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**SECOND ANNUAL
GI & LIVER**

Summit



What's New in GI Pharmacotherapy?

Philip Schoenfeld, MD, MSc, MEd, MSc (Epi)

Editor-in-Chief, Evidence-Based GI: An ACG Publication

Chief (Emeritus)-Gastroenterology Section

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Detroit, MI



EVIDENCE-BASED GI
AN ACG PUBLICATION

*Clinical take-aways and
evidence-based summaries of
articles in GI, Hepatology & Endoscopy*



AMERICAN COLLEGE OF GASTROENTEROLOGY





EVIDENCE-BASED GI | Clinical take-aways and
AN ACG PUBLICATION | evidence-based summaries of
articles in GI, Hepatology & Endoscopy



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Upadacitinib Is Effective for the Induction and Maintenance of Moderate-to-Severe Crohn's Disease

Rahul Dalal, MD, MPH; Jessica Allegretti, MD, MPH

Listen to the audio summary

In two 12-week, double-blind, placebo-controlled randomized control trials of moderate-to-severe Crohn's disease patients, upadacitinib 45 mg daily was more effective than placebo at inducing clinical remission: 50% vs 29% in U-EXCEL, and 39% vs 21% in U-EXCEED. In a 52-week, double-blind, placebo-controlled randomized control trial, upadacitinib 30 mg and upadacitinib 15 mg daily was more effective than placebo at maintaining clinical remission: 48% vs 37% vs 15%, respectively.

Summarizing Loftus EV Jr, Panés J, Lacerda AP, Peyrin-Biroulet L, et al. Upadacitinib Induction and Maintenance Therapy for Crohn's Disease. *N Engl J Med.* 2023 May 25;388(21):1966-1980. doi:



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Poor Bowel Preps: A Daily Frustration!



It's a Bad "Prep" Even Though the Patient Took It Correctly: Consider 15 mg Bisacodyl plus 4-Liter PEG Split Prep Before Next Colonoscopy



Philip Schoenfeld, MD, MSc (Epi)

Chief (Emeritus)-Gastroenterology Section, John D. Dingell VA Medical Center, Detroit, MI

This article reviews Sey MSL, Von Renteln D, Sultanian R, et al. A Multicenter Randomized Controlled Trial Comparing Bowel Cleansing Regimens for Colonoscopy After Failed Bowel Preparation. Clin Gastroenterol Hepatol 2022; In Press.

Philip Schoenfeld, MD, MSc (Epi)
Editor-in-Chief

Tweetorial provided by:

Zubair Khan, MD

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PGY-6, University of Texas
at Houston



Why Do Patients Have Bad Preps Even When They Are Compliant With Instructions and Complete the Prep?



❖ Risk Factors for colonic dysmotility and inadequate bowel preparation despite compliance

- Obesity
- Current opioid use
- Diabetes mellitus
- History of using constipation treatments
- Current use of anticholinergics (including TCA)
- Prior history of inadequate prep!

Why Do Patients Have Bad Preps Even When They Are Compliant With Instructions and Complete the Prep?



❖ Risk Factors for colonic dysmotility and inadequate bowel preparation despite compliance

- Obesity
- Current opioid use
- Diabetes mellitus
- History of using constipation treatments
- Current use of anticholinergics (including TCA)
- Prior history of inadequate prep!

Note: In non-compliant patient, additional patient education is probably more beneficial than prescribing supra-therapeutic bowel prep regimen.



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- No prior RCT assessing patients who successfully completed 4L PEG split-prep but still had inadequate cleansing.
- Multi-center, single-blind RCT
- Intervention: 4L PEG split prep + 15mg bisacodyl (taken at 2pm on day before scope) vs 6L PEG split prep + 15mg bisacodyl
- Outcome: Adequate bowel prep based on BBPS ≥ 6 with ≥ 2 in each segment
- Patient Demographics: 37% obese, 41% with IBS-C or CIC, 10% on opioids. Prior bowel prep: 35% used 4L PEG; 38% used 2L PEG; 12% used sodium picosulfate



It's a Bad "Prep" Even Though the Patient Took It Correctly: Consider 15 mg Bisacodyl plus 4-Liter PEG Split Prep Before Next Colonoscopy



Outcome		Split-dose 4L + bisacodyl (n=97)	Split-dose 6L + bisacodyl (n=99)	P-value
Adequate cleansing	Defined as BBPS ≥ 6	83 (91.2%)	78 (87.6%)	0.44
	Defined as adequate to identify polyps $>5\text{mm}$	82 (91.1%)	76 (85.4%)	0.24
Secondary endpoints	Cecal intubation rate, n (%)	87 (98.7%)	82 (92.1%)	0.19
	Adenoma detection rate, n (%)	34 (37.4%)	28 (31.5%)	0.41
Adherence	Diet + consumed 100% of prep	67 (81.7%)	53 (68.0%)	0.05
	Diet + consumed 80% of prep	71 (86.6%)	57 (73.1%)	0.03

My Practice:

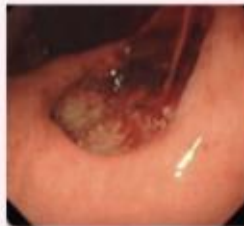


- For patients with history of poor bowel preparation or 2 risk factors for poor prep >>>>
- Prescribe 4 liters PEG split-prep + 15 mg bisacodyl at 2pm on day before procedure.

Forrest 1a, 1b, and IIa Bleeding Peptic Ulcers: Best Practices to Control Bleeding and Reduce Re-Bleeding



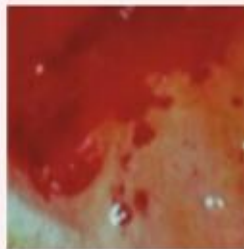
Acute Hemorrhage



1a

Active Spurting

Rebleeding Risk:
60 to 100%



1b

Active Oozing

Rebleeding Risk:
50%

Signs of Recent Hemorrhage



IIa

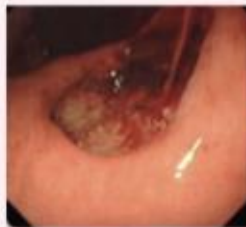
**Non-Bleeding
Visible Vessel**

Rebleeding Risk:
40 to 50%

Forrest 1a, 1b, and IIa Bleeding Peptic Ulcers: Best Practices to Control Bleeding and Reduce Re-Bleeding

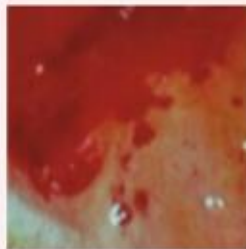


Acute Hemorrhage



1a

Active Spurting
Rebleeding Risk:
60 to 100%



1b

Active Oozing
Rebleeding Risk:
50%

Signs of Recent Hemorrhage



IIa

**Non-Bleeding
Visible Vessel**
Rebleeding Risk:
40 to 50%

These are re-bleeding risks without endoscopic intervention + PPIs. However, even with conventional therapy, failure to control bleeding or re-bleeding is 10-20%!

Over-the-Scope Clips Decrease Non-Variceal Upper GI Bleeding vs Standard Endoscopic Treatment... In the Right Patient



Jennifer M. Kolb MD, MS

Assistant Professor of Medicine, Division of Gastroenterology, Hepatology and Parenteral Nutrition, VA Greater Los Angeles Healthcare System, David Geffen School of Medicine at UCLA, Los Angeles, CA

This summary reviews Lau JYW, Li R, Tan C et al. Comparison of Over-the-Scope Clips to Standard Endoscopic Treatment as the Initial Treatment in Patients With Bleeding From a Nonvariceal Upper Gastrointestinal Cause: A Randomized Controlled Trial. *Ann Intern Med* 2023 Apr;176(4):455-462. doi: 10.7326/M22-1783

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Drs Kolb, Chauhan and Ezeani have no disclosures

Tweetorial Provided by:

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&

Kashyap Chauhan

 **@DrKashyapC**

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Over-The-Scope-Clip Highlights



- Nitinol clips mounted on a clear plastic cap
- Attached to the endoscope externally
- Target tissue suctioned into cap
- Deployed clip results in 8 opposing prongs to approximate anchoring into mucosal tissue

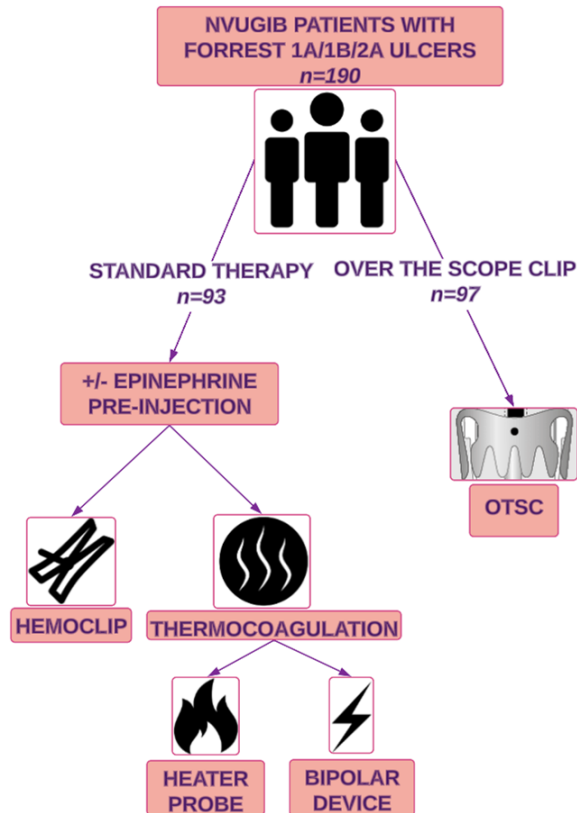
“Over-the-Scope clips decrease non-variceal upper GI bleed vs Standard Endoscopic Treatment... In the right Patient”
Summary of Lau JY, Tan C, Sun X, et al. Comparison of Over-the-Scope Clips to standard Endoscopic Treatment as the initial Treatment in patients with Bleeding from a Nonvariceal Upper Gastrointestinal Cause. Ann Intern. Med. 2023; 176(4):455-462

IMPORTANCE

10-20% Risk of Re-Bleed in High-Risk Lesions with Conventional Treatment

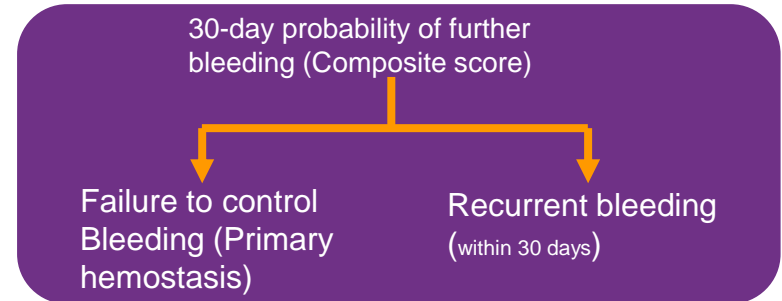
10-20% risk of re-bleed in high-risk lesions with conventional treatment

May affirm OTSC as new first line therapy for NVUGIB (vs. currently only after treatment failure).



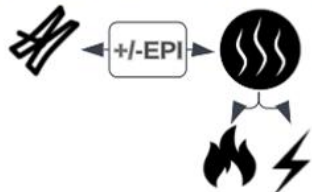
**Primary
Endpoint**

“Over-the-Scope clips decrease non-variceal upper GI bleed vs Standard Endoscopic Treatment... In the right Patient”
Summary of Lau JYW, Tan C, Sun X, et al. Comparison of Over-the-Scope Clips to standard Endoscopic Treatment as the initial Treatment in patients with Bleeding from a Nonvariceal Upper Gastrointestinal Cause. *Ann Intern. Med.* 2023; 176(4):455-462





STANDARD THERAPY



VS.

OTSC



STUDY RESULTS

14.6%	FURTHER BLEEDING 95% CI 3.3-20.0, $P=0.006$	3.2%
6.2%	PRIMARY HEMOSTASIS FAILURE 95% CI: 0.7-11.8	1.1%
8.8%	30-DAY RECURRENT BLEED 95% CI: -0.3 to 14.4	2.2%

“Over-the-Scope clips decrease non-variceal upper GI bleed vs Standard Endoscopic Treatment... In the right Patient”
Summary of Lau JY, Tan C, Sun X, et al. Comparison of Over-the-Scope Clips to standard Endoscopic Treatment as the initial Treatment in patients with Bleeding from a Nonvariceal Upper Gastrointestinal Cause. Ann Intern. Med. 2023; 176(4):455-462

My Practice:



- Worked with advanced endoscopists and endoscopy technicians to learn how to mount OTSC clip and practice placement.

Treating *Helicobacter pylori* Infection With Vonoprazan, A Potassium-Competitive Acid Blocker: A New Paradigm



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Chief Emeritus-Gastroenterology Section, John D. Dingell VA Medical Center, Detroit, Michigan

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Romy Chamoun, MD

 @RomyChamoun

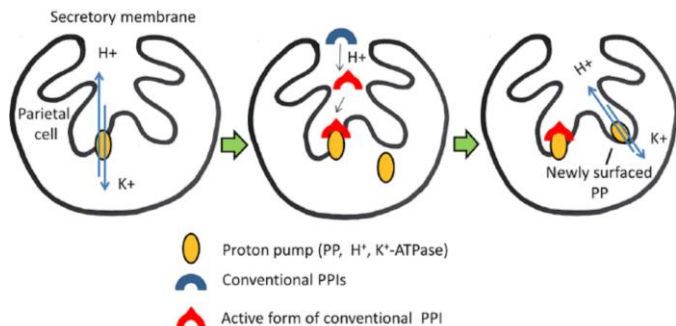
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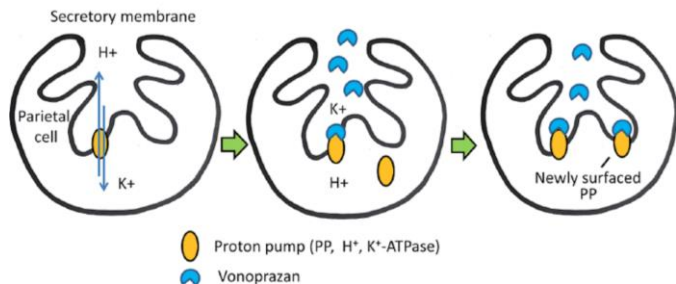




(a) Conventional PPI



(b) Vonoprazan



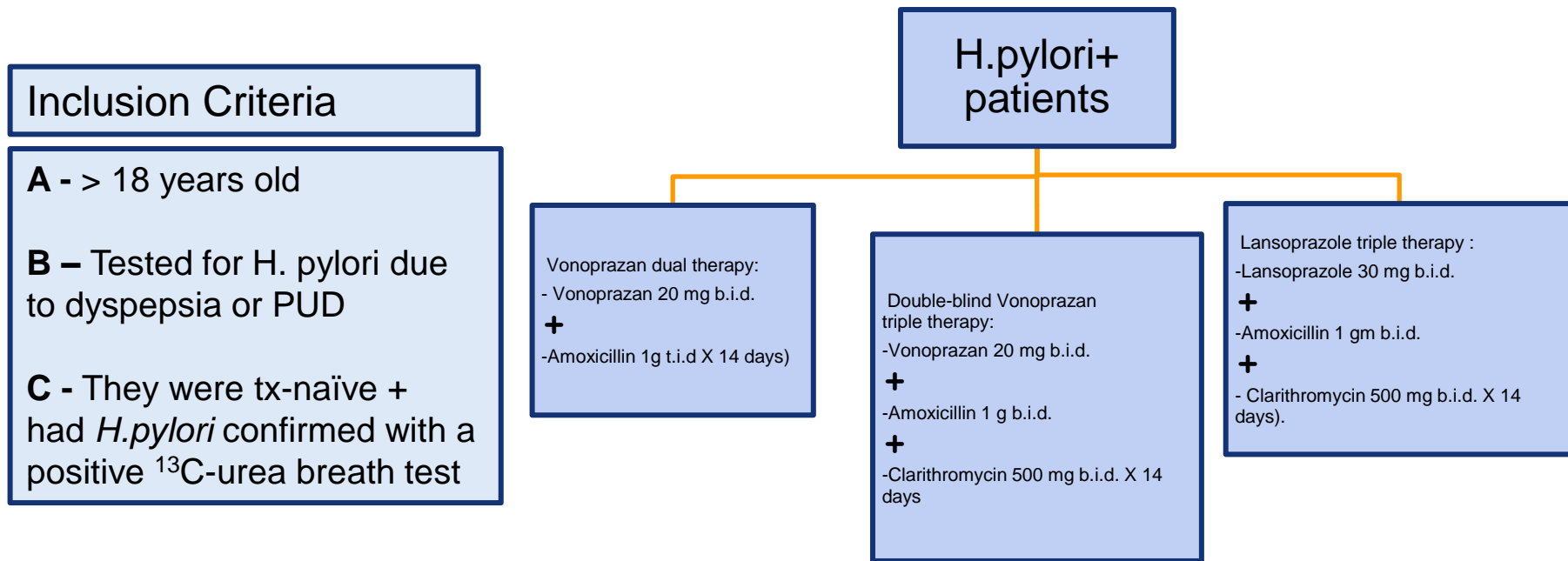
Conventional PPIs are

- Unstable in canaliculi
- Rapidly degraded
- Not able to inhibit new proton pumps (PPs) that surface after administration of the drug.

→ require a few days to reach their maximum effect

Vonoprazan, a potassium-competitive acid blocker acts differently:

- ✓ does not require acid activation
 - ✓ rapidly absorbed in the small intestine
 - ✓ binding to H⁺/K⁺-ATPase in a K⁺-competitive manner
 - ✓ more stable than conventional PPIs in the canaliculi
- fast and stable inhibition of gastric acid secretion





US and European phase 3 RCT comparing vonoprazan- and lansoprazole-based regimens

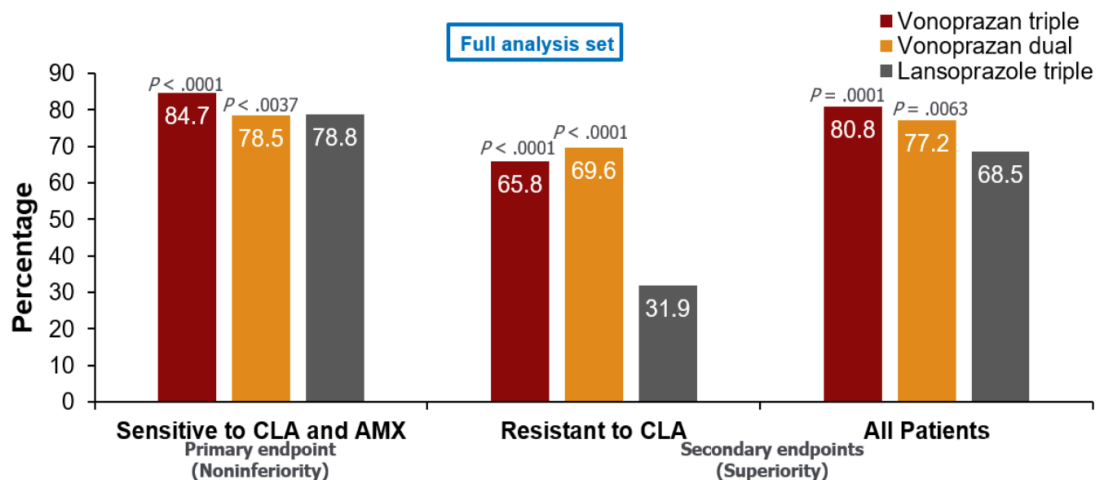


Figure 1: *Helicobacter pylori* eradication rates.

AMX, amoxicillin ; CLA, clarithromycin; RCT, randomized controlled trial.



US and European phase 3 RCT comparing vonoprazan- and lansoprazole-based regimens

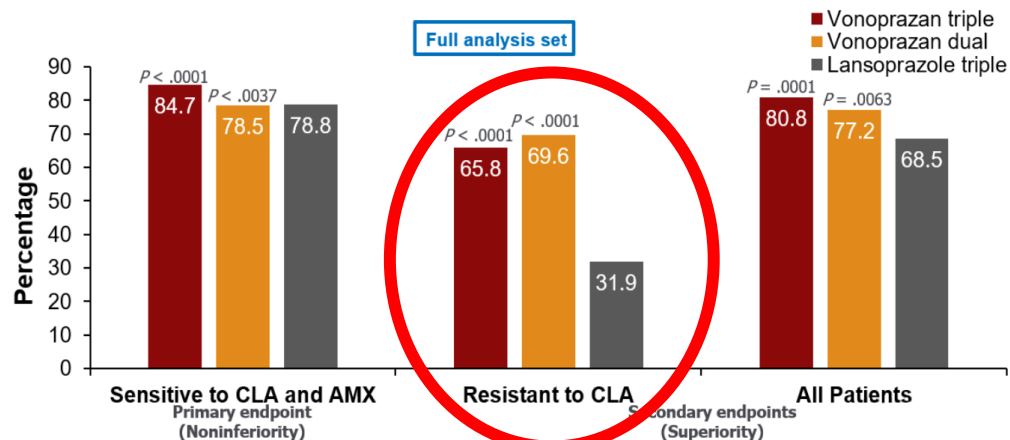


Figure 1: *Helicobacter pylori* eradication rates.

AMX, amoxicillin ; CLA, clarithromycin; RCT, randomized controlled trial.

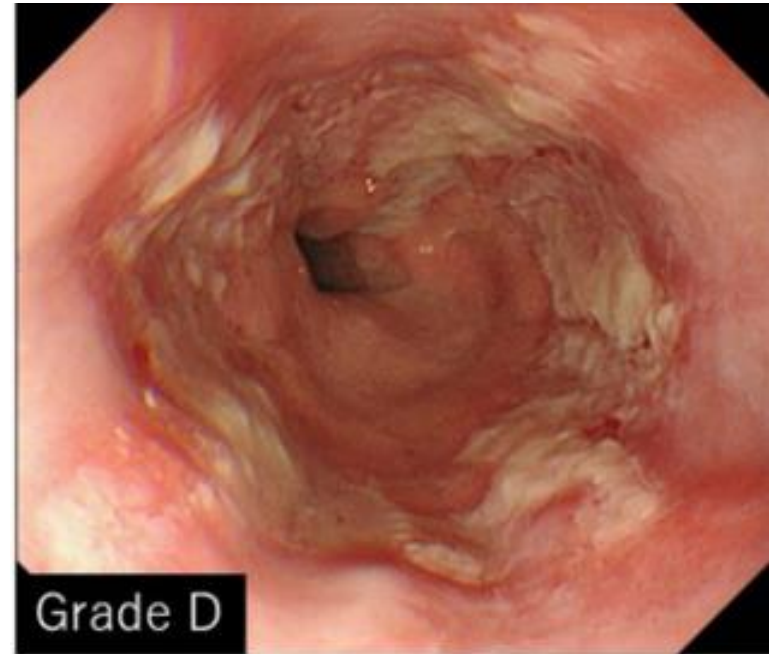
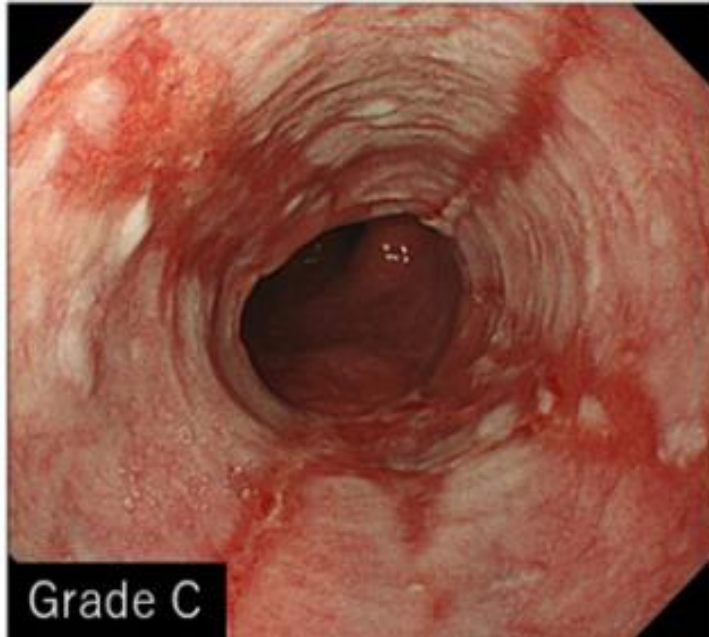
If clarithromycin resistance is > 15-20% in your region or if your patient previously tx with “-mycins” antibiotic, use vonoprazan-based regimen

My Practice:



- Worked I don't use clarithromycin-based h.pylori treatment regimens. Currently, I rely on bismuth-based quadruple regimens.
- When vonoprazan-based regimens become available in next 6 months, will use vonoprazan 20mg bid + 1000mg amoxicillin tid X 14 days as my primary regimen.

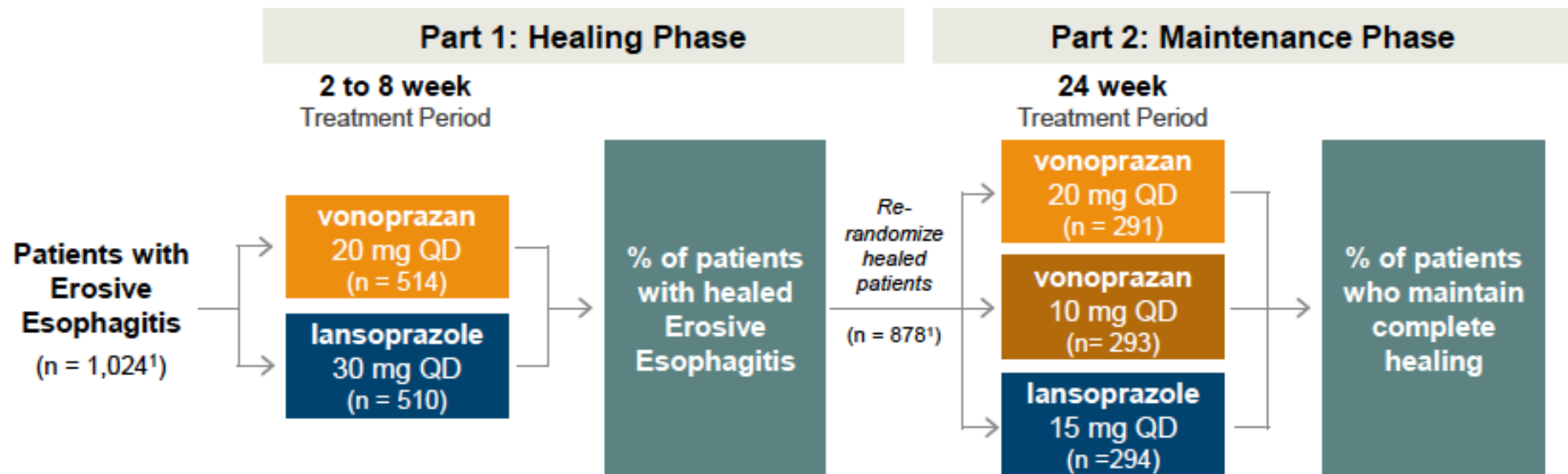
LA Class C & D Erosive Esophagitis: Best Practices for Healing & Maintenance of Healing



PHALCON-EE phase 3 study design

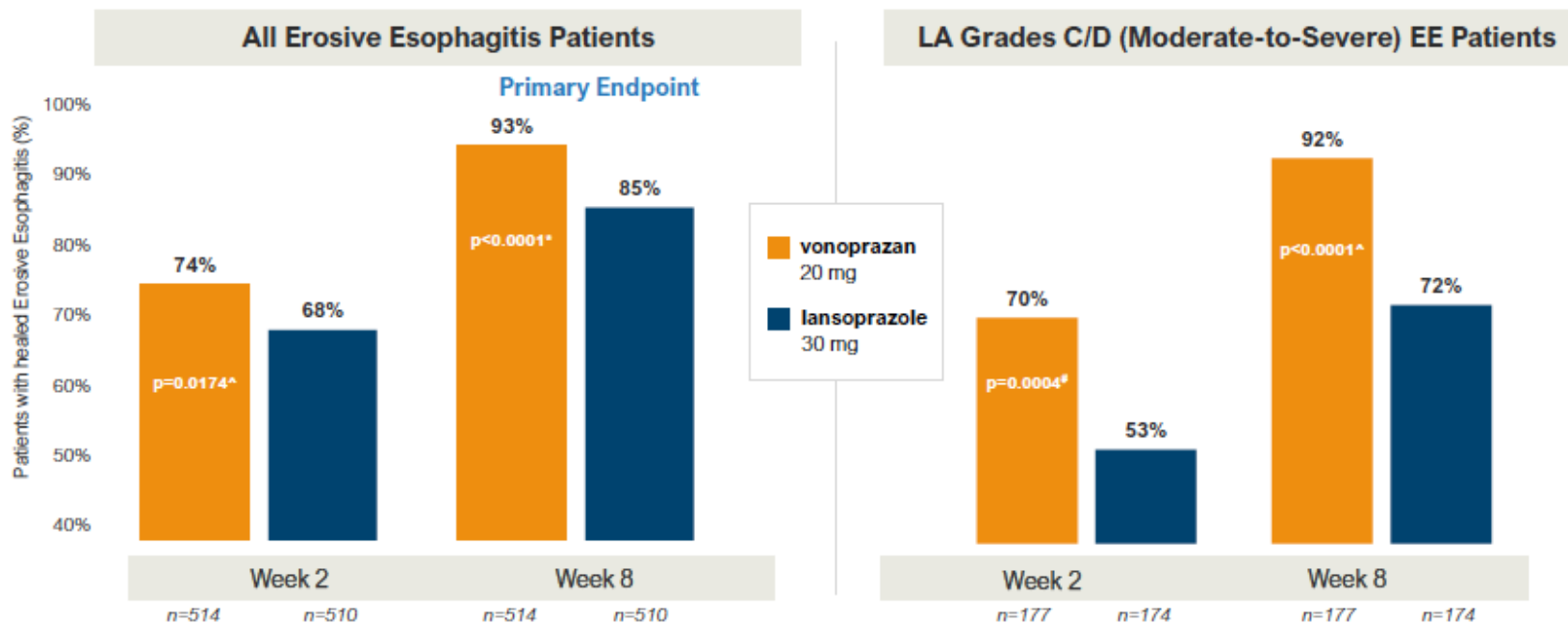
US/Europe study in Erosive Esophagitis

pHalcon^{ee}
A research study for Erosive Esophagitis



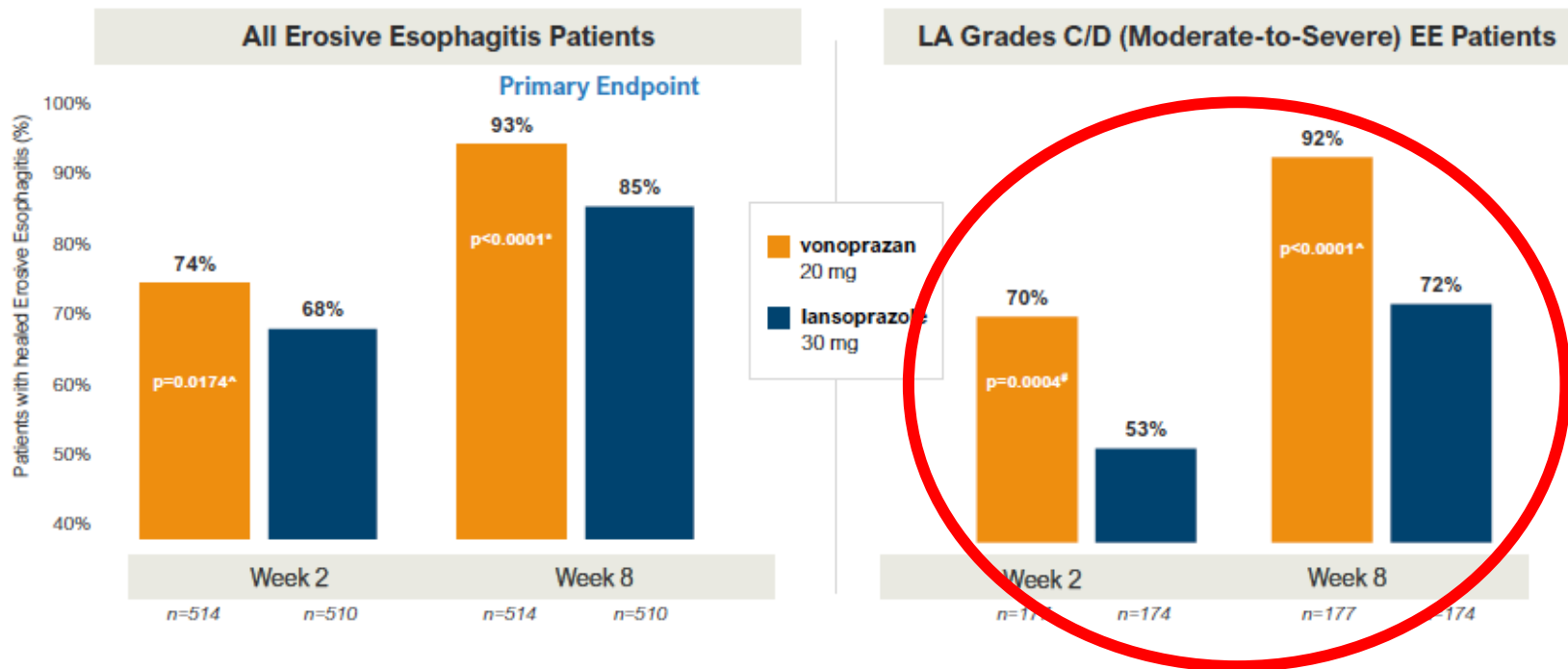


Healing endpoints



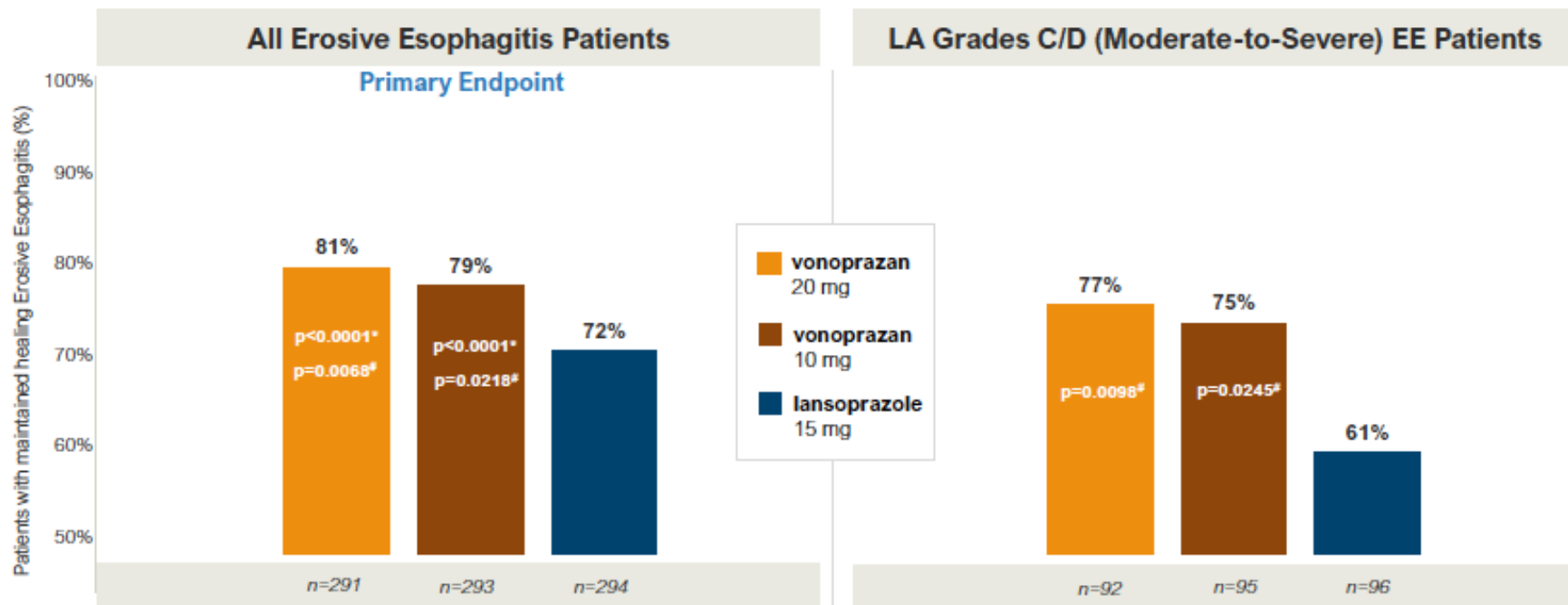


Healing endpoints





Maintenance of healing endpoints



Maintenance of healing endpoints

