



S.I.C.O.B.

XXXII CONGRESSO  
NAZIONALE SICOB

23 - 25 MAGGIO 2024  
GIARDINI  
NAXOS



# Reflusso Duodeno Gastro Esofageo e Prevenzione

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**GASTROENTEROLOGIA**

**UNIVERSITÀ DI SALERNO**



# What is BILE reflux?



When the material produced by the pancreas and the liver gets into the duodenum, it is mixed with duodenal fluid and form a solution that contains also bile



# What is GERD?



**When this solution gets into the stomach  
and then up into the esophagus,  
It is called BILE REFLUX.**



# What is BILE reflux?



**Bile reflux is ALSO named  
DUODENO-GASTRO-ESOPHAGEAL  
REFLUX**





Normally, there is always a certain amount of duodeno-pancreatic secretion that goes into the stomach, most commonly at night and postprandially. In the stomach, this solution is mixed with gastric content, which is, most of the time, acid.

Keanetal. 1981; Sonnenbergetal. 1982; Thompson 1982; Heading 1983; Muller-Lissner 1983),





# What is GERD?



**Reflux of gastric contents to the esophagus is a physiological event: a healthy person typically has reflux episodes.**



# What is GERD?



**GERD is a condition which develops when the reflux of stomach contents causes troublesome symptoms and/or complications**







**'troublesome symptoms' =  
MODERATE-SEVERE symptoms that  
occur more than once a week**

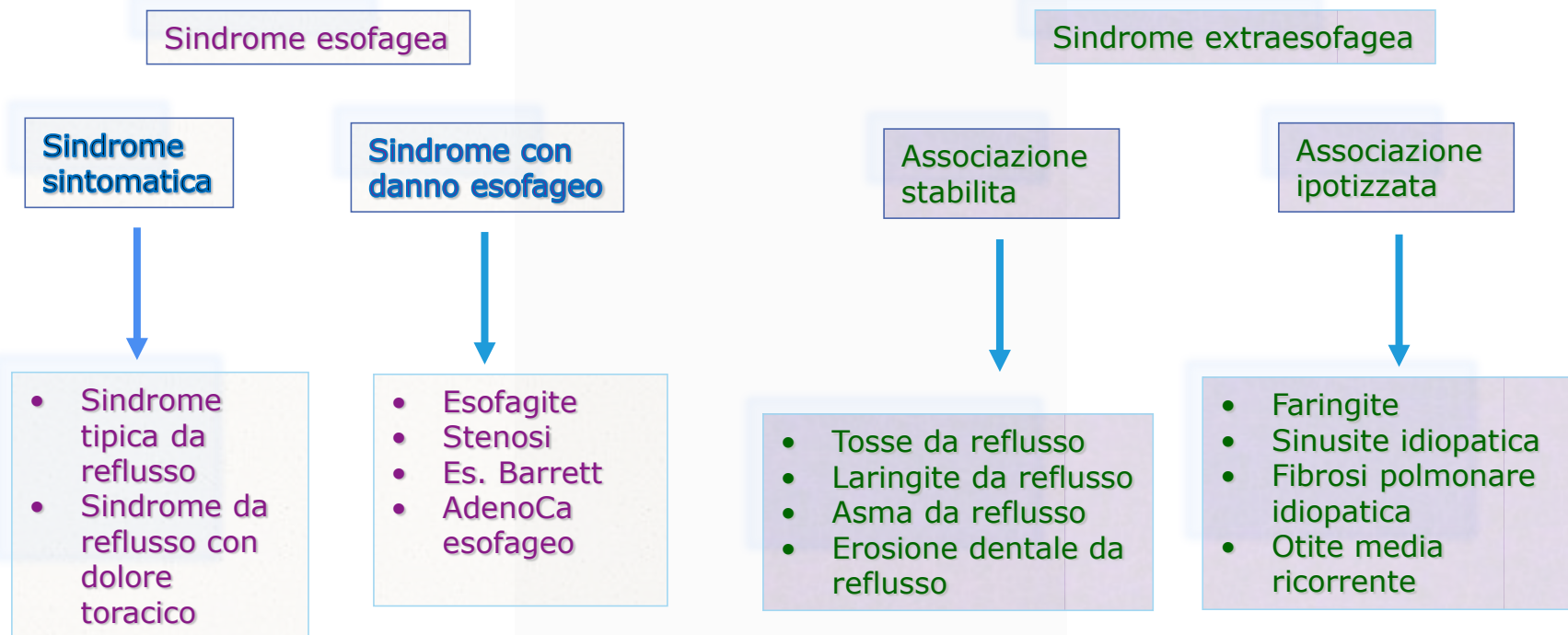


24%



Zagari et al., Gut, 2008

La MRGE è una condizione che si sviluppa quando un reflusso di contenuto gastrico causa sintomi o complicanze fastidiosi





# How is bile reflux distinguished from acidic reflux?

- ◎ **It is not possible to distinguish BILE REFLUX from ACIDIC REFLUX in terms of signs and symptoms.**



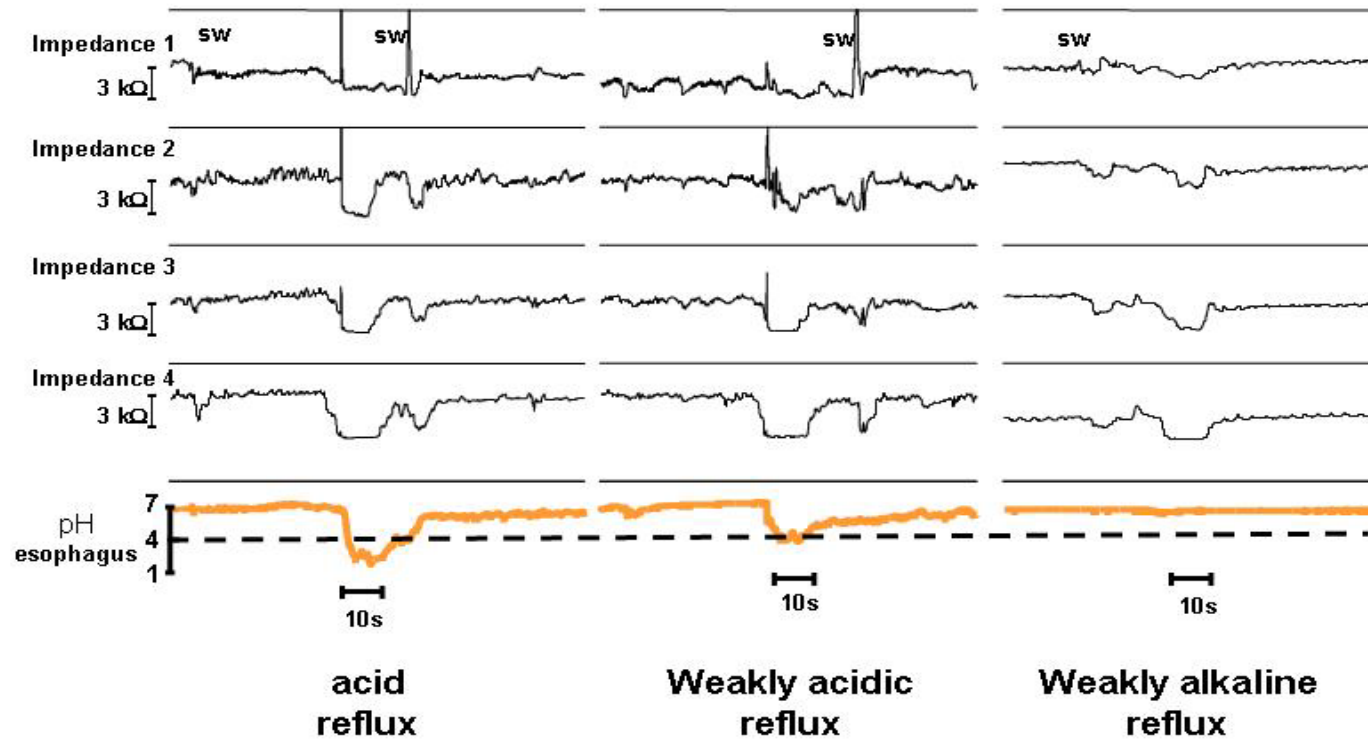
# Other than acidic reflux, are there any conditions with which bile reflux can be confused?


## Bile reflux $\neq$ nonacidic reflux

- ✓ **Nonacidic reflux is a type of refluxate that can be recognized only by pH-impedance monitoring.**
- ✓ **Nonacidic reflux might or might not contain bile.**
- ✓ **Bile is more often associated with acidic gastric juice than with a nonacidic component of gastric contents.**



# Patterns of esophageal reflux





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**symptoms**

**endoscopic  
changes,**

**histologic features of  
a chemical (reactive)  
gastritis)**



Method	Advantages	Disadvantages
Endoscopy	<ul style="list-style-type: none"><li>• Easy visualization</li></ul>	<ul style="list-style-type: none"><li>• Poor sensitivity/specificity of bile</li><li>• Positive predictive value</li><li>• Requires sedation</li><li>• High cost</li></ul>
Aspiration studies	<ul style="list-style-type: none"><li>• Less invasive than endoscopy</li><li>• No sedation</li><li>• Low cost</li><li>• Noninvasive</li></ul>	<ul style="list-style-type: none"><li>• Short duration of study</li><li>• Requires familiarity with enzymatic assay for BA</li></ul>
Scintigraphy	<ul style="list-style-type: none"><li>• Noninvasive</li></ul>	<ul style="list-style-type: none"><li>• Semiquantitative at best</li><li>• Radiation exposure</li><li>• High cost</li></ul>
pH monitoring	<ul style="list-style-type: none"><li>• Easy to perform</li><li>• Relatively noninvasive</li><li>• Prolonged monitoring</li><li>• Ambulatory</li></ul>	<ul style="list-style-type: none"><li>• pH &gt;7 not a marker for DGER</li><li>• Not specific for DGER</li></ul>
Bilirubin monitoring (Bilitec)	<ul style="list-style-type: none"><li>• Easy to perform</li><li>• Relatively noninvasive</li><li>• Prolonged monitoring</li><li>• Ambulatory</li><li>• Good correlation with gastric BA concentrations</li></ul>	<ul style="list-style-type: none"><li>• Current design underestimates DGER by ~30% in acidic medium (pH &lt;3.5)</li><li>• Requires modified diet</li></ul>

BA = bile acid.



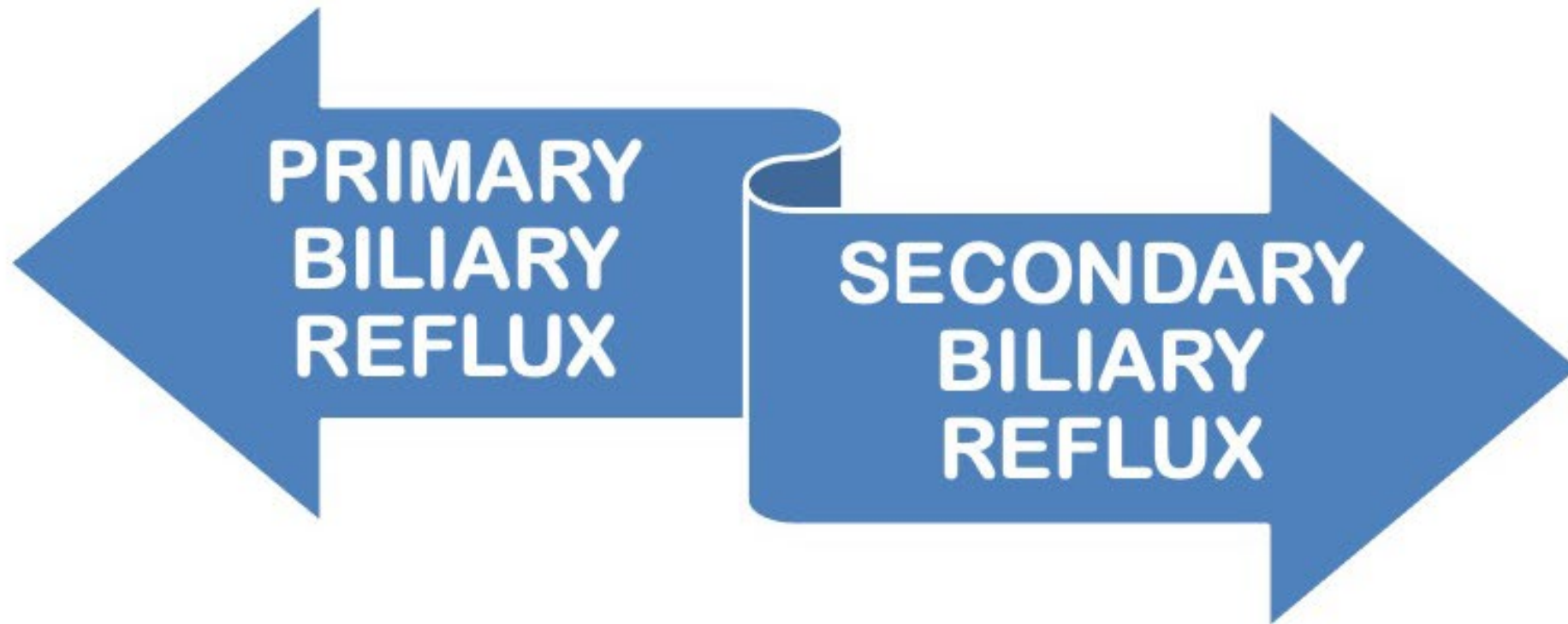


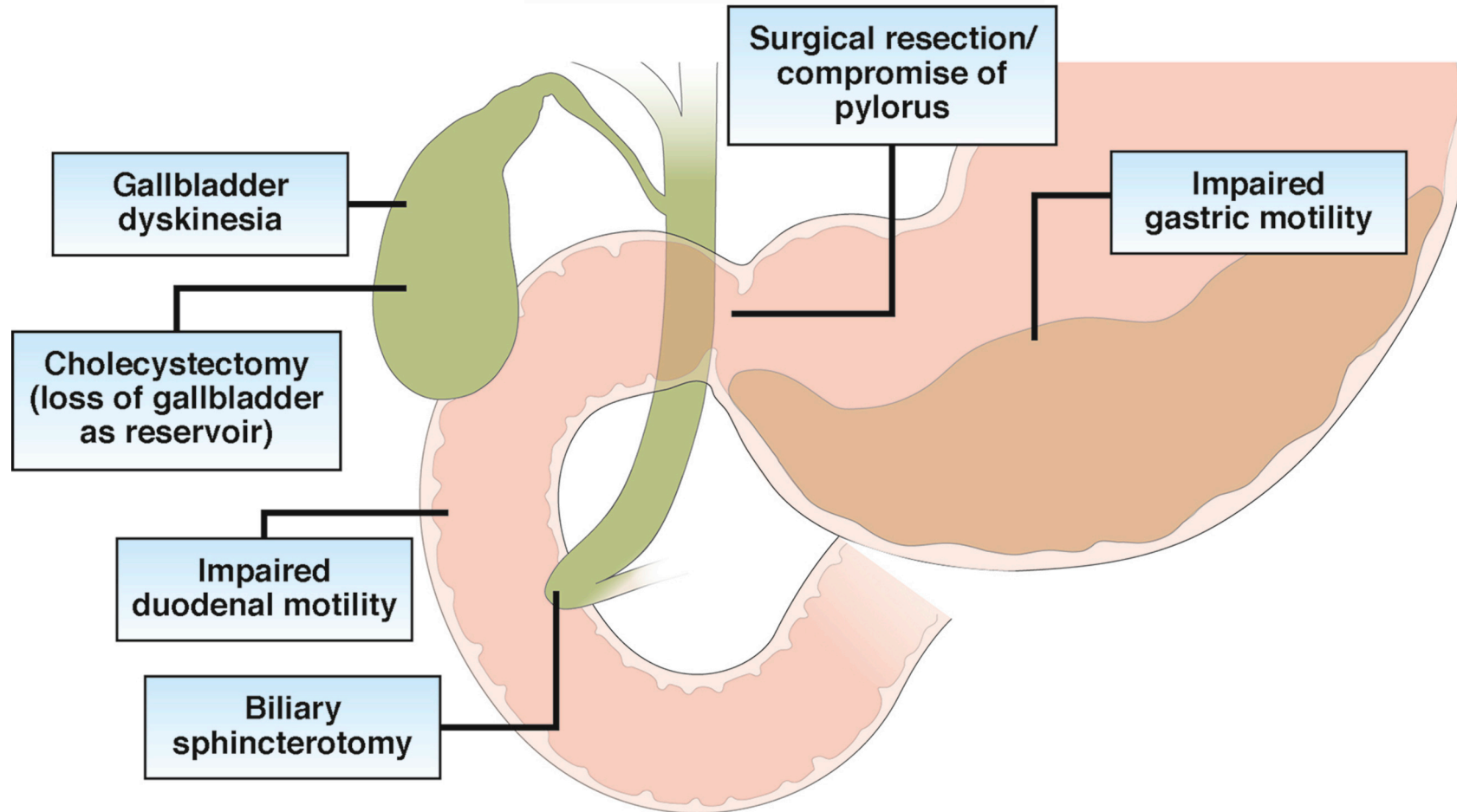
Findings	Grade				P value
		Grade 0	Grade 1	Grade 2	
Gastritis	Grade 0	180 (49.0%)	84 (23.0%)	12 (3.3%)	NS (0.6732)
	Grade 1	19 (5.0%)	8 (2.2%)	2 (0.5%)	
	Grade 2	20 (5.4%)	13 (3.6%)	2 (0.5%)	
	Grade 3	10 (2.7%)	4 (1.1%)	2 (0.5%)	
	Grade 4	0 (0.0%)	1 (0.3%)	0 (0.0%)	
Extent of gastritis	Grade 0	136 (77.7%)	0 (0.0%)	0 (0.0%)	<0.0001
	Grade 1	0 (0.0%)	60 (16.4%)	7 (1.9%)	
	Grade 2	0 (0.0%)	50 (13.7%)	10 (2.7%)	
	Grade 3	0 (0.0%)	0 (0.0%)	1 (0.3%)	
Bile reflux	Grade 0	237 (64.9%)	95 (26.0%)	15 (4.1%)	NS (0.6732)
	Grade 1	18 (4.9%)	15 (4.1%)	3 (0.8%)	

NS, Not significant

Kubo M et al Gastric Cancer (2002) 5: 83–89

In consideration of all of these difficulties in evaluating the functional state of the residual stomach by endoscopy alone, it seems that any attempt to determine the grade of gastritis with visual indices alone is an impossible







**Table 1 Initial reconstruction, interval, and location of remnant gastric cancer based on primary disease**

Ref.	Primary disease	No. of patients	Initial reconstruction (B- I /B-II/R-Y)	Interval (yr)	Location (Stomal/non-stomal)
Tanigawa <i>et al</i> <sup>[6]</sup> 2002	Benign	20	7/13	25.8	8/12
	Cancer	27	18/9	10.6	3/24
An <i>et al</i> <sup>[22]</sup> 2007	Benign	25	-	28.6	16/9
	Cancer	13	-	18.8	7/6
Ohashi <i>et al</i> <sup>[23]</sup> 2007	Cancer	108	71/28 <sup>1</sup>	7.5	14/94
Schaefer <i>et al</i> <sup>[24]</sup> 2007	Benign	19	1/18	34.0	11/8
Ahn <i>et al</i> <sup>[25]</sup> 2008	Benign	13	0/13	32.4	12/1
	Cancer	45	6/38 <sup>1</sup>	6.8	23/21
Firat <i>et al</i> <sup>[26]</sup> 2009	Benign	26	0/26	32.0	16/10
Ojima <i>et al</i> <sup>[27]</sup> 2010	Benign	17	12/5	22.0	8/9
	Cancer	21	16/5	9.0	2/19
Mezhir <i>et al</i> <sup>[3]</sup> 2011	Benign	105	B-II: 97	32.0	72/33
Komatsu <i>et al</i> <sup>[28]</sup> 2012	Benign	19	4/15	30.0	9/10
	Cancer	14	12/1 <sup>1</sup>	12.0	2/12
Li <i>et al</i> <sup>[29]</sup> 2013	Benign	88	28/60	32.1	55/33
	Cancer	24	14/10	16.8	9/15
Tokunaga <i>et al</i> <sup>[30]</sup> 2013	Benign	89	23/66	31.0	46/43
	Cancer	78	59/17 <sup>1</sup>	9.4	13/65
Leo <i>et al</i> <sup>[31]</sup> 2014	Benign	176	10/167	34.6	71/105



## Gastric stump cancer after distal gastrectomy for benign gastric ulcer in a population-based study

Jesper Lagergren<sup>1,2</sup>, Anna Lindam<sup>1</sup> and Robert M. Mason<sup>3</sup>

<sup>1</sup>Upper Gastrointestinal Research, Department of Molecular Medicine and Surgery, Karolinska Institutet, Stockholm, Sweden

<sup>2</sup>Division of Cancer Studies, King's College London, London, United Kingdom

<sup>3</sup>Department of Surgery, St. Thomas' Hospital, London, United Kingdom

**In conclusion, this large population based study revealed**

- **a prevalence 140/ 18912 (0.74%) of remnant gastric cancer**
- **an increased risk of cancer in the gastric remnant only 30 years or longer after gastric resection for benign disease, whereas other factors did not influence this risk.**

**Chronic damage attributed to duodenogastric reflux**

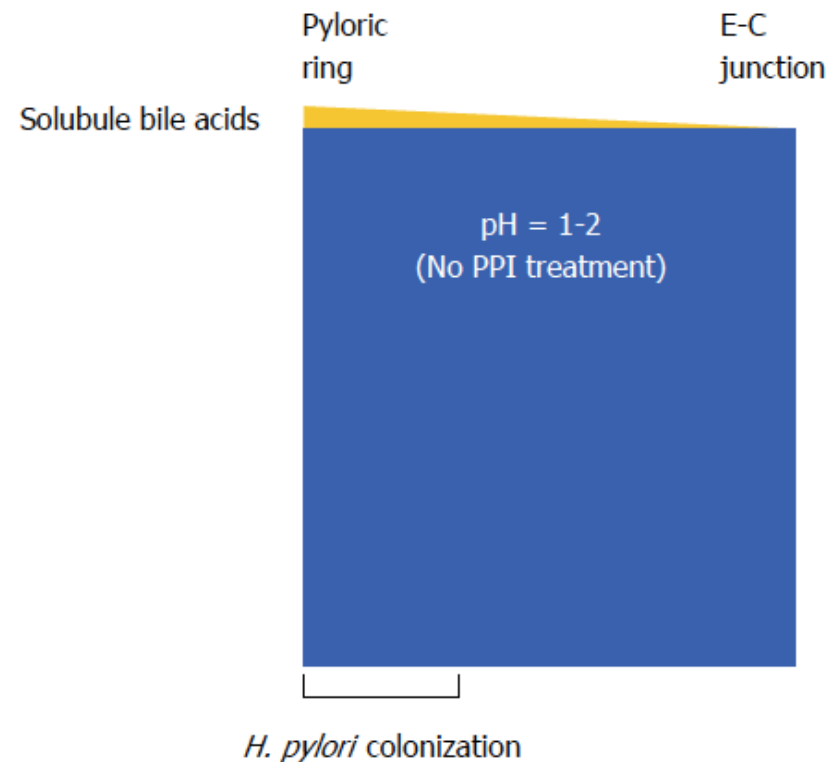
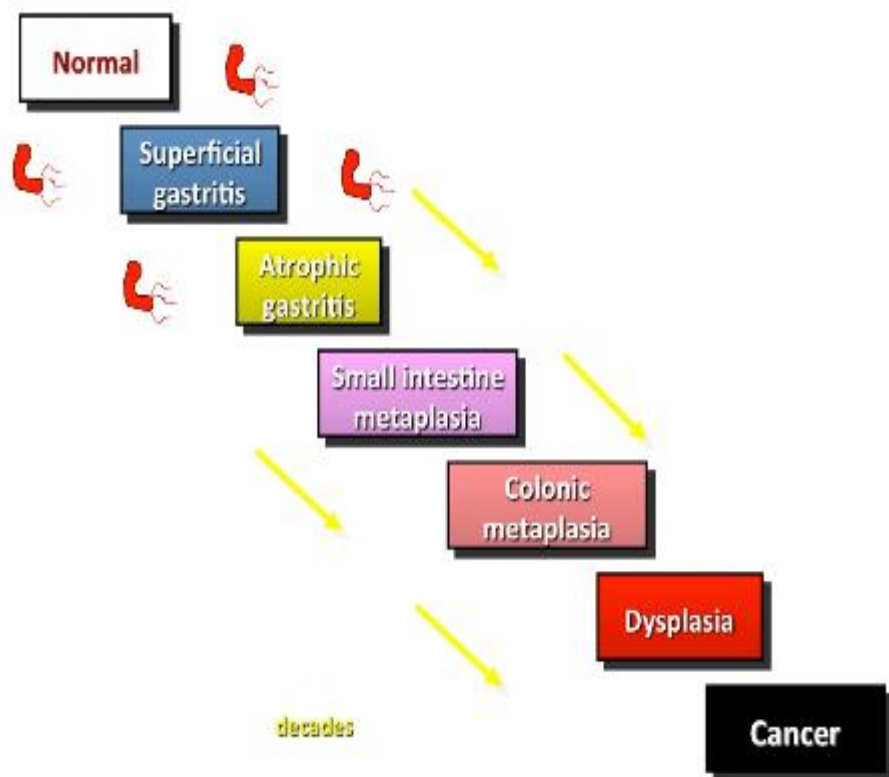


**is  
one of**

**the main factors responsible for  
changes affecting remnant gastric  
mucosa after distal gastrectomy**



Miwa K et al Carcinogenesis 1992;  
Kondo K et al Carcinogenesis 1995;  
Kondo K et al Gastric Cancer 2002;



Mukaisho et WJG 2014

© bile acids are chemorepellents for *H. pylori*

Worku ML et al J Med Microbiol 2004

© an inverse relationship exists between bile reflux and the presence of *H. pylori*, which may account for the absence of *H. pylori* in the stomach with persistent biliary reflux

Thao TD et al Biochem Pharmacol 2008



# Scandinavian Journal of Gastroenterology

2022, VOL. 57, NO. 12, 1430–1434

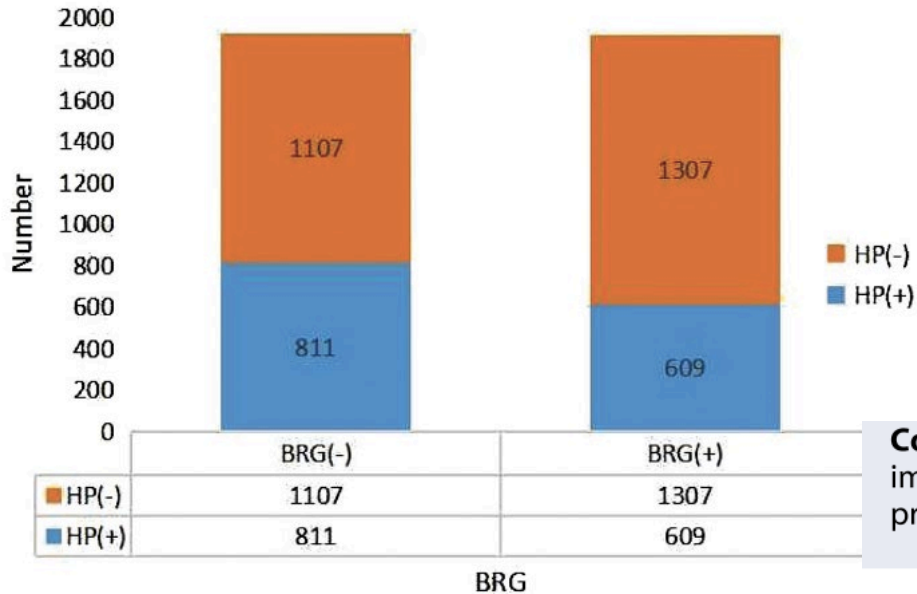
<https://doi.org/10.1080/00365521.2022.2094721>



ISSN: (Print) (Online) Journal homepage: [www.tandfonline.com/journals/igas20](http://www.tandfonline.com/journals/igas20)

## Negative correlations between bile reflux gastritis and *Helicobacter pylori* infection

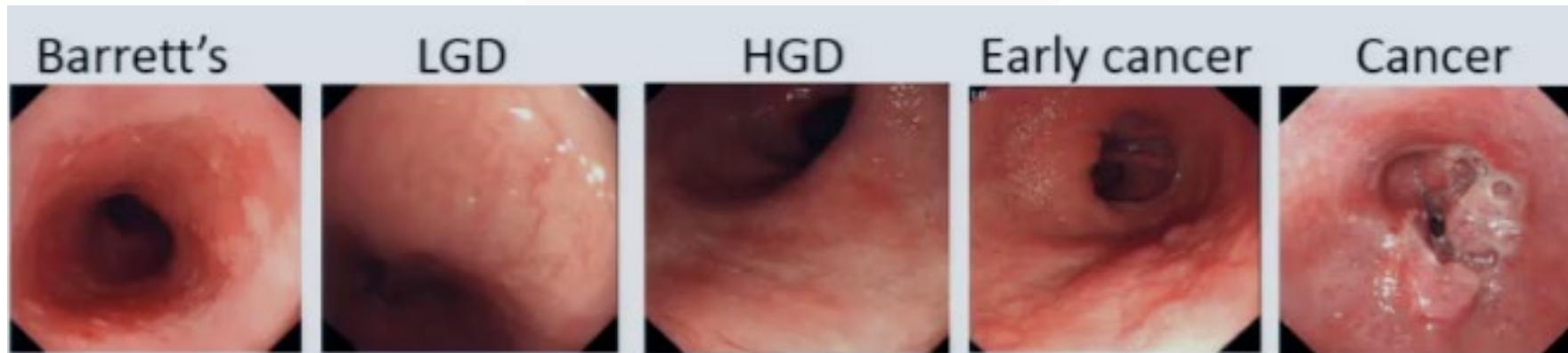
Xian-hua Zhuo, Jia-chen Sun, Wei-jie Zhong & Yi Lu

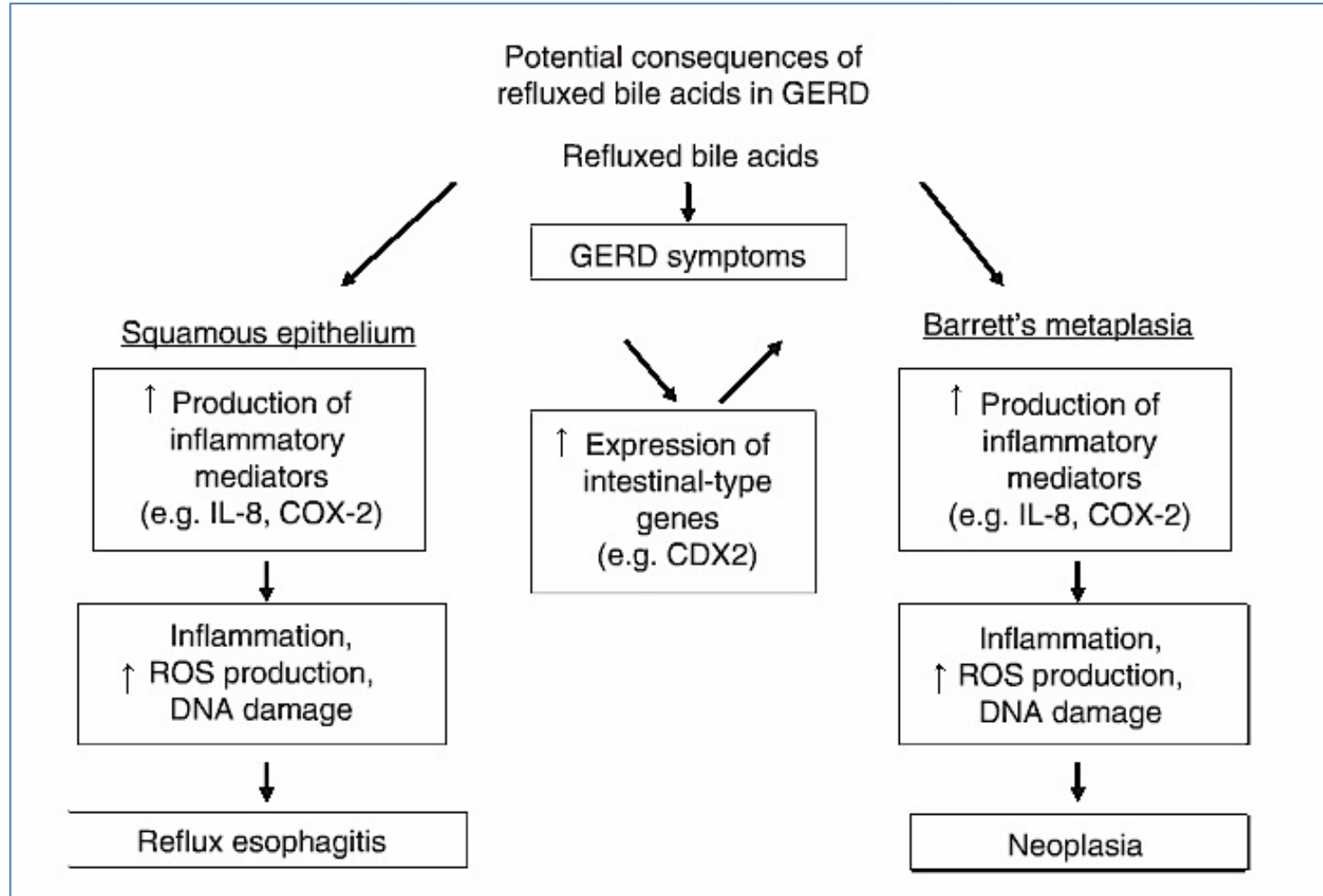


**Conclusion:** Patients with bile reflux may have less likely to get HP infection. HP eradication is an important thing for the prevention of gastric cancer and this study serves as a foundation and may provide directions for future research.



# Which is the impact of bile acids and bile salts on the esophageal mucosa and the development of Barrett's esophagus and cancer ?










# Which therapy?

## Systematic review: duodenogastroesophageal (biliary) reflux prevalence, symptoms, oesophageal lesions and treatment

Chamara Basnayake<sup>1,2</sup>  | Annelies Geeraerts<sup>1</sup> | Ans Pauwels<sup>1</sup> | Ger Koek<sup>3</sup> | Michael Vaezi<sup>4</sup> | Tim Vanuytsel<sup>1</sup>  | Jan Tack<sup>1</sup> 

- **PPIs** 9 studies
- **Prokinetic agents** 2 studies
- **Histamine receptor antagonist** 1 study
- **Baclofen** 1 study

10 studies Bilitec 2000 as the device to measure bile reflux,  
3 studies used a sodium ion electrode.

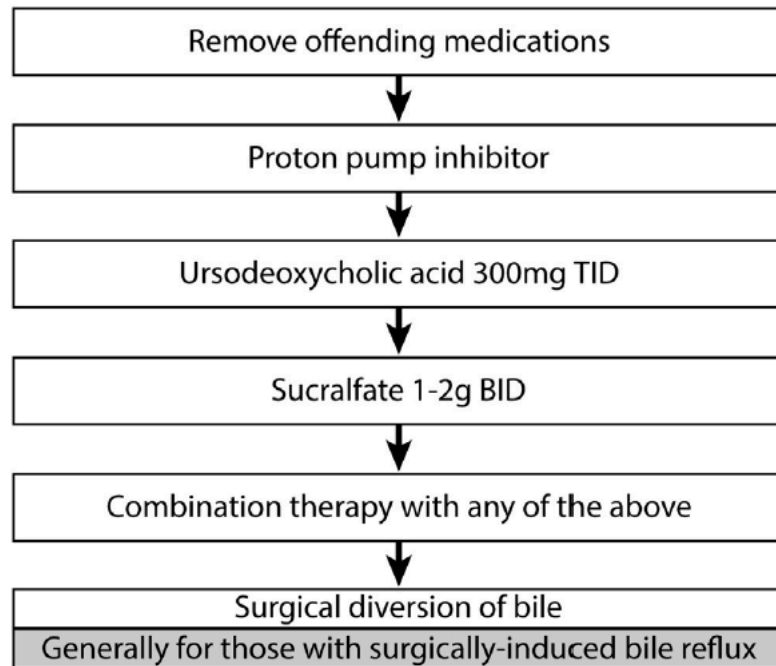


**Only 1 study utilised a randomised controlled trial design**

2021



# Which therapy?



**Figure 5.** Proposed treatment algorithm.



# Management advice for patients with reflux-like symptoms: an evidence-based consensus

Hungin, A. Pali<sup>a</sup>; Yadlapati, Rena<sup>b</sup>; Anastasiou, Foteini<sup>c</sup>; Bredenoord, Albert J.<sup>d</sup>; El Serag, Hashem<sup>e</sup>; Fracasso, Pierluigi<sup>f</sup>; Mendive, Juan M<sup>g</sup>; Savarino, Edoardo V.<sup>h</sup>; Sifrim, Daniel<sup>i</sup>; Udrescu, Mihaela<sup>j</sup>; Kahrilas, Peter J<sup>k</sup>

Author Information 

*European Journal of Gastroenterology & Hepatology* 36(1):p 13-25, January 2024. | DOI: 10.1097/MEG.0000000000002682 

**Statement 18:** Alginate-antacid combinations are an effective treatment for reflux-like symptoms. Agreement: 100% (6, 90.9%; 5, 9.1%; grade of evidence: A; strength of recommendation: high).

**Statement 21:** Products containing hyaluronic acid and chondroitin sulphate are an effective treatment option for reflux-like symptoms. Agreement: 81.8% (6, 82%; 5, 18%; grade of evidence: B; strength of recommendation: low).





# PROTEZIONE DELLA MUCOSA: NUOVE EVIDENZE



**AJG** The American Journal of GASTROENTEROLOGY

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ARTICLE: FUNCTIONAL GI DISORDERS

**Poliprotect vs Omeprazole in the Relief of Heartburn, Epigastric Pain, and Burning in Patients Without Erosive Esophagitis and Gastroduodenal Lesions: A Randomized, Controlled Trial**

Corazziari, Enrico Stefano MD<sup>1</sup>; Gasbarrini, Antonio MD<sup>2</sup>; D'Alba, Lucia MD<sup>3</sup>; D'Ovidio, Valeria MD<sup>4</sup>; Riggio, Oliviero MD<sup>5</sup>; Passaretti, Sandro MD<sup>6</sup>; Annibale, Bruno MD<sup>7</sup>; Cicala, Michele MD<sup>8</sup>; Repici, Alessandro MD<sup>9</sup>; Bassotti, Gabrio MD<sup>10</sup>; Ciacci, Carolina MD<sup>11</sup>; Di Sabatino, Antonio MD<sup>12</sup>; Neri, Matteo MD<sup>13</sup>; Bragazzi, Maria Consiglia MD<sup>14</sup>; Ribichini, Emanuela MD<sup>6</sup>; Radocchia, Giulia MA Biothec<sup>15</sup>; Iovino, Paola MD<sup>11</sup>; Marazzato, Massimiliano PhD<sup>15</sup>; Schippa, Serena Msc Biol<sup>15</sup>; Badiali, Danilo MD<sup>5</sup>

Author Information

*The American Journal of Gastroenterology* ( ):10.14309/ajg.0000000000002360, September 11, 2023. | DOI: 10.14309/ajg.0000000000002360

prodotto 100 % naturale costituito da Poliprotect e da una frazione flavonoidica da Glycyrrhiza glabra and Matricaria recutita)

Poliprotect è costituito da:

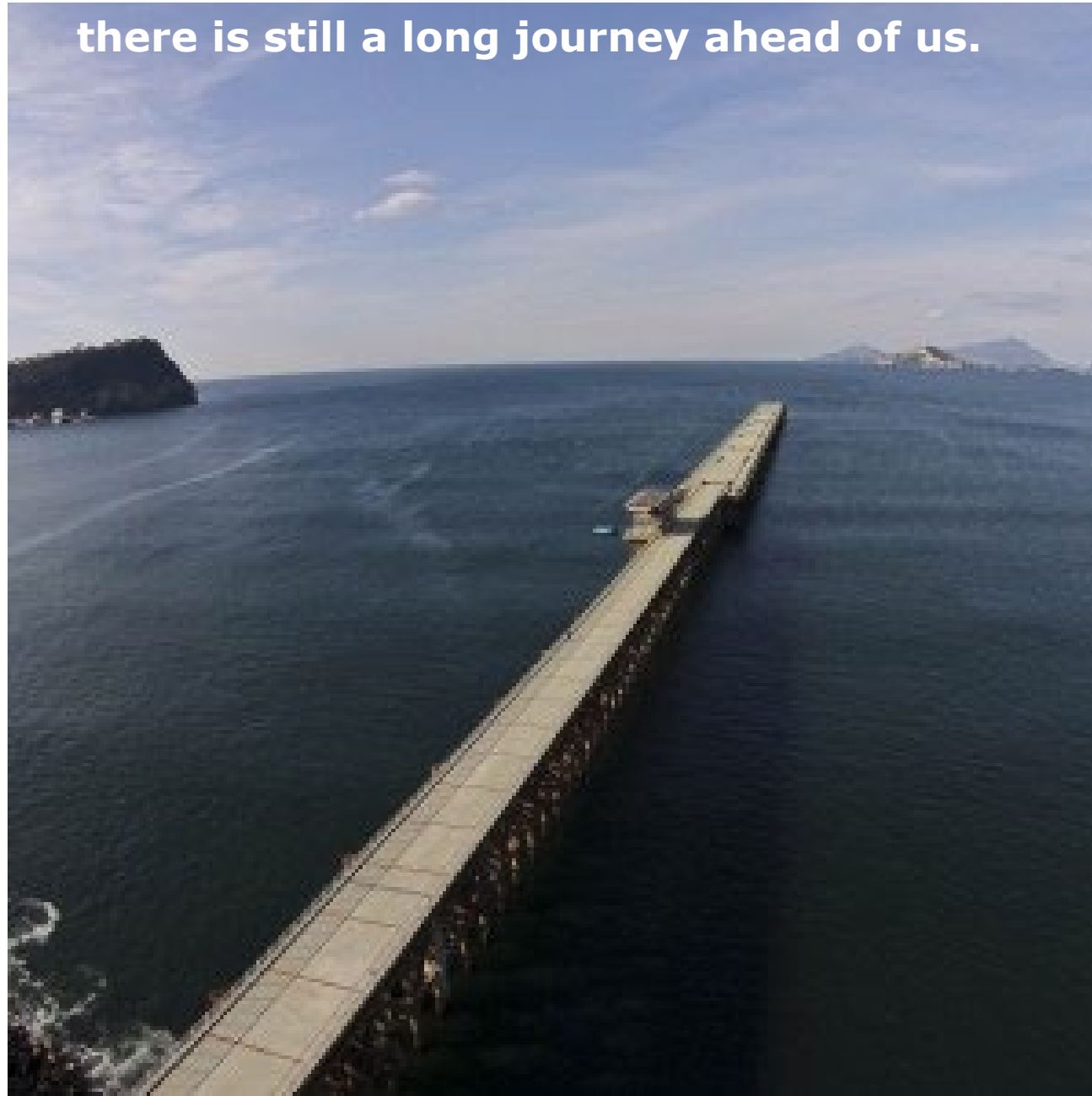
- Frazione polisaccaridica aderente all'epitelio da *Aloe vera*, *Malva sylvestris* e *Althaea officinalis*  
**per rafforzare la barriera epiteliale**
- Componenti antiacidi dai minerali naturali limestone e nahcolite incorporati nella frazione polisaccaridica  
**per tamponare l'acido sull'epitelio al quale aderisce**

# Is biliary reflux increased after OAGB?





there is still a long journey ahead of us.







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**Grazie**