data appear. Clinical considerations may justify a course of action at variance to these recommendations. ESGE and ESPGHAN guidelines are intended to be an educational device to provide information that may assist endoscopists in providing care to patients. They are not rules and should not be construed as establishing a legal standard of care or as encouraging, advocating, requiring, or discouraging any particular treatment.

#### REFERENCES

- Koletzko S, Jones NL, Goodman KJ, et al. Evidence-based guidelines from ESPGHAN and NASPGHAN for *Helicobacter pylori* infection in children. J Pediatr Gastroenterol Nutr 2011;53:230–43.
- 2. Turner D, Levine A, Escher JC, et al. Management of pediatric ulcerative colitis: joint ECCO and ESPGHAN evidence-based consensus guidelines. *J Pediatr Gastroenterol Nutr* 2012;55: 340–61.
- Guyatt GH, Oxman AD, Vist GE, et al. GRADE: an emerging consensus on rating quality of evidence and strength of recommendations. *BMJ* 2008;336:924–6.
- Heuschkel RB, Gottrand F, Devarajan K, et al. ESPGHAN position paper on management of percutaneous endoscopic gastrostomy in children and adolescents. *J Pediatr Gastroenterol Nutr* 2015;60: 131–41.
- Dumonceau JM, Hassan C, Riphaus A, et al. European Society of Gastrointestinal Endoscopy (ESGE) Guideline Development Policy. *Endoscopy* 2012;44:626–9.
- ASGE Standards of Practice CommitteeLightdale JR, Acosta R, Shergill AK, et al. Modifications in endoscopic practice for pediatric patients. *Gastrointest Endosc* 2014;79:699–710.
- Samer Ammar M, Pfefferkorn MD, Croffie JM, et al. Complications after outpatient upper GI endoscopy in children: 30-day follow-up. *Am J Gasterol* 2003;98:1508–11.
- Lee WS, Zainuddin H, Boey CC, et al. Appropriateness, endoscopic findings and contributive yield of pediatric gastrointestinal endoscopy. *World J Gastroenterol* 2013;19:9077–83.
- Hummel TZ, ten Kate FJ, Reitsma JB, et al. Additional value of upper GI tract endoscopy in the diagnostic assessment of childhood IBD. J Pediatr Gastroenterol Nutr 2012;54:753–7.
- Kori M, Gladish V, Ziv-Sokolovskaya N, et al. The significance of routine duodenal biopsies in pediatric patients undergoing upper intestinal endoscopy. *J Clin Gastroenterol* 2003;37:39–41.

- 11. Volonaki E, Sebire NJ, Borrelli O, et al. Gastrointestinal endoscopy and mucosal biopsy in the first year of life: indications and outcome. *J Pediatr Gastroenterol Nutr* 2012;55:62–5.
- Mangiavillano B, Masci E, Parma B, et al. Bulb biopsies for the diagnosis of celiac disease in pediatric patients. *Gastrointest Endosc* 2010;72:564–8.
- Husby S, Koletzko S, Korponay-Szabo IR, et al. European Society for Pediatric Gastroenterology, Hepatology, and Nutrition guidelines for the diagnosis of coeliac disease. *J Pediatr Gastroenterol Nutr* 2012;54:136–60.
- Papadopoulou A, Koletzko S, Heuschkel R, et al. Management guidelines of eosinophilic esophagitis in childhood. J Pediatr Gastroenterol Nutr 2014;58:107–18.
- Levine A, Koletzko S, Turner D, et al. ESPGHAN revised porto criteria for the diagnosis of inflammatory bowel disease in children and adolescents. *Pediatr Gastroenterol Nutr* 2014;58:795–806.
- Van Beek EJ, Leroy PL. Safe and effective procedural sedation for gastrointestinal endoscopy in children. J Pediatr Gastroenterol Nutr 2012;54:171–85.
- Hayat JO, Sirohi R, Gorard DA. Paediatric endoscopy performed by adult-service gastroenterologists. *Eur J Gastroenterol Hepatol* 2008;20:648–52.
- Committee AT, Barth BA, Banerjee S, et al. Equipment for pediatric endoscopy. *Gastrointest Endosc* 2012;76:8–17.
- Beggs AD, Latchford AR, Vasen HF, et al. Peutz-Jeghers syndrome: a systematic review and recommendations for management. *Gut* 2010;59:975–86.
- Bülow S, Leeds Castle Polyposis Group. Guidelines for the surveillance and management of familial adenomatous polyposis (FAP): a world wide survey among 41 registries. *Colorectal Dis* 1999;1:214–21.
- Dabadie A, Bellaiche M, Cardey J, et al. Current indications of ileocolonoscopy in children in 2012. Arch Pediatr 2012;19:1247–51.
- Thakkar K, El-Serag HB, Mattek N, et al. Complications of pediatric colonoscopy: a five-year multicenter experience. *Clin Gastroenterol Hepatol* 2008;6:515–20.
- Hassan C, Bretthauer M, Kaminski MF, et al. Bowel preparation for colonoscopy: European Society of Gastrointestinal Endoscopy (ESGE) guideline. *Endoscopy* 2013;45:142–50.
- Turner D, Levine A, Weiss B, et al. Evidence-based recommendations for bowel cleansing before colonoscopy in children: a report from a national working group. *Endoscopy* 2010;42:1063–70.
- Hunter A, Mamula P. Bowel preparation for pediatric colonoscopy procedures. J Pediatr Gastroenterol Nutr 2010;51:254–61.
- Di Nardo G, Aloi M, Cucchiara S, et al. Bowel preparations for colonoscopy: an RCT. *Pediatrics* 2014;134:249–56.
- Manfredi MA, Jiang H, Borges LF, et al. Good agreement between endoscopic findings and biopsy reports supports limited tissue sampling during pediatric colonoscopy. *J Pediatr Gastroenterol Nutr* 2014;58:773–8.
- Liu X, Xiao SY, Plesec TP, et al. Collagenous colitis in children and adolescents: study of 7 cases and literature review. *Mod Pathol* 2013;26:881–7.
- Singh P, Das P, Jain AK, et al. Microscopic colitis in children with chronic diarrhea. J Pediatr Gastroenterol Nutr 2013;57:240–4.
- Castellaneta SP, Afzal NA, Greenberg M, et al. Diagnostic role of upper gastrointestinal endoscopy in pediatric inflammatory bowel disease. J Pediatr Gastroenterol Nutr 2004;39:257–61.

- Mowat C, Cole A, Windsor A, et al. Guidelines for the management of inflammatory bowel disease in adults. *Gut* 2011;60:571–607.
- Geboes K, Van Eyken P. Inflammatory bowel disease unclassified and indeterminate colitis: the role of the pathologist. J Clin Pathol 2009;62:201–5.
- Sultan M, Ramprasad J, Jensen MK, et al. Endoscopic diagnosis of pediatric acute gastrointestinal graft-versus-host disease. J Pediatr Gastroenterol Nutr 2012;55:417–20.
- Kreisel W, Dahlberg M, Bertz H, et al. Endoscopic diagnosis of acute intestinal GVHD following allogeneic hematopoietic SCT: a retrospective analysis in 175 patients. *Bone Marrow Transplant* 2012;47:430–8.
- Wang WL, Wu ZH, Sun Q, et al. Meta-analysis: the use of carbon dioxide insufflation vs. room air insufflation for gastrointestinal endoscopy. *Aliment Pharmacol Ther* 2012;35:1145–54.
- Wu J, Hu B. The role of carbon dioxide insufflation in colonoscopy: a systematic review and meta-analysis. *Endoscopy* 2012;44:128–36.
- Steiner SJ, Pfefferkorn MD, Fitzgerald JF. Patient-reported symptoms after pediatric outpatient colonoscopy or flexible sigmoidoscopy under general anesthesia. J Pediatr Gastroenterol Nut 2006; 43:483–6.
- Homan M, Mahkovic D, Orel R, et al. Randomized, double-blind trial of CO2 versus air insufflation in children undergoing colonoscopy. *Gastrointest Endosc* 2016;83:993–7.
- Thomson M, Venkatesh K, Elmalik K, et al. Double balloon enteroscopy in children: diagnosis, treatment, and safety. *World J Gastroenterol* 2010;16:56–62.
- Urs AN, Martinelli M, Rao P, et al. Diagnostic and therapeutic utility of double-balloon enteroscopy in children. J Pediatr Gastroenterol Nutr 2014;58:204–12.
- 41. De Bie CI, Buderus S, Sandhu BK, et al. Diagnostic workup of paediatric patients with inflammatory bowel disease in Europe: results of a 5-year audit of the EUROKIDS registry. *J Pediatr Gastroenterol Nutr* 2012;54:374–80.
- Zuber-Jerger I, Endlicher E, Gelbmann CM. Factors affecting cecal and ileal intubation time in colonoscopy. *Med Klin (Munich)* 2008;103:477–81.
- Sawczenko A, Sandhu BK. Presenting features of inflammatory bowel disease in Great Britain and Ireland. Arch Dis Child 2003;88:995– 1000.
- Thakkar K, Alsarraj A, Fong E, et al. Prevalence of colorectal polyps in pediatric colonoscopy. *Dig Dis Sci* 2012;57:1050–5.
- Lee CK, Shim JJ, Jang JY. Cold snare polypectomy vs. cold forceps polypectomy using double-biopsy technique for removal of diminutive colorectal polyps: a prospective randomized study. *Am J Gastroenterol* 2013;108:1593–600.
- Deenadayalu VP, Rex DK. Colon polyp retrieval after cold snaring. Gastrointest Endosc 2005;62:253–6.
- 47. Chino A, Karasawa T, Uragami N, et al. A comparison of depth of tissue injury caused by different modes of electrosurgical current in a pig colon model. *Gastrointest Endosc* 2004;59:374–9.
- Monkemuller KE, Fry LC, Jones BH, et al. Histological quality of polyps resected using the cold versus hot biopsy technique. *Endoscopy* 2004;36:432–6.
- ASGE Standards of Practice CommitteeIkenberry SO, Jue TL, Anderson MA, et al. . Management of ingested foreign bodies and food impactions. *Gastrointest Endosc* 2011;73:1085–91.
- Chu KM, Choi HK, Tuen HH, et al. A prospective randomized trial comparing the use of the flexible gastroscope versus the bronchoscope in the management of foreign body ingestion. *Gastrointest Endosc* 1998;47:23-7.
- Kim JK, Kim SS, Kim JI, et al. Management of foreign bodies in the gastrointestinal tract: an analysis of 104 cases in children. *Endoscopy* 1999;31:302–4.
- Jiraki K. Aortoesophageal conduit due to a foreign body. Am J Forensic Med Pathol 1996;17:347–8.
- Jayachandra S, Eslick GD. A systematic review of paediatric foreign body ingestion: presentation, complications, and management. *Int J Pediatr Otorhinolaryngol* 2013;77:311–7.
- Arana A, Hauser B, Hachimi-Idrissi S, et al. Management of ingested foreign bodies in childhood and review of the literature. *Eur J Pediatrics* 2001;160:468–72.

- Wright CC, Closson FT. Updates in pediatric gastrointestinal foreign bodies. *Pediatr Clin North Am* 2013;60:1221–39.
- Guelfguat M, Kaplinskiy V, Reddy SH, et al. Clinical guidelines for imaging and reporting ingested foreign bodies. *AJR Am J Roentgenol* 2014;203:37–53.
- 57. Hunter TB, Taljanovic MS. Foreign bodies. *Radiographics* 2003;23: 731–757.
- Hong SJ, Goo HW, Roh JL. Utility of spiral and cine CT scans in pediatric patients suspected of aspirating radiolucent foreign bodies. *Otolaryngol Head Neck Surg* 2008;138:576–80.
- Lee JB, Ahmad S, Gale CP. Detection of coins ingested by children using a handheld metal detector: a systematic review. *Emerg Med J* 2005;22:839–44.
- Moammar H, Al-Edreesi M, Abdi R. Sonographic diagnosis of gastricoutlet foreign body: case report and review of literature. *J Family Community Med* 2009;16:33–6.
- Ingraham CR, Mannelli L, Robinson JD, et al. Radiology of foreign bodies: how do we image them? *Emerg Radiol* 2015;22:425–30.
- 62. Loh KS, Tan LK, Smith JD, et al. Complications of foreign bodies in the esophagus. *Otolaryngol Head Neck Surg* 2000;123:613–6.
- 63. Park JH, Park CH, Park JH, et al. Review of 209 cases of foreign bodies in the upper gastrointestinal tract and clinical factors for successful endoscopic removal. *Korean J Gastroenterol* 2004;43:226–33.
- 64. Denney W, Ahmad N, Dillard B, et al. Children will eat the strangest things: a 10-year retrospective analysis of foreign body and caustic ingestions from a single academic center. *Pediatr Emerg Care* 2012;28:731–4.
- Chen X, Milkovich S, Stool D, et al. Pediatric coin ingestion and aspiration. Int J Pediatr Otorhinolaryngol 2006;70:325–9.
- Waltzman ML, Baskin M, Wypij D, et al. A randomized clinical trial of the management of esophageal coins in children. *Pediatrics* 2005;116:614–9.
- Tander B, Yazici M, Rizalar R, et al. Coin ingestion in children: which size is more risky? J Laparoendosc Adv Surg Tech A 2009;19:241–3.
- Kramer RE, Lerner DG, Lin T, et al. Management of ingested foreign bodies in children: a clinical report of the NASPGHAN Endoscopy Committee. J Pediatr Gastroenterol Nutr 2015;60:562–74.
- Reilly BK, Stool D, Chen X, et al. Foreign body injury in children in the twentieth century: a modern comparison to the Jackson collection. *Int J Pediatr Otorhinolaryngol* 2003;67(suppl 1):S171–4.
- Paul RI, Christoffel KK, Binns HJ, et al. Foreign body ingestions in children: risk of complication varies with site of initial health care contact. Pediatric Practice Research Group. *Pediatrics* 1993;91:121–7.
- Gregori D, Scarinzi C, Morra B, et al. Ingested foreign bodies causing complications and requiring hospitalization in European children: results from the ESFBI study. *Pediatr Intr* 2010;52:26–32.
- Farmakakis T, Dessypris N, Alexe DM, et al. Magnitude and objectspecific hazards of aspiration and ingestion injuries among children in Greece. *Int J Pediatr Otorhinolaryngol* 2007;71:317–24.
- Braumann C, Goette O, Menenakos C, et al. Laparoscopic removal of ingested pin penetrating the gastric wall in an immunosuppressed patient. *Surg Endosc* 2004;18:870.
- Mehran A, Podkameni D, Rosenthal R, et al. Gastric perforation secondary to ingestion of a sharp foreign body. JSLS 2005;9:91–3.
- Goh BK, Chow PK, Quah HM, et al. Perforation of the gastrointestinal tract secondary to ingestion of foreign bodies. World J Surg 2006;30:372–7.
- Akcam M, Kockar C, Tola HT, et al. Endoscopic removal of an ingested pin migrated into the liver and affixed by its head to the duodenum. *Gastrointest Endosc* 2009;69:382–4.
- Garcia-Segui A, Bercowsky E, Gomez-Fernandez I, et al. Late migration of a toothpick into the bladder: initial presentation with urosepsis and hydronephrosis. *Arch Esp Urol* 2012;65:626–9.
- Karadayi SSE, Nadir A, Kaptanoglu M. Wandering pins: case report. Cumhuriyet Med J 2009;31:300–2.
- Sai Prasad TR, Low Y, Tan CE, et al. Swallowed foreign bodies in children: report of four unusual cases. Ann Acad Med Singapore 2006;35:49–53.
- Pinero Madrona A, Fernandez Hernandez JA, Carrasco Prats M, et al. Intestinal perforation by foreign bodies. *Eur J Surg* 2000;166:307–9.
- Palta R, Sahota A, Bemarki A, et al. Foreign-body ingestion: characteristics and outcomes in a lower socioeconomic population with predominantly intentional ingestion. *Gastrointest Endosc* 2009;69: 426–33.

- Hameed K, Kamal Hassan M, Rehman S. Management of foreign bodies in the upper gastrointestinal tract with flexible endoscope. J Postgrad Med Inst 2011;25:433–5.
- Tokar B, Cevik AA, Ilhan H. Ingested gastrointestinal foreign bodies: predisposing factors for complications in children having surgical or endoscopic removal. *Pediatr Surg Int* 2007;23:135–9.
- Rodríguez-Hermosa JI, Codina-Cazador A, Sirvent JM, et al. Surgically treated perforations of the gastrointestinal tract caused by ingested foreign bodies. *Colorectal Dis* 2008;10:701–7.
- Litovitz T, Whitaker N, Clark L. Preventing battery ingestions: an analysis of 8648 cases. *Pediatrics* 2010;125:1178–83.
- Ferrante JOBC, Osterhout C, Gilchrist J. Injuries from batteries among children aged < 13 years—United States, 1995–2010. MMWR Morb Mortal Wkly Rep 2012:661–6.
- Spiers A, Jamil S, Whan E, et al. Survival of patient after aortooesophageal fistula following button battery ingestion. ANZ J Surg 2012;82:186–7.
- Honda S, Shinkai M, Usui Y, et al. Severe gastric damage caused by button battery ingestion in a 3-month-old infant. *J Pediatr Surg* 2010;45:e23–6.
- Brumbaugh DE, Colson SB, Sandoval JA, et al. Management of button battery-induced hemorrhage in children. J Pediatr Gastroenterol Nutr 2011;52:585–9.
- Litovitz T, Schmitz BF. Ingestion of cylindrical and button batteries: an analysis of 2382 cases. *Pediatrics* 1992;89:747–57.
- Bronstein AC, Spyker DA, Cantilena LR Jr et al. 2011 Annual report of the American Association of Poison Control Centers' National Poison Data System (NPDS): 29th Annual Report. *Clin Toxicol* 2012;50:911–1164.
- Sahn B, Mamula P, Ford CA. Review of foreign body ingestion and esophageal food impaction management in adolescents. J Adolesc Health 2014;55:260–6.
- Centers for Disease Control and Prevention. Gastrointestinal injuries from magnet ingestion in children—United States, 2003–2006. MMWR Morb Mortal Weekly Rep 2006;55:1296–300.
- Longstreth GF, Longstreth KJ, Yao JF. Esophageal food impaction: epidemiology and therapy. A retrospective, observational study. *Gastrointest Endosc* 2001;53:193–8.
- Byrne KR, Panagiotakis PH, Hilden K, et al. Retrospective analysis of esophageal food impaction: differences in etiology by age and gender. *Dig Dis Sci* 2007;52:717–21.
- Cheung KM, Oliver MR, Cameron DJ, et al. Esophageal eosinophilia in children with dysphagia. J Pediatr Gastroenterol Nutr 2003;37:498–503.
- 97. Lao J, Bostwick HE, Berezin S, et al. Esophageal food impaction in children. *Pediatr Emerg Care* 2003;19:402–7.
- Hurtado CW, Furuta GT, Kramer RE. Etiology of esophageal food impactions in children. J Pediatr Gastroenterol Nutr 2011;52:43–6.
- Weant KA, Weant MP. Safety and efficacy of glucagon for the relief of acute esophageal food impaction. Am J Health Syst Pharm 2012;69:573–7.
- Thimmapuram J, Oosterveen S, Grim R. Use of glucagon in relieving esophageal food bolus impaction in the era of eosinophilic esophageal infiltration. *Dysphagia* 2013;28:212–6.
- Beno S, Calello D, Baluffi A, et al. Pediatric body packing: drug smuggling reaches a new low. *Pediatr Emerg Care* 2005; 21:744–6.
- Chaves DM, Ishioka S, Felix VN, et al. Removal of a foreign body from the upper gastrointestinal tract with a flexible endoscope: a prospective study. *Endoscopy* 2004;36:887–92.
- 103. Gmeiner D, Von Rahden BH, Meco C, et al. Flexible versus rigid endoscopy for treatment of foreign body impaction in the esophagus. *Surg Endosc* 2007;21:2026–9.
- Russell R, Lucas A, Johnson J, et al. Extraction of esophageal foreign bodies in children: rigid versus flexible endoscopy. *Pediatr Surg Int* 2014;30:417–22.
- 105. Cetinkursun S, Sayan A, Demirbag S, et al. Safe removal of upper esophageal coins by using Magill forceps: two centers' experience. *Clin Pediatr* 2006;45:71–3.
- 106. Mehta D, Attia M, Quintana E, et al. Glucagon use for esophageal coin dislodgment in children: a prospective, double-blind, placebo-controlled trial. Acad Emerg Med 2001;8:200–3.

- Park KS. Evaluation and management of caustic injuries from ingestion of Acid or alkaline substances. *Clin Endosc* 2014;47:301–7.
- Contini S, Swarray-Deen A, Scarpignato C. Oesophageal corrosive injuries in children: a forgotten social and health challenge in developing countries. *Bull World Health Organ* 2009;87:950–4.
- 109. Riffat F, Cheng A. Pediatric caustic ingestion: 50 consecutive cases and a review of the literature. *Dis Esophagus* 2009;22:89–94.
- De la Esperanza Rueda-Valencia M, Sosa EV, Fernandez S, et al. Pediatric caustic ingestion: eight years experience. *Indian J Pediatr* 2015;82:381–2.
- 111. Johnson CM, Brigger MT. The public health impact of pediatric caustic ingestion injuries. *Arch Otolaryngol Head Neck Surg* 2012;138:1111–5.
- 112. Sahin S, Carman KB, Dinleyici EC. Acute poisoning in children: data of a pediatric emergency unit. *Iran J Ped* 2011;21:479–84.
- 113. Sanchez-Ramirez CA, Larrosa-Haro A, Vasquez-Garibay EM, et al. Socio-demographic factors associated with caustic substance ingestion in children and adolescents. *Int J Pediatr Otorhinolaryngol* 2012;76:253–6.
- Dogan Y, Erkan T, Cokugras FC, et al. Caustic gastroesophageal lesions in childhood: an analysis of 473 cases. *Clin Pediatr (Phila)* 2006;45:435–8.
- 115. Arevalo-Silva C, Eliashar R, Wohlgelernter J, et al. Ingestion of caustic substances: a 15-year experience. *Laryngoscope* 2006;116:1422–6.
- 116. Bramuzzo M, Amaddeo A, Facchina G, et al. Liquid detergent capsule ingestion: a new pediatric epidemic? *Pediatr Emerg Care* 2013;29:410–1.
- 117. Fraser L, Wynne D, Clement WA, et al. Liquid detergent capsule ingestion in children: an increasing trend. *Arch Dis Child* 2012;97:1007.
- 118. Lee HJ, Lee JH, Seo JM, et al. A single center experience of selfbougienage on stricture recurrence after surgery for corrosive esophageal strictures in children. *Yonsei Med J* 2010;51:202–5.
- 119. Karagiozoglou-Lampoudi T, Agakidis CH, Chryssostomidou S, et al. Conservative management of caustic substance ingestion in a pediatric department setting, short-term and long-term outcome. *Dis Esophagus* 2011;24:86–91.
- Temiz A, Oguzkurt P, Ezer SS, et al. Predictability of outcome of caustic ingestion by esophagogastroduodenoscopy in children. World J Gastroenterol 2012;18:1098–103.
- Baskin D, Urganci N, Abbasoglu L, et al. A standardised protocol for the acute management of corrosive ingestion in children. *Pediatr Surg Int* 2004;20:824–8.
- Havanond C, Havanond P. Initial signs and symptoms as prognostic indicators of severe gastrointestinal tract injury due to corrosive ingestion. J Emerg Med 2007;33:349–53.
- 123. Boskovic A, Stankovic I. Predictability of gastroesophageal caustic injury from clinical findings: is endoscopy mandatory in children? *Eur J Gastroenterol Hepatol* 2014;26:499–503.
- 124. Lamireau T, Rebouissoux L, Denis D, et al. Accidental caustic ingestion in children: is endoscopy always mandatory? J Pediatr Gastroenterol Nutr 2001;33:81–4.
- Gupta SK, Croffie JM, Fitzgerald JF. Is esophagogastroduodenoscopy necessary in all caustic ingestions? J Pediatr Gastroenterol Nutr 2001;32:50–3.
- Betalli P, Falchetti D, Giuliani S, et al. Caustic ingestion in children: is endoscopy always indicated? The results of an Italian multicenter observational study. *Gastrointest Endosc* 2008;68:434–9.
- 127. Kay M, Wyllie R. Caustic ingestions in children. *Curr Opin Pediatr* 2009;21:651-4.
- Chen TY, Ko SF, Chuang JH, et al. Predictors of esophageal stricture in children with unintentional ingestion of caustic agents. *Chang Gung Med J* 2003;26:233–9.
- 129. Zargar SA, Kochhar R, Mehta S, et al. The role of fiberoptic endoscopy in the management of corrosive ingestion and modified endoscopic classification of burns. *Gastrointest Endosc* 1991;37:165–9.
- Temiz A, Oguzkurt P, Ezer SS, et al. Management of pyloric stricture in children: endoscopic balloon dilatation and surgery. *Surg Endosc* 2012;26:1903–8.
- 131. Usta M, Erkan T, Cokugras FC, et al. High doses of methylprednisolone in the management of caustic esophageal burns. *Pediatrics* 2014;133:E1518–24.

- Boukthir S, Fetni I, Mrad SM, et al. High doses of steroids in the management of caustic esophageal burns in children. *Arch Pediatr* 2004;11:13–7.
- 133. Pelclova D, Navratil T. Do corticosteroids prevent oesophageal stricture after corrosive ingestion? *Toxicol Rev* 2005;24:125–9.
- Pearson EG, Downey EC, Barnhart DC, et al. Reflux esophageal stricture—a review of 30 years' experience in children. J Pediatr Surg 2010;45:2356–60.
- 135. Ozdemir R, Bayrakci B, Teksam O, et al. Thirty-three-year experience on childhood poisoning. *Turk J Pediatr* 2012;54:251–9.
- 136. Urganci N, Usta M, Kalyoncu D, et al. Corrosive substance ingestion in children. *Indian J Pediatr* 2014;81:675–9.
- 137. Lan LC, Wong KK, Lin SC, et al. Endoscopic balloon dilatation of esophageal strictures in infants and children: 17 years' experience and a literature review. *J Pediatr Surg* 2003;38:1712–5.
- 138. Wilsey MJ Jr, Scheimann AO, Gilger MA. The role of upper gastrointestinal endoscopy in the diagnosis and treatment of caustic ingestion, esophageal strictures, and achalasia in children. *Gastrointest Endosc Clin N Am* 2001;11:767–87vii–viii.
- 139. Koivusalo A, Turunen P, Rintala RJ, et al. Is routine dilatation after repair of esophageal atresia with distal fistula better than dilatation when symptoms arise? Comparison of results of two European pediatric surgical centers. J Pediatr Surg 2004;39:1643–7.
- Koivusalo A, Pakarinen MP, Rintala RJ. Anastomotic dilatation after repair of esophageal atresia with distal fistula. Comparison of results after routine versus selective dilatation. *Dis Esophagus* 2009;22:190–4.
- 141. Jayakrishnan VK, Wilkinson AG. Treatment of oesophageal strictures in children: a comparison of fluoroscopically guided balloon dilatation with surgical bouginage. *Pediatr Radiol* 2001;31:98–101.
- 142. Thyoka M, Timmis A, Mhango T, et al. Balloon dilatation of anastomotic strictures secondary to surgical repair of oesophageal atresia: a systematic review. *Pediatr Radiol* 2013;43:898–901.
- 143. Uygun I, Arslan MS, Aydogdu B, et al. Fluoroscopic balloon dilatation for caustic esophageal stricture in children: an 8-year experience. *J Pediatr Surg* 2013;48:2230–4.
- Cakmak M, Boybeyi O, Gollu G, et al. Endoscopic balloon dilatation of benign esophageal strictures in childhood: a 15-year experience. *Dis Esophagus* 2016;29:179–84.
- Lang T, Hummer HP, Behrens R. Balloon dilation is preferable to bougienage in children with esophageal atresia. *Endoscopy* 2001;33:329–35.
- 146. Poddar U, Thapa BR. Benign esophageal strictures in infants and children: results of Savary-Gilliard bougie dilation in 107 Indian children. *Gastrointest Endosc* 2001;54:480–4.
- Alshammari J, Quesnel S, Pierrot S, et al. Endoscopic balloon dilatation of esophageal strictures in children. *Int J Pediatr Otorhinolaryngol* 2011;75:1376–9.
- Kochman ML, McClave SA, Boyce HW. The refractory and the recurrent esophageal stricture: a definition. *Gastrointest Endosc* 2005;62:474–5.
- 149. Uhlen S, Fayoux P, Vachin F, et al. Mitomycin C: an alternative conservative treatment for refractory esophageal stricture in children? *Endoscopy* 2006;38:404–7.
- 150. Chapuy L, Pomerleau M, Faure C. Topical mitomycin-C application in recurrent esophageal strictures after surgical repair of esophageal atresia. *J Pediatr Gastroenterol Nutr* 2014;59:608–11.
- 151. Berger M, Ure B, Lacher M. Mitomycin C in the therapy of recurrent esophageal strictures: hype or hope? *Eur J Pediatr Surg* 2012;22:109–16.
- Rosseneu S, Afzal N, Yerushalmi B, et al. Topical application of mitomycin-C in oesophageal strictures. J Pediatr Gastroenterol Nutr 2007;44:336–41.
- 153. El-Asmar KM, Hassan MA, Abdelkader HM, et al. Topical mitomycin C application is effective in management of localized caustic esophageal stricture: a double-blinded, randomized, placebo-controlled trial. *J Pediatr Surg* 2013;48: 1621–1627.
- 154. Kramer RE, Quiros JA. Esophageal stents for severe strictures in young children: experience, benefits, and risk. *Curr Gastroenterol Rep* 2010;12:203–10.
- 155. Manfredi MA, Jennings RW, Anjum MW, et al. Externally removable stents in the treatment of benign recalcitrant strictures and esophageal perforations in pediatric patients with esophageal atresia. *Gastrointest Endosc* 2014;80:246–52.

- Bicakci U, Tander B, Deveci G, et al. Minimally invasive management of children with caustic ingestion: less pain for patients. *Pediatr Surg Int* 2010;26:251–5.
- 157. Hirdes MM, Van Hooft JE, Koornstra JJ, et al. Endoscopic corticosteroid injections do not reduce dysphagia after endoscopic dilation therapy in patients with benign esophagogastric anastomotic strictures. *Clin Gastroenterol Hepatol* 2013;11: 795–801.
- 158. Connor MJ, Springford LR, Kapetanakis VV, et al. Esophageal atresia and transitional care-step 1: a systematic review and meta-analysis of the literature to define the prevalence of chronic long-term problems. *Am J Surg* 2015;209:747–59.
- 159. Vergouwe FW, H IJ, Wijnen RM, et al. Screening and surveillance in esophageal atresia patients: current knowledge and future perspectives. *Eur J Pediatr Surg* 2015;25:345–52.
- Castilloux J, Bouron-Dal Soglio D, Faure C. Endoscopic assessment of children with esophageal atresia: lack of relationship of esophagitis and esophageal metaplasia to symptomatology. *Can J Gastroenterol* 2010;24:312–6.
- 161. Schneider A, Gottrand F, Bellaiche M, et al. Prevalence of Barrett esophagus in adolescents and young adults with esophageal atresia. *Ann Surg* 2016 May 26 [Epub ahead of print].
- 162. Enns RA, Gagnon YM, Barkun AN, et al. Validation of the Rockall scoring system for outcomes from non-variceal upper gastrointestinal bleeding in a Canadian setting. *World J Gastroenterol* 2006;12: 7779–7785.
- 163. Custodio Lima J, Garcia Montes C, Kibune Nagasako C, et al. Performance of the Rockall scoring system in predicting the need for intervention and outcomes in patients with nonvariceal upper gastrointestinal bleeding in a Brazilian setting: a prospective study. *Digestion* 2013;88:252–7.
- 164. Bessa X, O'Callaghan E, Balleste B, et al. Applicability of the Rockall score in patients undergoing endoscopic therapy for upper gastrointestinal bleeding. *Dig Liver Dis* 2006;38:12–7.
- 165. Thomson MA, Leton N, Belsha D. Acute upper gastrointestinal bleeding in childhood: development of the Sheffield Scoring System to predict need for endoscopic therapy. J Pediatr Gastroenterol Nutr 2015;60:632–6.
- 166. Fritscher-Ravens A, Scherbakov P, Bufler P, et al. The feasibility of wireless capsule endoscopy in detecting small intestinal pathology in children under the age of 8 years: a multicentre European study. *Gut* 2009;58:1467–72.
- 167. Thomson M, Fritscher-Ravens A, Mylonaki M, et al. Wireless capsule endoscopy in children: a study to assess diagnostic yield in small bowel disease in paediatric patients. J Pediatr Gastroenterol Nutr 2007;44:192–7.
- 168. Yachha SK, Khanduri A, Sharma BC, et al. Gastrointestinal bleeding in children. J Gastroenterol Hepatol 1996;11:903–7.
- Gralnek IM, Barkun AN, Bardou M. Management of acute bleeding from a peptic ulcer. N Engl J Med 2008;359:928–37.
- Spiegel BM, Vakil NB, Ofman JJ. Endoscopy for acute nonvariceal upper gastrointestinal tract hemorrhage: is sooner better? A systematic review. Arch Intern Med 2001;161:1393–404.
- 171. Cooper GS, Chak A, Way LE, et al. Early endoscopy in upper gastrointestinal hemorrhage: associations with recurrent bleeding, surgery, and length of hospital stay. *Gastrointest Endosc* 1999;49: 145–52.
- 172. Colle I, Wilmer A, Le Moine O, et al. Upper gastrointestinal tract bleeding management: Belgian guidelines for adults and children. *Acta Gastroenterol Belg* 2011;74:45–66.
- 173. Adamopoulos AB, Baibas NM, Efstathiou SP, et al. Differentiation between patients with acute upper gastrointestinal bleeding who need early urgent upper gastrointestinal endoscopy and those who do not. A prospective study. *Eur J Gastroenterol Hepatol* 2003;15:81–7.
- 174. Gralnek IM, Dumonceau JM, Kuipers EJ, et al. Diagnosis and management of nonvariceal upper gastrointestinal hemorrhage: European Society of Gastrointestinal Endoscopy (ESGE) Guideline. *Endoscopy* 2015;47:a1–46.
- 175. Van Leerdam ME, Rauws EA, Geraedts AA, et al. Management in peptic ulcer hemorrhage: a Dutch national inquiry. *Endoscopy* 2000;32:935–42.

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- 176. Jensen DM, Machicado GA, Jutabha R, et al. Urgent colonoscopy for the diagnosis and treatment of severe diverticular hemorrhage. N Engl J Med 2000;342:78–82.
- 177. Elta GH. Urgent colonoscopy for acute lower-GI bleeding. *Gastro-intest Endosc* 2004;59:402–8.
- Gershman G, Thomson M. Practical Pediatric Gastrointestinal Endoscopy. Wiley-Blackwell: ; 2012:88–94.
- 179. Senger JL, Kanthan R. The evolution of Dieulafoy's lesion since 1897: then and now—a journey through the lens of a pediatric lesion with literature review. *Gastroenterol Res Pract* 2012;2012:432517.
- Valera JM, Pino RQ, Poniachik J, et al. Endoscopic band ligation of bleeding Dieulafoy lesions: the best therapeutic strategy. *Endoscopy* 2006;38:193–4.
- Garg R. Bleeding from a gastric Dieulafoy lesion. *Emerg Med J* 2007;24:520.
- Lokesh Babu TG, Jacobson K, Phang M, et al. Endoscopic hemostasis in a neonate with a bleeding duodenal ulcer. *J Pediatr Gastroenterol Nut* 2005;41:244–6.
- 183. Khan K, Schwarzenberg SJ, Sharp H, et al. Argon plasma coagulation: clinical experience in pediatric patients. *Gastrointest Endosc* 2003;7:110–2.
- 184. [Anonymous]. NICE Guideline recommendation for Non-variceal bleeding endoscopic treatment. https://www.nice.org.uk/Guidance/ CG141. Accessed May 1, 2016.
- 185. Zargar SA, Javid G, Khan BA, et al. Endoscopic ligation compared with sclerotherapy for bleeding esophageal varices in children with extrahepatic portal venous obstruction. *Hepatology* 2002;36:666–72.
- Shneider BL, Abel B, Haber B, et al. Portal hypertension in children and young adults with biliary atresia. J Pediatr Gastroenterol Nutr 2012;55:567–73.
- 187. Kim SJ, Oh SH, Jo JM, et al. Experiences with endoscopic interventions for variceal bleeding in children with portal hypertension: a single center study. *Pediatr Gastroenterol Hepatol Nutr* 2013;16: 248–253.
- 188. Duche M, Ducot B, Ackermann O, et al. Experience with endoscopic management of high-risk gastroesophageal varices, with and without bleeding, in children with biliary atresia. *Gastroenterology* 2013;145:801–7.
- Rivet C, Robles-Medranda C, Dumortier J, et al. Endoscopic treatment of gastroesophageal varices in young infants with cyanoacrylate glue: a pilot study. *Gastrointest Endosc* 2009;69:1034–8.
- 190. McKiernan PJ, Beath SV, Davison SM. A prospective study of endoscopic esophageal variceal ligation using a multiband ligator. J Pediatr Gastroenterol Nut 2002;34:207–11.
- 191. Shaheen NJ, Stuart E, Schmitz SM, et al. Pantoprazole reduces the size of postbanding ulcers after variceal band ligation: a randomized, controlled trial. *Hepatology* 2005;41:588–94.
- 192. Zurera LJ, Espejo JJ, Lombardo S, et al. Safety and efficacy of expanded polytetrafluoroethylene-covered transjugular intrahepatic portosystemic shunts in children with acute or recurring upper gastrointestinal bleeding. *Pediatr Radiol* 2015;45:422–9.
- 193. Tam YH, Lee KH, To KF, et al. *Helicobacter pylori*-positive versus *Helicobacter pylori*-negative idiopathic peptic ulcers in children with their long-term outcomes. *J Pediatr Gastroenterol Nut* 2009;48:299–305.
- 194. Nuutinen H, Kolho KL, Salminen P, et al. Capsule endoscopy in pediatric patients: technique and results in our first 100 consecutive children. *Scan J Gastroenterol* 2011;46:1138–43.
- 195. Cohen SA, Klevens AI. Use of capsule endoscopy in diagnosis and management of pediatric patients, based on meta-analysis. *Clin Gastroenterol Hepatol* 2011;9:490–6.
- 196. Shen R, Sun B, Gong B, et al. Double-balloon enteroscopy in the evaluation of small bowel disorders in pediatric patients. *Dig Endosc* 2012;24:87–92.
- 197. Vegting IL, Tabbers MM, Taminiau JA, et al. Is endoscopic retrograde cholangiopancreatography valuable and safe in children of all ages? *J Pediatr Gastroenterol Nutr* 2009;48:66–71.
- Limketkai BN, Chandrasekhara V, Kalloo AN, et al. Comparison of performance and safety of endoscopic retrograde cholangiopancreatography across pediatric age groups. *Dig Dis Sci* 2013;58:2653–60.
- 199. Philpott C, Rosenbaum J, Moon A, et al. Paediatric MRCP: 10 year experience with 195 patients. *Eur J Radiol* 2013;82:699–706.

- Delaney L, Applegate KE, Karmazyn B, et al. MR cholangiopancreatography in children: feasibility, safety, and initial experience. *Pediatr Radiol* 2008;38:64–75.
- Varadarajulu S, Wilcox CM, Hawes RH, et al. Technical outcomes and complications of ERCP in children. *Gastrointest Endosc* 2004;60:367–71.
- Iqbal CW, Baron TH, Moir CR, et al. Post-ERCP pancreatitis in pediatric patients. J Pediatr Gastroenterol Nutr 2009;49:430–4.
- Teng R, Yokohata K, Utsunomiya N, et al. Endoscopic retrograde cholangiopancreatography in infants and children. J Gastroenterol 2000;35:39–42.
- Poddar U, Thapa BR, Bhasin DK, et al. Endoscopic retrograde cholangiopancreatography in the management of pancreaticobiliary disorders in children. J Gastroenterol Hepatol 2001;16:927–31.
- Pfau PR, Chelimsky GG, Kinnard MF, et al. Endoscopic retrograde cholangiopancreatography in children and adolescents. J Pediatr Gastroenterol Nutr 2002;35:619–23.
- Cheng CL, Fogel EL, Sherman S, et al. Diagnostic and therapeutic endoscopic retrograde cholangiopancreatography in children: a large series report. *J Pediatr Gastroenterol Nutr* 2005;41:445–53.
- Issa H, Al-Haddad A, Al-Salem AH. Diagnostic and therapeutic ERCP in the pediatric age group. *Pediatr Surg Int* 2007;23:111–6.
- Jang JY, Yoon CH, Kim KM. Endoscopic retrograde cholangiopancreatography in pancreatic and biliary tract disease in Korean children. *World J Gastroenterol* 2010;16:490–5.
- Otto AK, Neal MD, Slivka AN, et al. An appraisal of endoscopic retrograde cholangiopancreatography (ERCP) for pancreaticobiliary disease in children: our institutional experience in 231 cases. *Surg Endosc* 2011;25:2536–40.
- Enestvedt BK, Tofani C, Lee DY, et al. Endoscopic retrograde cholangiopancreatography in the pediatric population is safe and efficacious. J Pediatr Gastroenterol Nutr 2013;57:649–54.
- 211. Halvorson L, Halsey K, Darwin P, et al. The safety and efficacy of therapeutic ERCP in the pediatric population performed by adult gastroenterologists. *Dig Dis Sci* 2013;58:3611–9.
- 212. Troendle DM, Barth BA. ERCP can be safely and effectively performed by a pediatric gastroenterologist for choledocholithiasis in a pediatric facility. *J Pediatr Gastroenterol Nutr* 2013;57:655–8.
- 213. Kieling CO, Hallal C, Spessato CO, et al. Changing pattern of indications of endoscopic retrograde cholangiopancreatography in children and adolescents: a twelve-year experience. *World J Pediatr* 2015;11:154–9.
- 214. Troendle DM, Abraham O, Huang R, et al. Factors associated with post-ERCP pancreatitis and the effect of pancreatic duct stenting in a pediatric population. *Gastrointest Endosc* 2015;81:1408–16.
- Dumonceau JM, Delhaye M, Tringali A, et al. Endoscopic treatment of chronic pancreatitis: European Society of Gastrointestinal Endoscopy (ESGE) Clinical Guideline. *Endoscopy* 2012;44:784–800.
- Agarwal J, Nageshwar Reddy D, Talukdar R, et al. ERCP in the management of pancreatic diseases in children. *Gastrointest Endosc* 2014;79:271–8.
- 217. Oracz G, Pertkiewicz J, Kierkus J, et al. Efficiency of pancreatic duct stenting therapy in children with chronic pancreatitis. *Gastrointest Endosc* 2014;80:1022–9.
- 218. Moyer V, Freese DK, Whitington PF, et al. Guideline for the evaluation of cholestatic jaundice in infants: recommendations of the North American Society for Pediatric Gastroenterology, Hepatology and Nutrition. J Pediatr Gastroenterol Nutr 2004;39:115–28.
- 219. Siles P, Aschero A, Gorincour G, et al. A prospective pilot study: can the biliary tree be visualized in children younger than 3 months on magnetic resonance cholangiopancreatography? *Pediatr Radiol* 2014;44:1077–84.
- 220. Liu B, Cai J, Xu Y, et al. Three-dimensional magnetic resonance cholangiopancreatography for the diagnosis of biliary atresia in infants and neonates. *PLoS One* 2014;9:e88268.
- 221. Keil R, Snajdauf J, Rygl M, et al. Diagnostic efficacy of ERCP in cholestatic infants and neonates—a retrospective study on a large series. *Endoscopy* 2010;42:121–6.
- 222. Iinuma Y, Narisawa R, Iwafuchi M, et al. The role of endoscopic retrograde cholangiopancreatography in infants with cholestasis. *J Pediatr Surg* 2000;35:545–9.
- 223. Aabakken L, Aagenaes I, Sanengen T, et al. Utility of ERCP in neonatal and infant cholestasis. J Laparoendosc Adv Surg Tech A 2009;19:431–6.

- 224. Petersen C, Meier PN, Schneider A, et al. Endoscopic retrograde cholangiopancreaticography prior to explorative laparotomy avoids unnecessary surgery in patients suspected for biliary atresia. *J Hepatol* 2009;51:1055–60.
- 225. Shanmugam NP, Harrison PM, Devlin J, et al. Selective use of endoscopic retrograde cholangiopancreatography in the diagnosis of biliary atresia in infants younger than 100 days. J Pediatr Gastroenterol Nutr 2009;49:435–41.
- Shteyer E, Wengrower D, Benuri-Silbiger I, et al. Endoscopic retrograde cholangiopancreatography in neonatal cholestasis. J Pediatr Gastroenterol Nutr 2012;55:1425.
- 227. Saito T, Terui K, Mitsunaga T, et al. Role of pediatric endoscopic retrograde cholangiopancreatography in an era stressing less-invasive imaging modalities. *J Pediatric Gastroenterol Nutr* 2014; 59:204–9.
- 228. Green JA, Scheeres DE, Conrad HA, et al. Pediatric ERCP in a multidisciplinary community setting: experience with a fellowship-trained general surgeon. *Surg Endosc* 2007;21:2187–92.
- Paris C, Bejjani J, Beaunoyer M, et al. Endoscopic retrograde cholangiopancreatography is useful and safe in children. *J Pediatr Surg* 2010;45:938–42.
- Leichtner AM, Gillis LA, Gupta S, et al. NASPGHAN guidelines for training in pediatric gastroenterology. J Pediatr Gastroenterol Nutr 2013;56(suppl 1):S1–8.
- Dumonceau JM, Andriulli A, Elmunzer BJ, et al. Prophylaxis of post-ERCP pancreatitis: European Society of Gastrointestinal Endoscopy (ESGE) Guideline—updated June 2014. *Endoscopy* 2014;46: 799–815.
- Dumonceau JM, Garcia-Fernandez FJ, Verdun FR, et al. Radiation protection in digestive endoscopy: European Society of Digestive Endoscopy (ESGE) guideline. *Endoscopy* 2012;44:408–21.
- Scheers I, Ergun M, Aouattah T, et al. Diagnostic and therapeutic roles of endoscopic ultrasound in pediatric pancreaticobiliary disorders. *J Pediatr Gastroenterol Nutr* 2015;61:238–47.
- Romeo E, Foschia F, De Angelis P, et al. Endoscopic management of congenital esophageal stenosis. J Pediatr Surg 2011;46:838–41.
- 235. Bocus P, Realdon S, Eloubeidi MA, et al. High-frequency miniprobes and 3-dimensional EUS for preoperative evaluation of the etiology of congenital esophageal stenosis in children (with video). *Gastrointest Endosc* 2011;74:204–7.
- Fox VL, Nurko S, Teitelbaum JE, et al. High-resolution EUS in children with eosinophilic "allergic" esophagitis. *Gastrointest Endosc* 2003;57:30–6.
- McKiernan PJ, Sharif K, Gupte GL. The role of endoscopic ultrasound for evaluating portal hypertension in children being assessed for intestinal transplantation. *Transplantation* 2008;86:1470–3.

- De Angelis P, Romeo E, Rea F, et al. Miniprobe EUS in management of pancreatic pseudocyst. World J Gastrointest Endosc 2013;5:255–60.
- 239. Takamizawa S, Tsugawa C, Mouri N, et al. Congenital esophageal stenosis: therapeutic strategy based on etiology. *J Pediatr Surg* 2002;37:197–201.
- 240. Usui N, Kamata S, Kawahara H, et al. Usefulness of endoscopic ultrasonography in the diagnosis of congenital esophageal stenosis. *J Pediatr Surg* 2002;37:1744–6.
- 241. Varadarajulu S, Wilcox CM, Eloubeidi MA. Impact of EUS in the evaluation of pancreaticobiliary disorders in children. *Gastrointest Endosc* 2005;62:239–44.
- 242. Attila T, Adler DG, Hilden K, et al. EUS in pediatric patients. *Gastrointest Endosc* 2009;70:892-8.
- 243. Al-Rashdan A, LeBlanc J, Sherman S, et al. Role of endoscopic ultrasound for evaluating gastrointestinal tract disorders in pediatrics: a tertiary care center experience. J Pediatr Gastroenterol Nutr 2010;51:718–22.
- 244. Buxbaum JL, Eloubeidi MA, Varadarajulu S. Utility of EUS-guided FNA in the management of children with idiopathic fibrosing pancreatitis. *J Pediatr Gastroenterol Nutr* 2011;52:482–4.
- 245. Fujii LL, Chari ST, El-Youssef M, et al. Pediatric pancreatic EUSguided trucut biopsy for evaluation of autoimmune pancreatitis. *Gastrointest Endosc* 2013;77:824–8.
- 246. Sheers I, Ergun MA, Tarik, et al. Diagnostic and therapeutic role of endoscopic ultrasound in pediatric pancreaticobiliary disorders. *J Pediatr Gastroenterol Nutr* 2015;61:238–47.
- Terui K, Saito T, Mitsunaga T, et al. Endoscopic management for congenital esophageal stenosis: a systematic review. World J Gastrointest Endosc 2015;7:183–91.
- 248. Dalby K, Nielsen RG, Kruse-Andersen S, et al. Gastroesophageal reflux disease and eosinophilic esophagitis in infants and children. A study of esophageal pH, multiple intraluminal impedance and endoscopic ultrasound. *Scand J Gastroenterol* 2010;45:1029–35.
- El-Karaksy HM, El-Koofy NM, Okasha H, et al. A comparative study of endoscopic ultrasonography versus endoscopic retrograde cholangiopancreatography in children with chronic liver disease. *Indian J Med Sci* 2008;62:345–51.
- 250. Ramesh J, Bang JY, Trevino J, et al. Endoscopic ultrasound-guided drainage of pancreatic fluid collections in children. *J Pediatr Gastroenterol Nutr* 2013;56:30–5.
- 251. Jazrawi SF, Barth BA, Sreenarasimhaiah J. Efficacy of endoscopic ultrasound-guided drainage of pancreatic pseudocysts in a pediatric population. *Dig Dis Sci* 2011;56:902–8.
- 252. McCann F, Michaud L, Aspirot A, et al. Congenital esophageal stenosis associated with esophageal atresia. *Dis Esophagus* 2015;28:211–5.