

**FIRENZE 29 NOVEMBRE 1 DICEMBRE 2018**

# **Histologic Scoring Indices In Ulcerative Colitis: A Critical Reappraisal**



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Anatomia Patologica  
Spedali Civili  
Brescia**



**“Terms such as non-specific chronic inflammation or signs of chronic inflammatory bowel disease but non-diagnostic should be avoided”**

**“It is important to define the terminology as clearly as possible and to use the terminology consistently to avoid confusion”**



# **CLINICAL INFORMATIONS**

## **MULTIPLE BIOPSIES**

**from five sites around the colon  
(including the rectum and the ileum)**

**Multiple implies a minimum of  
two samples....correctly oriented!!!!**



## Microscopic Features of Ulcerative Colitis

A diagnosis of established ulcerative colitis is based upon the combination of :

**A) Crypt**

**Arch**

**Dis**

**B) Heavy Diffuse**

**Trans**

**Prop**

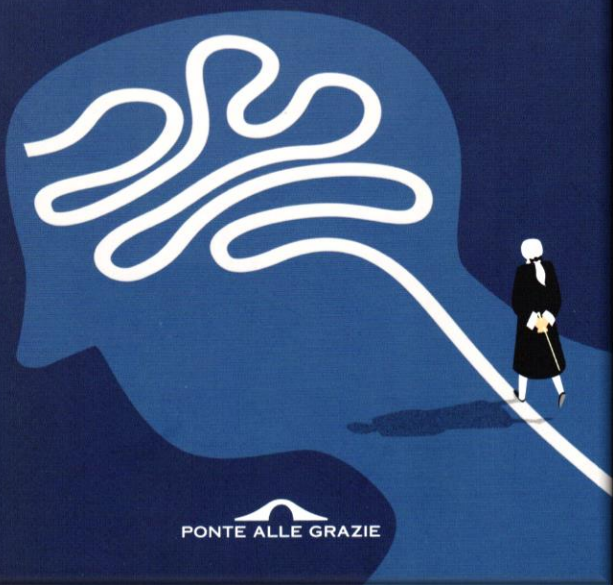
**C) Basal  
Plasmacytosis**



ROGER-POL DROIT

# La passeggiata di Kant

*Filosofia del camminare in 27 ritratti*



**Ah, tu credi questo, ma ne sei proprio sicuro?**

**Questa insinuazione è una vera e propria destabilizzazione.**

**La messa in dubbio che incrina le evidenze dà sulle prime, l'impressione che si cadrà.**

**D'un tratto, il mondo familiare e sicuro si rivela incerto.**

**I pilastri ai quali ci aggrappavamo ci sfuggono.**

**Credete di sapere? Embè, non sapete!**

**Siete sicuri che tutto è familiare?**

**Guardate in modo diverso, sarete sorpresi .**

## Microscopic Features of Ulcerative Colitis

**A) Heavy Diffuse**

**B)**

**D) Crypt**

**Architectural Distortion**





Heavy Diffuse Transmucosal Lamina Propria Cell Increase

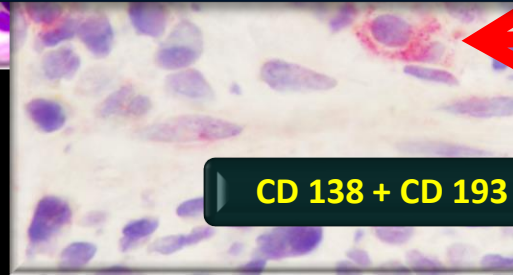



Basal Plasmacytosis

Plasmacells

Eosinophils

# DIAGNOSIS OF ULCERATIVE COLITIS !!!

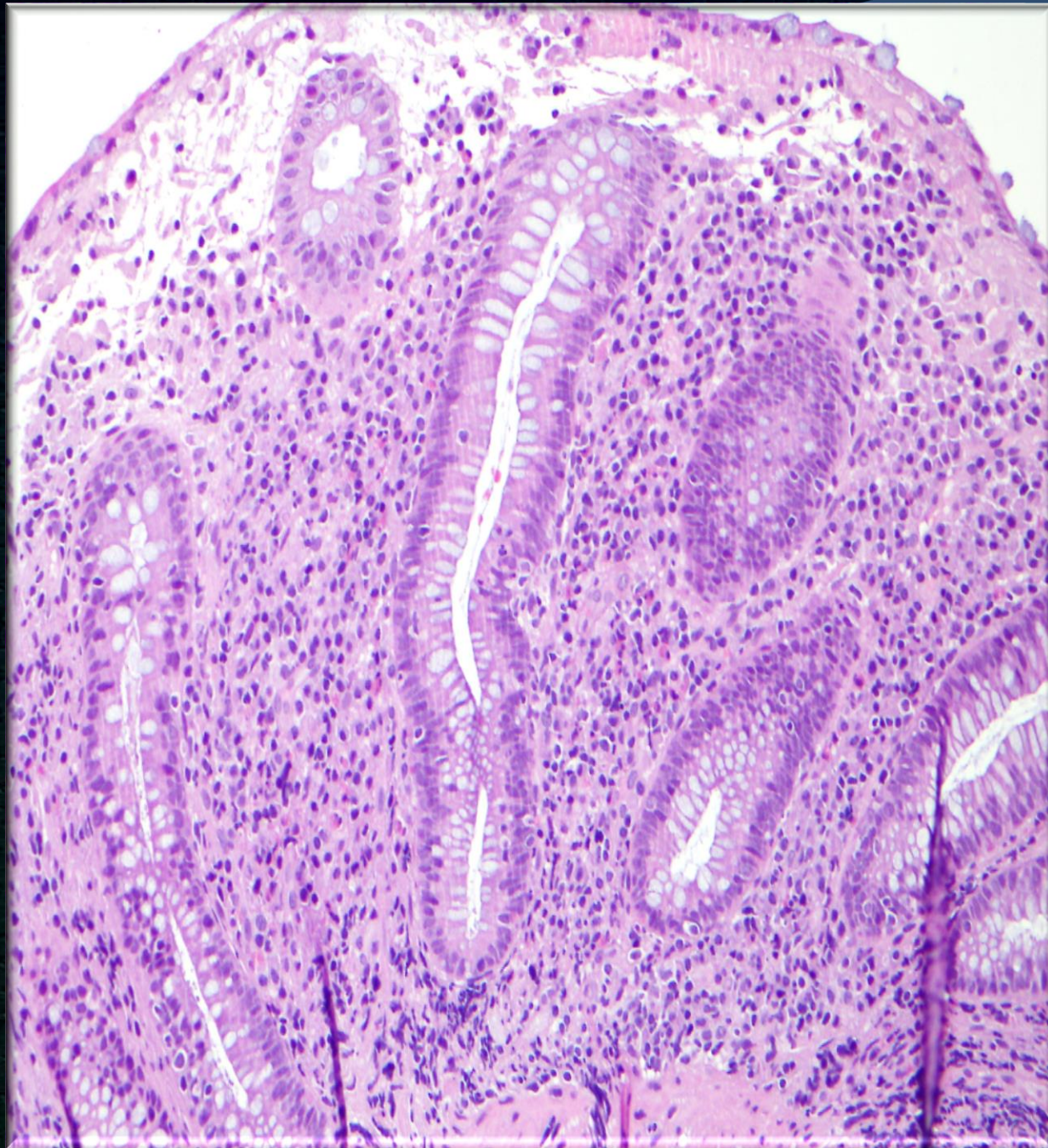


CD 138 + CD 193

Crypt Architectural Distortion

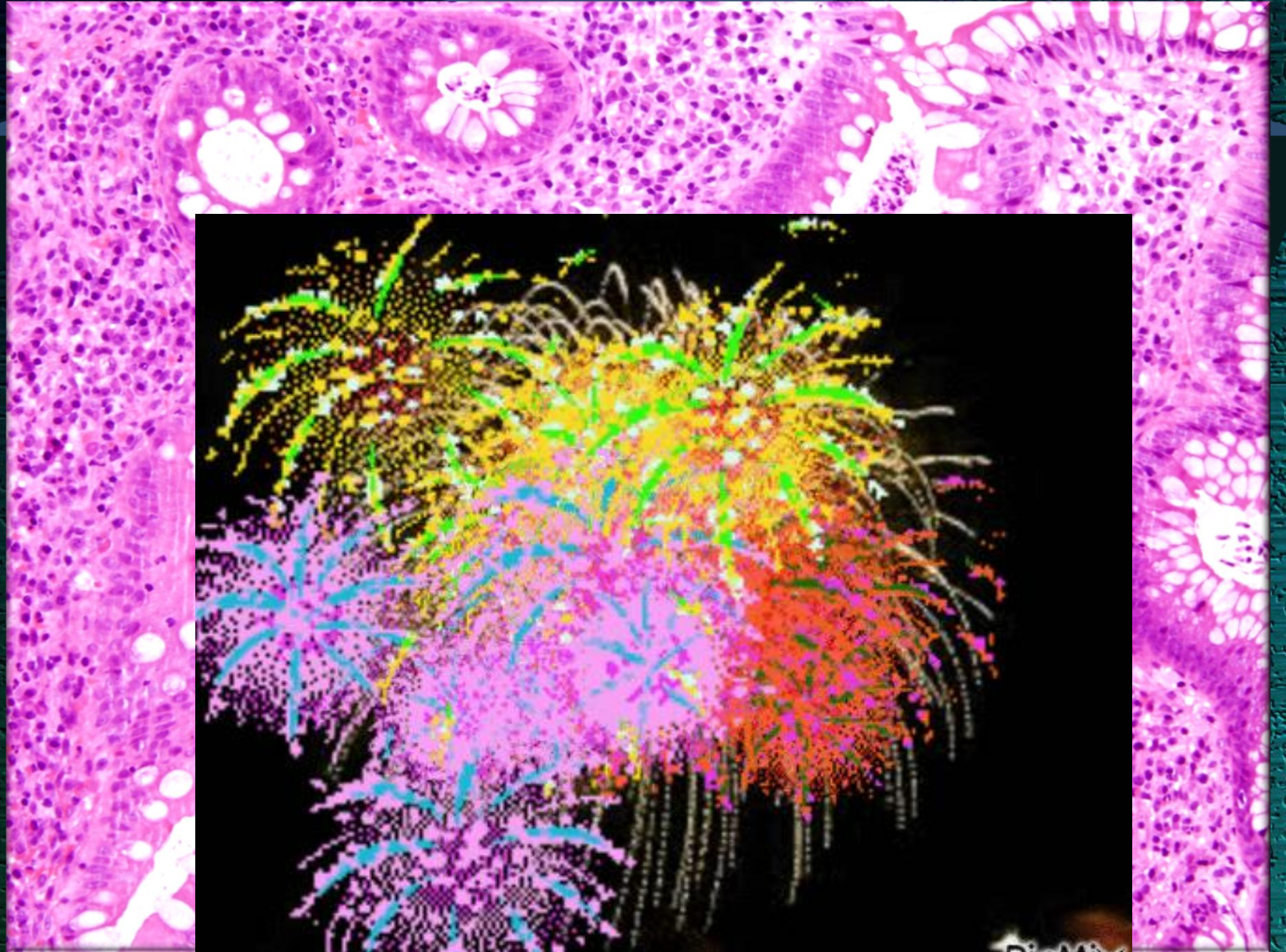








# ACTIVITY OF UC !





**Corticosteroids**

**Salicylates**

**Immunomodulators**

**Biological Agents**



**è nu passaggio dal sonoro al muto.  
E quanno s'è stutata 'a lampetella  
significa ca ll'opera è fernuta  
e 'o primm'attore s'è ghiuto a cuccà.**



**poesia del grande Totò**

# **WHEN TO STOP TREATMENT WITH BIOLOGICAL DRUGS?**

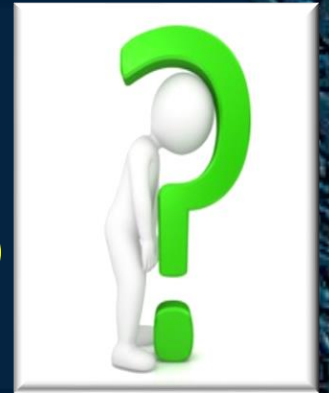




**When it stops working?**



**When it is working but associated with toxicity?**



**When it is still working?**





# When to stop therapy?

ECCO	BSG	ACG	CAG
<b>"no recommendations can be given for the duration of treatment with MTX or anti-TNF although prolonged use of these medications may be considered if needed"</b>	<b>"NICE recommends that maintenance therapy with anti-TNF therapy should continue 12 months or until treatment failure"</b>	<b>"the recommendation uses strong language regarding the benefit of continuing anti-TNF therapy ...the quality of the evidence was high"</b>	<b>"for the patient who has responded favourably to 52 weeks of therapy, the benefit of continued therapy appears to outweigh the risks"</b>

ACG, American College of Gastroenterology;  
BSG, British Society of Gastroenterology;  
CAG, Canadian Association of Gastroenterology;  
ECCO, European Crohn's and Colitis Organisation

Leading Change in

**There are no established guidelines as to whether and when biological therapy can be discontinued**



# When to stop therapy?

## IG-IBD Statement 5F

Maintenance therapy after successful anti-TNF $\alpha$  agents induction is mandatory. Infliximab [EL 1b, RG A], adalimumab [EL 1b, RG B], or AZA/6-MP [EL 2b, RG C], with drained sepsis [EL 4, RG B], should be used as maintenance therapy. All the maintenance therapy should be used for at least 1 year [EL 1b, RG A]; adalimumab could be used up to 3 years [EL 2, RG B]





# The London Position Statement of the World Congress of Gastroenterology on Biological Therapy for IBD With the European Crohn's and Colitis Organization: When to Start, When to Stop, Which Drug to Choose, and How to Predict Response?

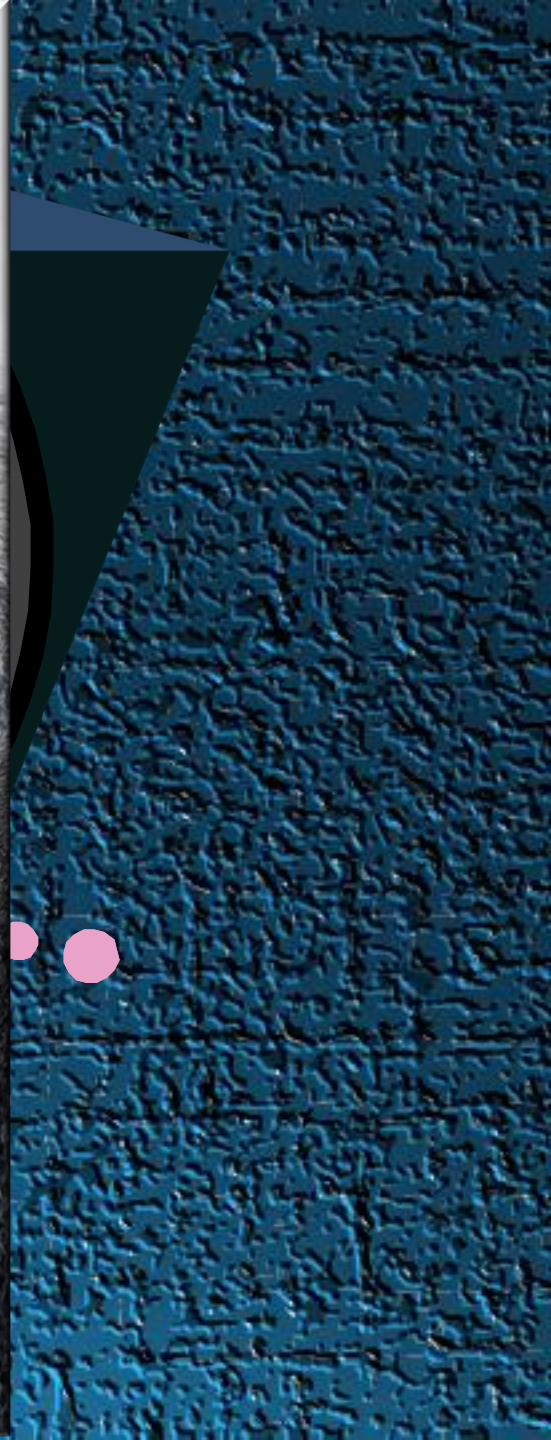
Geert R. D'Haens, MD, PhD<sup>1</sup>, Remo Panaccione, MD<sup>2</sup>, Peter D.R. Higgins, MD<sup>3</sup>, Severine Vermeire, MD, PhD<sup>4</sup>, Miquel Gassull, MD, PhD<sup>5</sup>, Yehuda Chowers, MD<sup>6</sup>, Stephen B. Hanauer, MD<sup>7</sup>, Hans Herfarth, MD<sup>8</sup>, Daan W. Hommes, MD, PhD<sup>9</sup>, Michael Kamm, MD<sup>10,11</sup>, Robert Löfberg, MD<sup>12</sup>, A. Quay<sup>13</sup>, Bruce Sands, MD<sup>14</sup>, A. Sood, MD<sup>15</sup>, G. Watermayer<sup>16</sup>, Bret Lashner, MD<sup>17</sup>, Marc Lémann, MD<sup>18</sup>, Scott Plevy<sup>19</sup>, Walter Reinisch, MD<sup>20</sup>, Stefan Schreiber, MD, PhD<sup>21</sup>, Corey Siegel, MD<sup>22</sup>, Stephen Targan, MD<sup>23</sup>, M. Watanabe, MD<sup>24</sup>, Brian Feagan, MD<sup>25</sup>, William J. Sandborn, MD<sup>26</sup>, Jean Frédéric Colombel, MD, PhD<sup>27</sup> and Simon Travis, MD<sup>28</sup>

*Am J Gastroenterol* 2011; 106:199–212; doi:10.1038/ajg.2010.392

## WCOG Statement 1.31

In patients with UC or CD who have responded to a year of anti-TNF therapy, the benefits of continuing therapy should be weighed against the risks of discontinuation. Withdrawal of therapy is possible in patients with CD who have both complete mucosal healing and no biological evidence of inflammation [EL 2b]. The previous pattern of disease and response to different therapies are essential considerations. There are no data for UC.





Oddio  
Mò che  
ci  
combina  
questo





# STRIDE: treat-to-target recommendations in CD

## Composite endpoint

### Clinical/PRO remission

- Resolution of abdominal pain and normalization of bowel habit
- Should be assessed at a minimum of 3 months during active disease
- Patients' individual goals (e.g. QoL, mood disorders, fatigue, work productivity) should also be addressed: normalisation of QoL as ultimate goal

*AND*

### Endoscopic remission

Absence of ulceration is the target

*OR*

Resolution of findings of inflammation on cross-sectional imaging

Adjunctive measures of disease activity that may be useful in selected cases. **BUT NOT ALONE**

- Histopathology is not a target due to lack of evidence of clinical utility
- Available biomarkers including CRP and faecal calprotectin are not targets



## **Mayo Endoscopic Score**

## **Endoscopic Finding**

**Score 0**

**Normal**

**Score 1**      **Mild disease: erythema, decreased vascularity**

**Score 2**      **Moderate disease: marked erythema, absent vascular pattern,  
friability, erosions**

**Score 3**      **Severe disease: marked erythema, granularity, spontaneous  
bleeding, ulcerations**



**It is of paramount importance to consider that an endoscopic description of Mucosal Healing does not necessarily imply histological healing of the mucosa**

**Several authors stressed the concept that microscopic evidence of inflammation persists in 16-100% of patients with endoscopically quiescent colitis . In addition, active histological inflammation predicts clinical relapse during 12 months of follow-up, whereas endoscopic features did not.**



**Therefore, even considering the potentiality of the new therapeutic approaches, the importance of achieving also histological healing might add further value to future trials**

**On the basis of these evidences, the optimal treatment goal should be the complete resolution of the inflammatory process.**

**This is reached only when confirmed by histological assessment.**





# Beyond Endoscopic Appearance !!!







**NUMBER AND SITE OF BIOPSIES**

**ACTIVITY AND INACTIVITY OF IBD**



**SCORES OF MUCOSAL HEALING**





# **FIRST PROBLEM !**

**NUMBER AND  
SITE OF BIOPSIES**





## Histologic Remission: The Ultimate Therapeutic Goal in Ulcerative Colitis?

Laurent Peyrin-Biroulet,<sup>\*,‡</sup> Aude Bressenot,<sup>\*,§</sup> and Wendy Kampman<sup>||</sup>

<sup>\*</sup>Inserm, U954, France; <sup>‡</sup>Department of Hepato-Gastroenterology, <sup>§</sup>Department of Pathology, University Hospital of Nancy, University of Nancy

## Microscopic activity in ulcerative colitis: what does it mean?

S A Riley, V Mani, M J Goodman, S Dutt, M E Herd

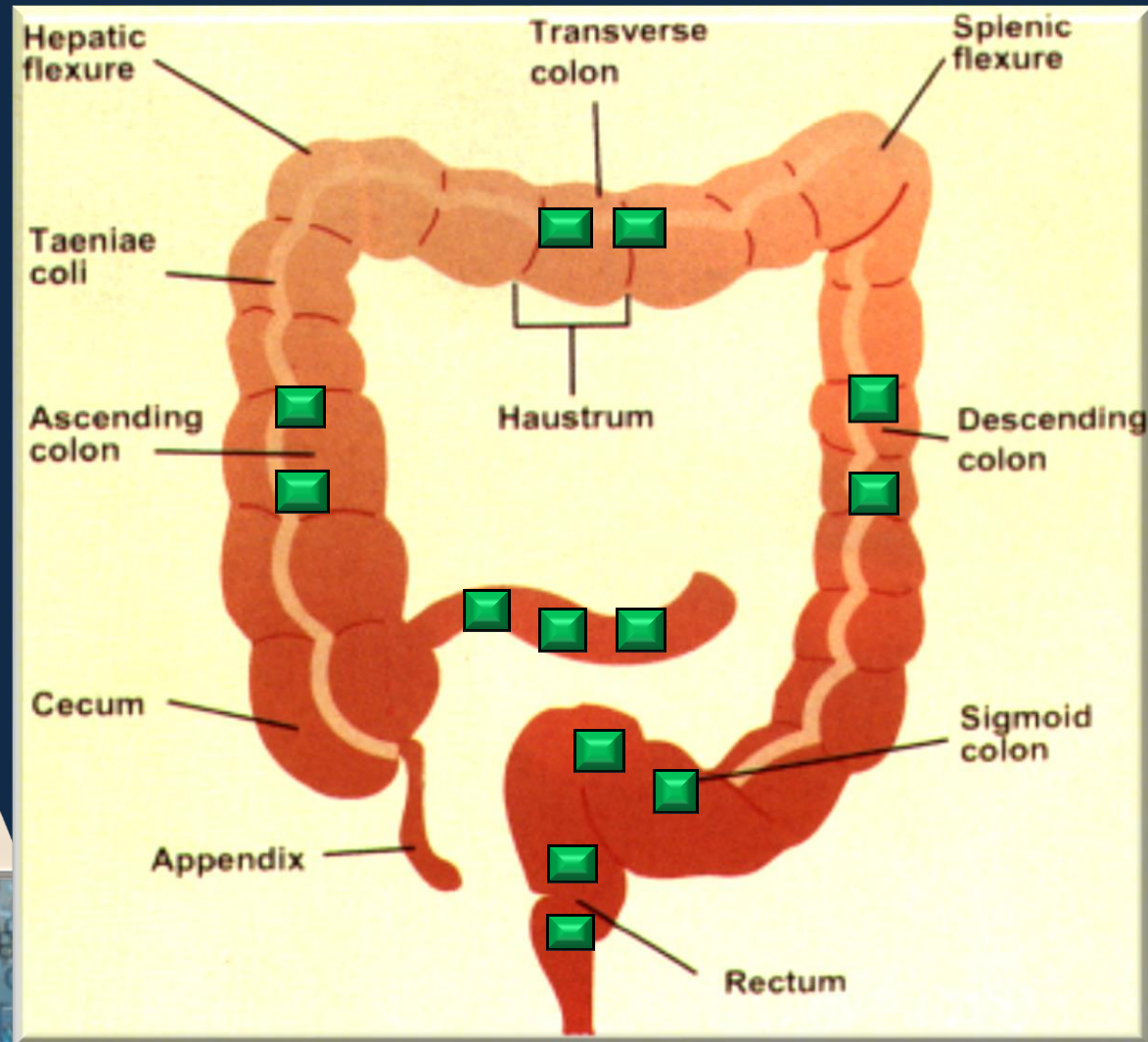
**No indications about the number of biopsies and the number of different type of cells to evaluate !!!**

## Systematic review with meta-analysis: mucosal healing is associated with improved long-term outcomes in Crohn's disease

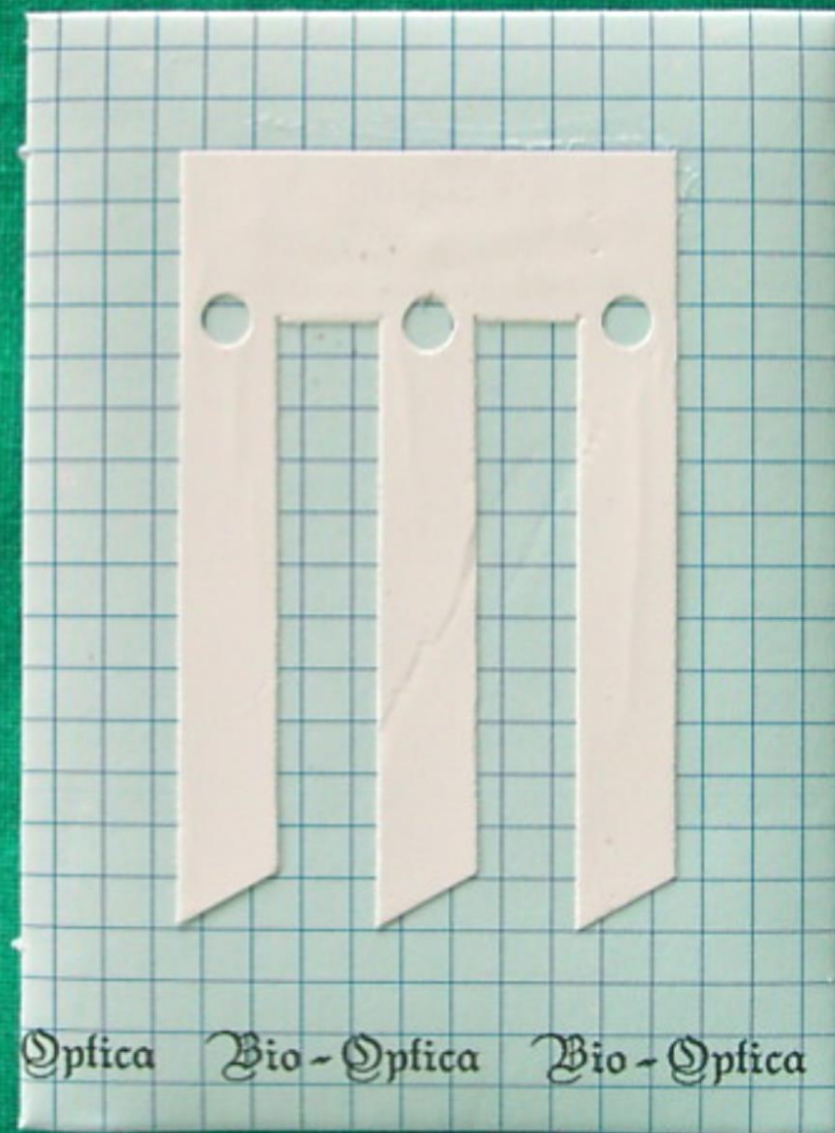
S. C. Shah, J.-F. Colombel, B. E. Sands & N. Narula



# ADEQUATE SAMPLING



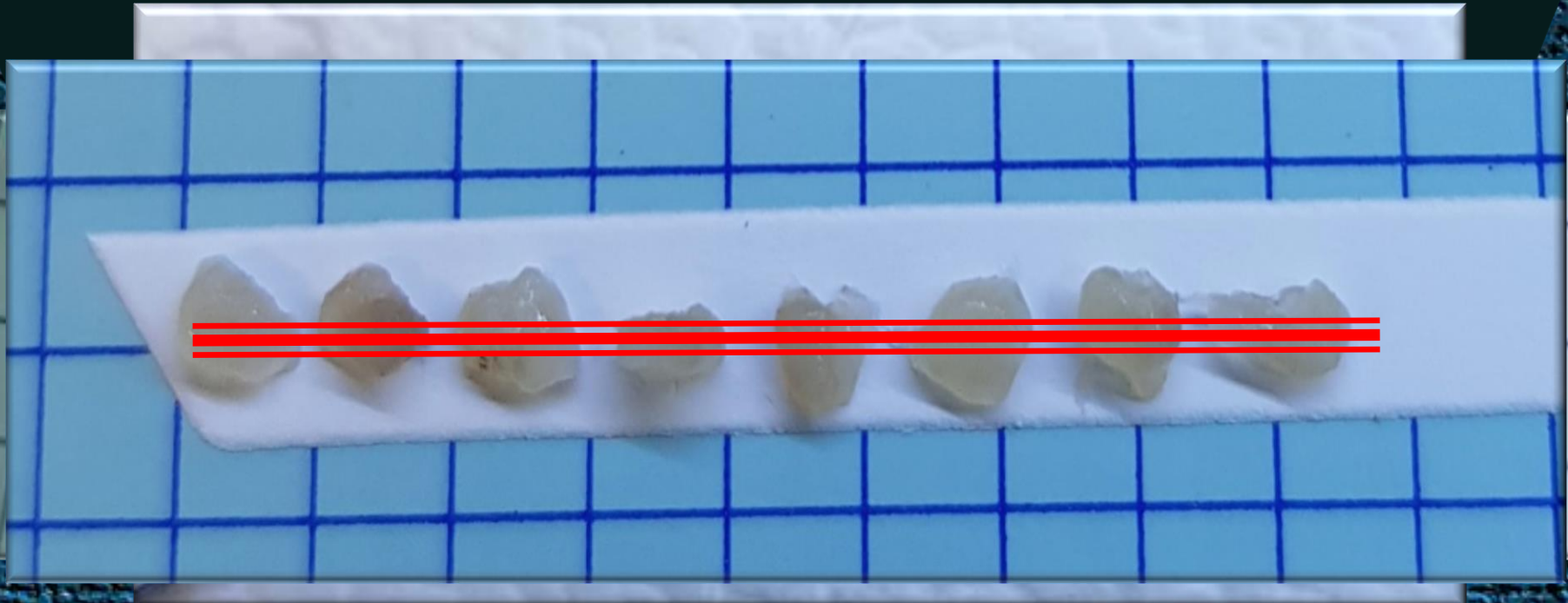




Optica Bio - Optica Bio - Optica



## Ileo Colon

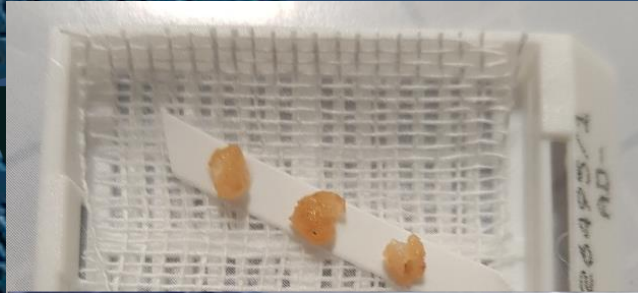








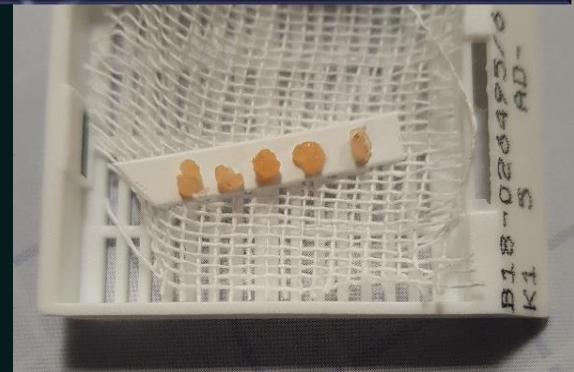
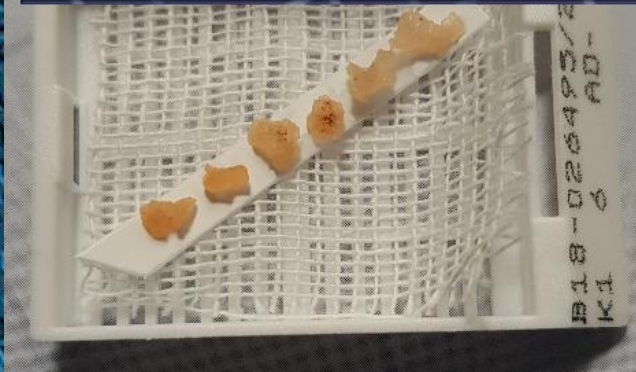
ILEO



SIGMA-RETTO



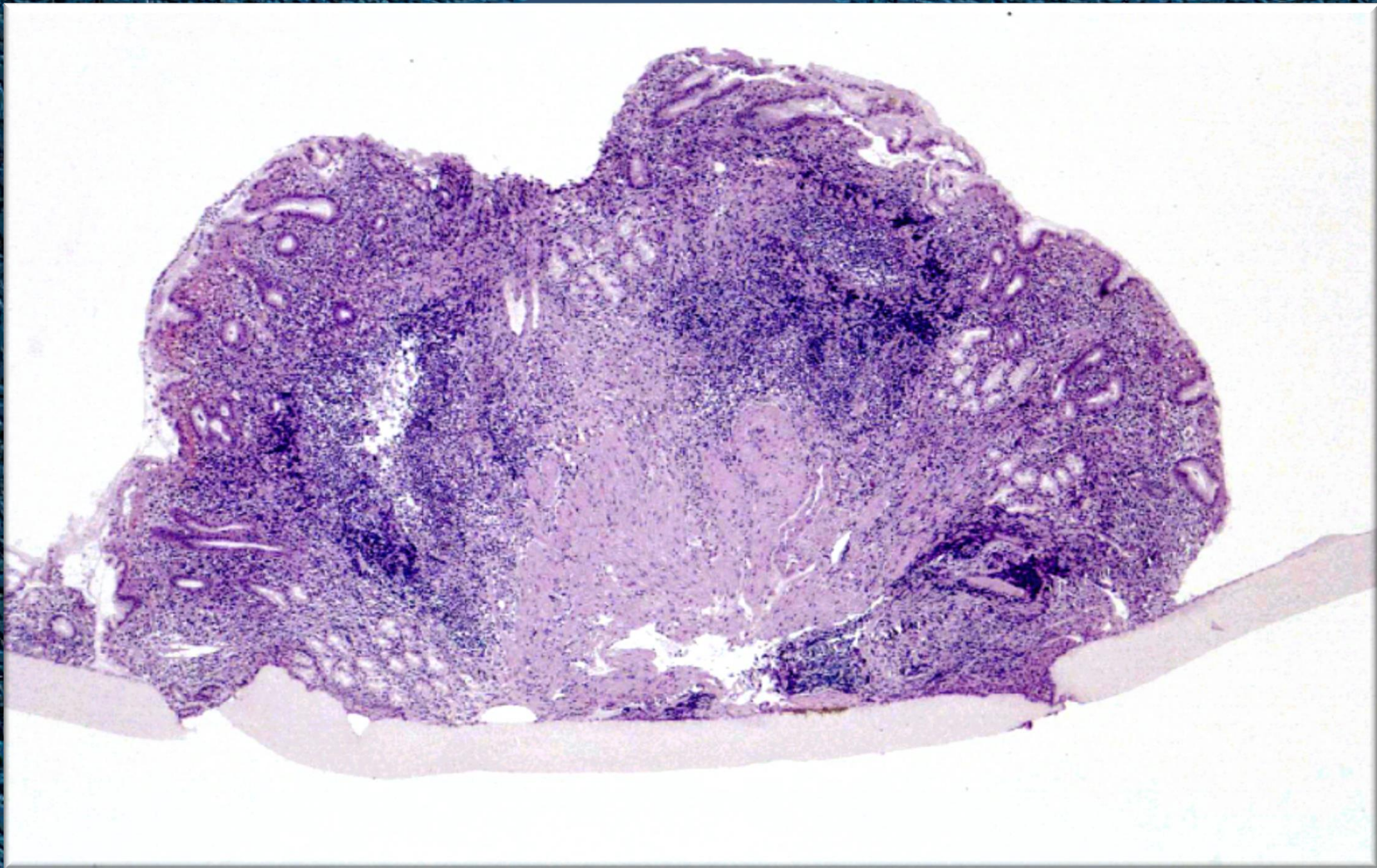
DUODENO-BULBO













## BIOPSIA COLORETTALE

SET DI DATI CLINICI PER  
SOSPETTA COLITE IDIOPATICA

Nome del paziente

Sesso F ☐ M ☐

Data di nascita

Data dell'esame

Ospedale

### SINTOMI PRINCIPALI

Diarrea

Acquosa ☐ Ematica ☐

Durata totale della malattia

Durata del presente episodio

Stato clinico attuale

Aspetto sigmoidoscopico

Caratteristiche della mucosa



Estensione della malattia

☐

retto

☐

retto-sigma

☐

lato sinistro

☐

trasverso

☐

ascendente

☐

pancolite

☐

Pattern

continuo

☐

segmentale

☐

### SEDE DELLA BIOPSIA/E

Opinione clinica

Informazioni aggiuntive (se appropriate)

Precedenti interventi G.I.

Terapia

Risultato colture fecali

Altre malattie presenti

OF

ES



# **SECOND PROBLEM !**

**ACTIVITY AND  
INACTIVITY OF  
IBD**





## Features of “activity”

- Cryptitis, crypt abscesses
- Epithelial injury
- Mucodepletion, cuboidal shape, nuclear enlargement, ulcer



### Neutrophils



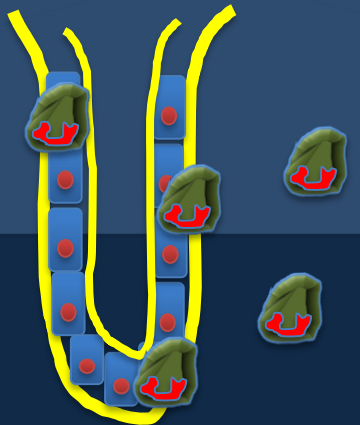
(cryptitis / crypt abscess formation)

markers of disease activity

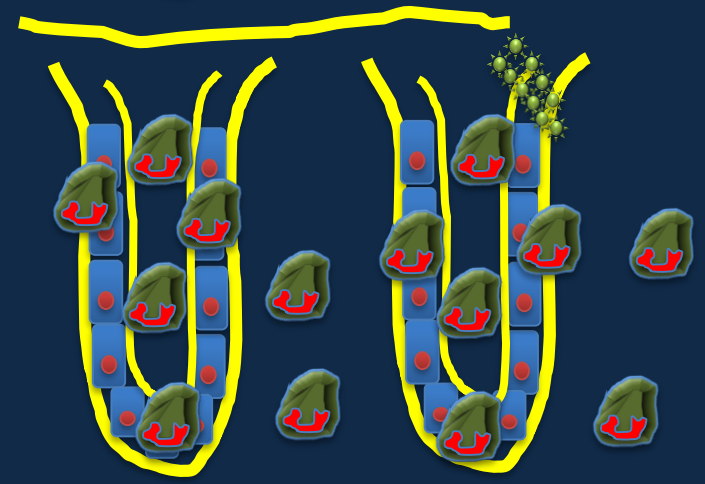


# Active Diseases

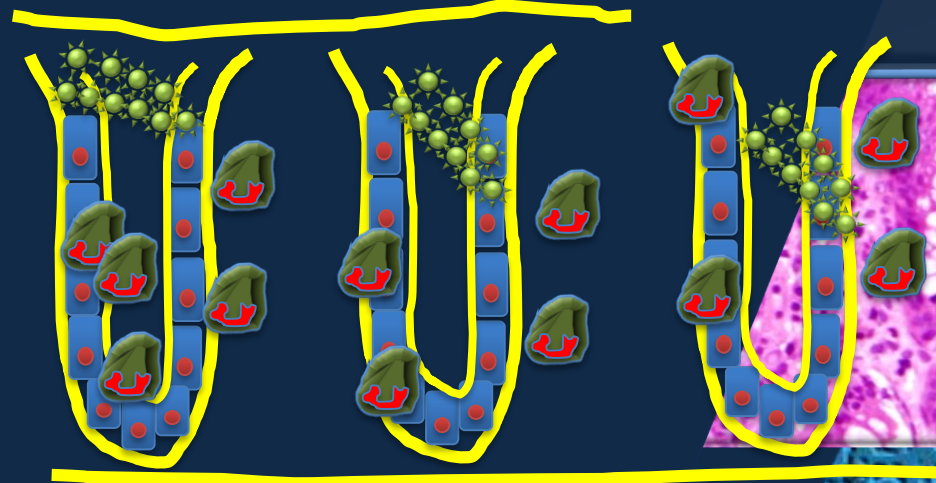
LOW



MODERATE



SEVERE



e





**ABSENCE OF  
NEUTROPHILS!!!**



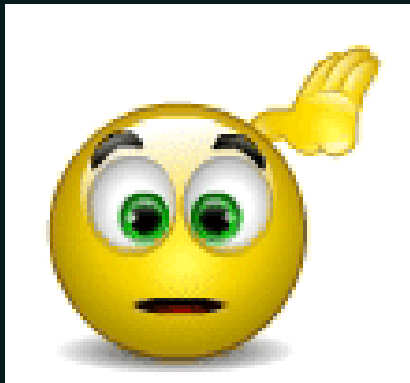


**THERE IS A PROBLEM..**





**Several authors stressed the concept that microscopic evaluation of Mucosal Healing is based not only on neutrophils but also on other cellular elements such as plasma cells and eosinophils...**







Available online at [www.sciencedirect.com](http://www.sciencedirect.com)

ScienceDirect



## REVIEW ARTICLE

# Systematic review: Histological remission in inflammatory bowel disease. Is 'complete' remission the new treatment paradigm? An IOIBD initiative

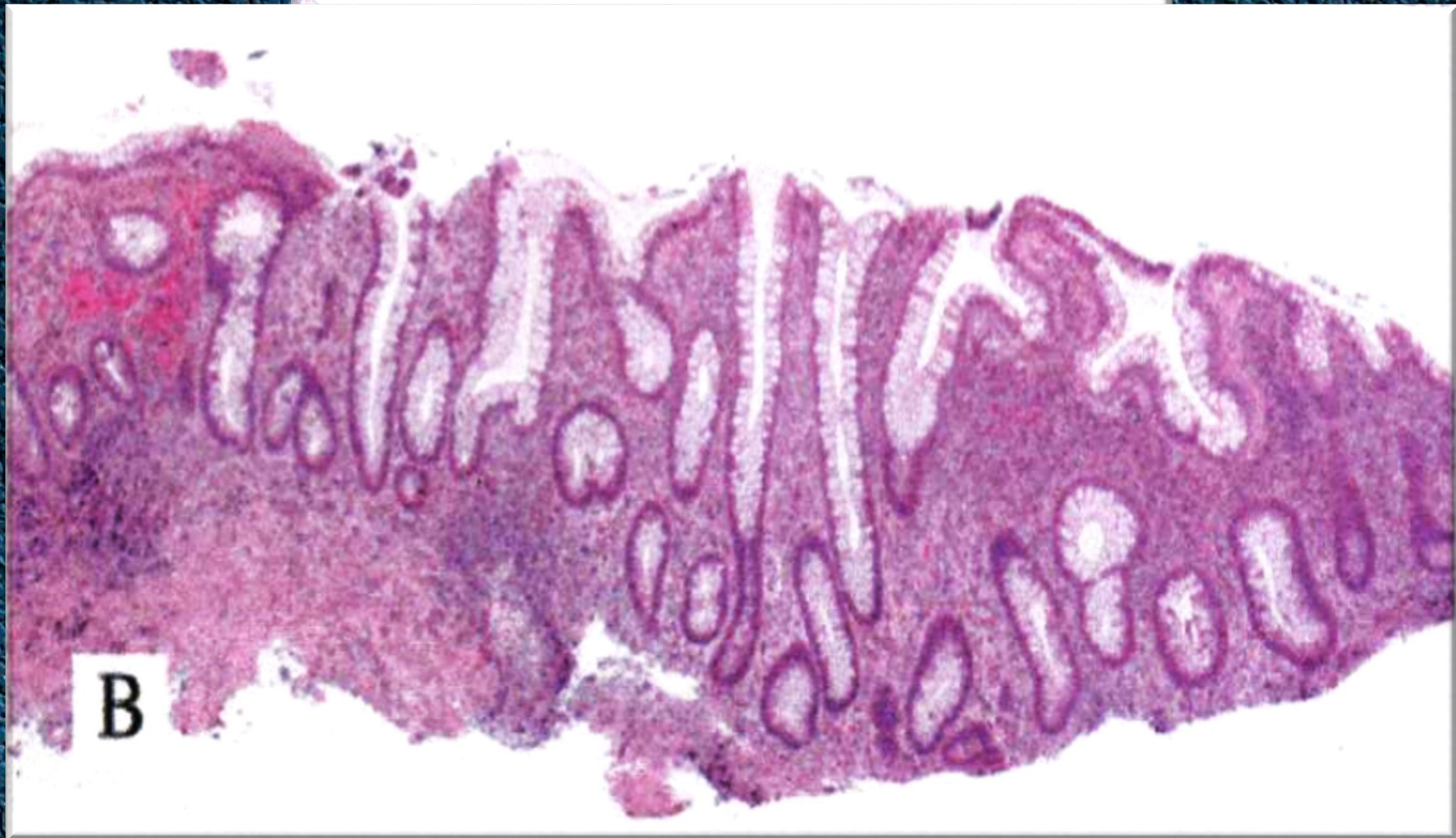


R.V. Bryant<sup>a</sup>, S. Winer<sup>b</sup>, SPLTravis<sup>a</sup>, R.H. Riddell<sup>b,\*</sup>

<sup>a</sup> Translational Gastroenterology Unit, John Radcliffe Hospital, Oxford University Hospitals, United Kingdom

<sup>b</sup> Department of Pathology and Laboratory Medicine, Mt Sinai Hospital, 600 University Avenue, Toronto, ON M5G 1X5, Canada







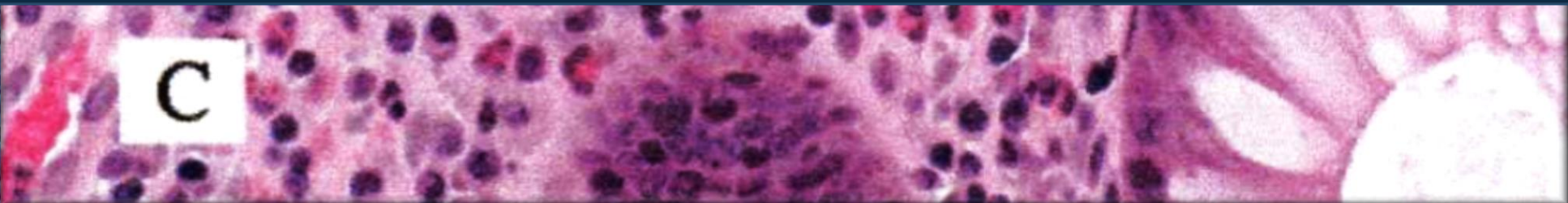


**The International Organization of Inflammatory Bowel Disease  
Has embraced histologic remission and defines it as:**

**A) The absence of neutrophils ( both in the crypts and lamina propria)**

**WHICH IS THE NORMAL NUMBER OF PLASMA CELLS ?**

**WHICH IS THE NORMAL NUMBER OF EOSINOPHILS ?**





## Study Highlights

1684

SEASE

### WHAT IS CURRENT KNOWLEDGE

- ✓ Mucosal healing is a key endpoint to medical therapy in ulcerative colitis (UC).
- ✓ Mucosal healing reduces risk of hospitalization and colectomy.

**No indications about the number of biopsies and the number of basal plasma cells to evaluate !!!**

- ✓ There is a poor correlation between endoscopic and microscopic disease activity.
- ✓ All patients with mucosal healing should have biopsies to assess the histologic disease activity.
- ✓ Closer follow-up and optimization of medical therapy should be considered in patients with basal plasmacytosis.



**I DISAGREE !!**

**BECAUSE BASAL PLASMACYTOSIS IS  
ONLY A DIAGNOSTIC MARKER AND  
NOT INDICATIVE OF REMISSION!!!!**



## **Microscopic Features of Ulcerative Colitis**

**Basal plasmacytosis at the initial onset has a high predictive value for the diagnosis of IBD**

**Basal plasmacytosis is a good diagnostic feature in established ulcerative colitis.**





# EOSINOPHILIC GASTROENTERITIS

Table 4: Proposed quantitative criteria for eosinophilic gastritis and eosinophilic gastroenteritis

Author(s)	Diagnosis	Criteria
Hurrell et al. <sup>78</sup>	Histologic eosinophilic gastritis	≥30 eosinophils per HPF in at least 5 separate HPFs (if H. pylori present, eosinophilia must persist several months post eradication)
Collins <sup>77</sup>	Eosinophilic gastritis	≥30 eosinophils per HPF in at least 5 separate HPFs
Ko et al. <sup>72</sup>	Eosinophilic gastritis (pediatric)	≥70 eosinophils per HPF involving >3 HPFs
Bischoff and Ulmer <sup>76</sup>	Eosinophilic gastroenteritis	>20 eosinophils per HPF
Collins <sup>77</sup>	Eosinophilic enteritis	>52 eosinophils per HPF in the duodenum OR >56 eosinophils per HPF in the ileum
Collins <sup>77</sup>	Eosinophilic colitis	>100 eosinophils per HPF in the right colon OR >84 eosinophils per HPF in the transverse or descending colon OR >64 eosinophils per HPF in the rectosigmoid colon
Turner et al. <sup>84</sup>	Colonic eosinophilia	>50 eosinophils per HPF in the right colon >35 eosinophils per HPF in the transverse colon >25 eosinophils per HPF in the left colon

**HOW MANY  
EOSINOPHILS?**

**THERE ARE NO  
ABSOLUTE  
CUT-OFF**



Poor prognostic factors for UC patients

Clinical factors



**EOSINOPHILS**



Disease and treatment factors

Reinisch W, et al. Clin Gastroenterol Hepatol. 2015;13:635–42



# Severe eosinophilic infiltration in colonic biopsies predicts patients with ulcerative colitis not responding to medical therapy

**P. Zezos\*†, K. Patsiaoura‡, A. Nakos\*, A. Mpoumponaris\*, T. Vassiliadis\*, O. Giouleme\*, M. Pitiakoudis§, G. Kouklakis† and N. Evgenidis\***

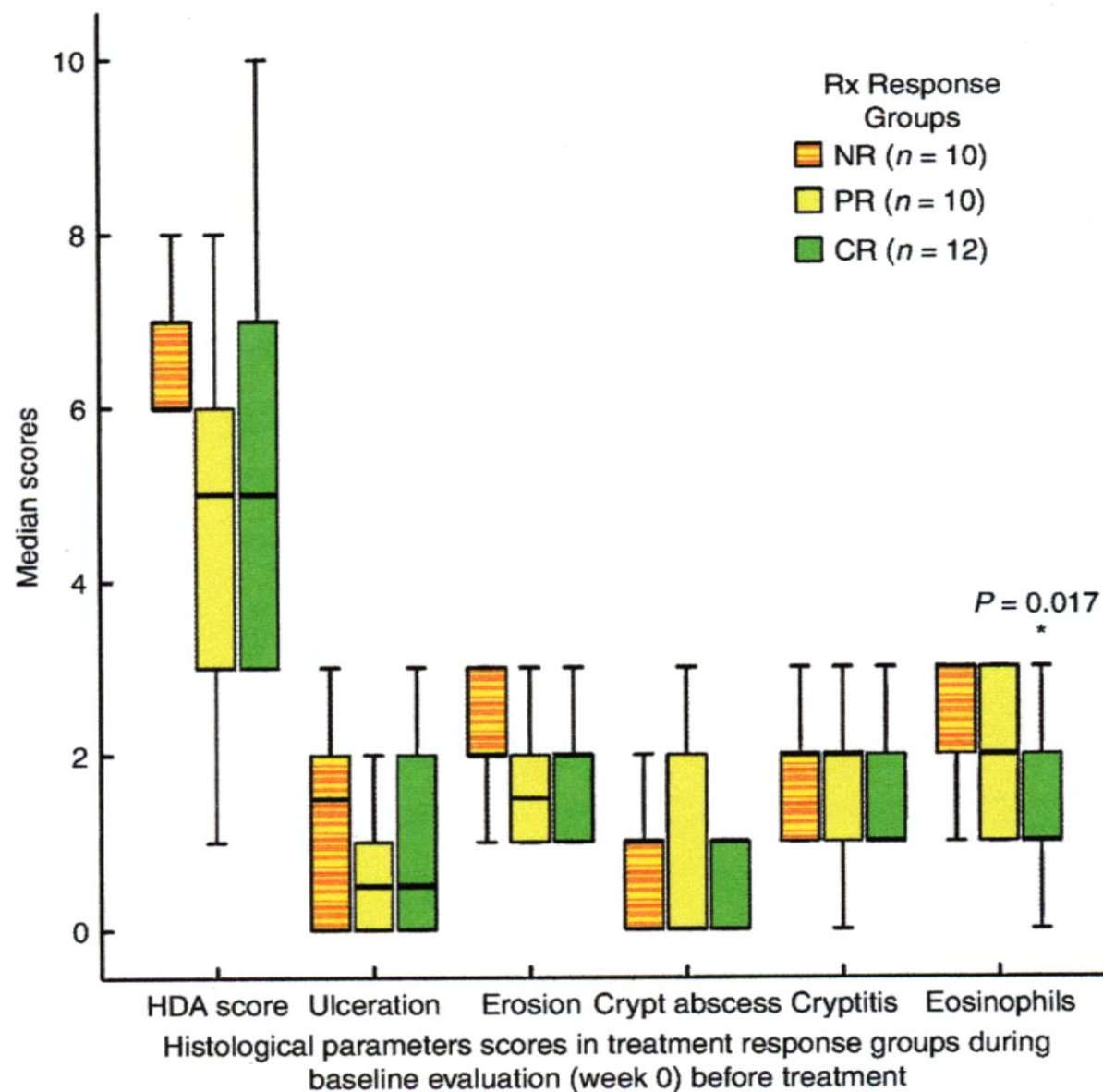
\*Division of Gastroenterology, 2<sup>nd</sup> Propaedeutic Department of Internal Medicine, "Hippokration" General Hospital, Aristotle University of Thessaloniki, Thessaloniki, Greece, †Gastrointestinal Endoscopy Unit, Democritus University of Thrace, University General Hospital of Alexandroupolis, Alexandroupolis, Greece, ‡Department of Pathology, "Hippokration" General Hospital, Thessaloniki, Greece and §2<sup>nd</sup> Department of Surgery, Democritus University of Thrace, University General Hospital of Alexandroupolis, Alexandroupolis, Greece

Received 17 March 2014; accepted 5 June 2014; Accepted Article online 14 July 2014

**COLORECTAL DISEASE**



**Figure 4** Histological parameters scores, according to treatment (Rx) response group, during baseline evaluation (week 0). \*In patients with complete response, eosinophilic infiltration of the lamina propria was significantly less severe compared with that of patients with no response to treatment. No other significant differences in histological parameters were observed between Rx response groups. Kruskal–Wallis nonparametric ANOVA and the Mann–Whitney *U*-test were used for statistical evaluation. Boxes display the median value along with 25th–75th percentiles and whiskers display minimum–maximum values. CR, complete response; NR, no response; PR, partial response.





Grading of histological disease activity (HDA) was performed semi-quantitatively for each of four histological feature (ulceration, erosion, crypt abscess and cryptitis), using a scale with scores from 0 to 3 (no or minimal HDA = 0, mild HDA = 1, moderate HDA = 2 and severe HDA = 3), resulting in a total HDA score of 0–12.

This total HDA score was categorized into four grades of histological disease activity: inactive/minimal (score 0–3); mild (score 4–6); moderate (score 7–9); and severe (score 10–12).

Moreover, in each case it was estimated whether there was a predominance of eosinophils among inflammatory cells (lymphocytes, plasmacytes, neutrophils and eosinophils) infiltrating the colonic lamina propria. The presence and the degree of eosinophil predominance was scored as:  
0, none or minimal; 1, mild; 2, moderate; and 3, severe (Fig. 1a, b).

Histological evaluation and scoring were performed under light microscopy, at x400 magnification, by one experienced gastrointestinal pathologist (KP) who knew the diagnosis but was blinded to the clinical status.

The most inflamed site in the left colon or rectum was used for the assessment of endoscopic and histological scores.



**28 cases of UC**

**21 Cases: 12 M + 9 F (median age 31,57)**

**7 Controls: 5 M + 2 F (median age 46,85)**

**> 60 in two biopsies at HPF 40x**

**Duration of disease**

**Cases: 2 - 13 years**

**Controls: 2 – 5 years**

**Therapy**

**Cases: inefficacy of therapy → IFX**

**Controls: efficacy of therapy → remission**



Edited by  
JAMES J. LEE AND HELENE F. ROSENBERG

# EOSINOPHILS IN HEALTH AND DISEASE



COLONIC HYP  
PREDIC

I am pleased  
pu

MAY HELP TO  
ERAPY

accepted for  
/.

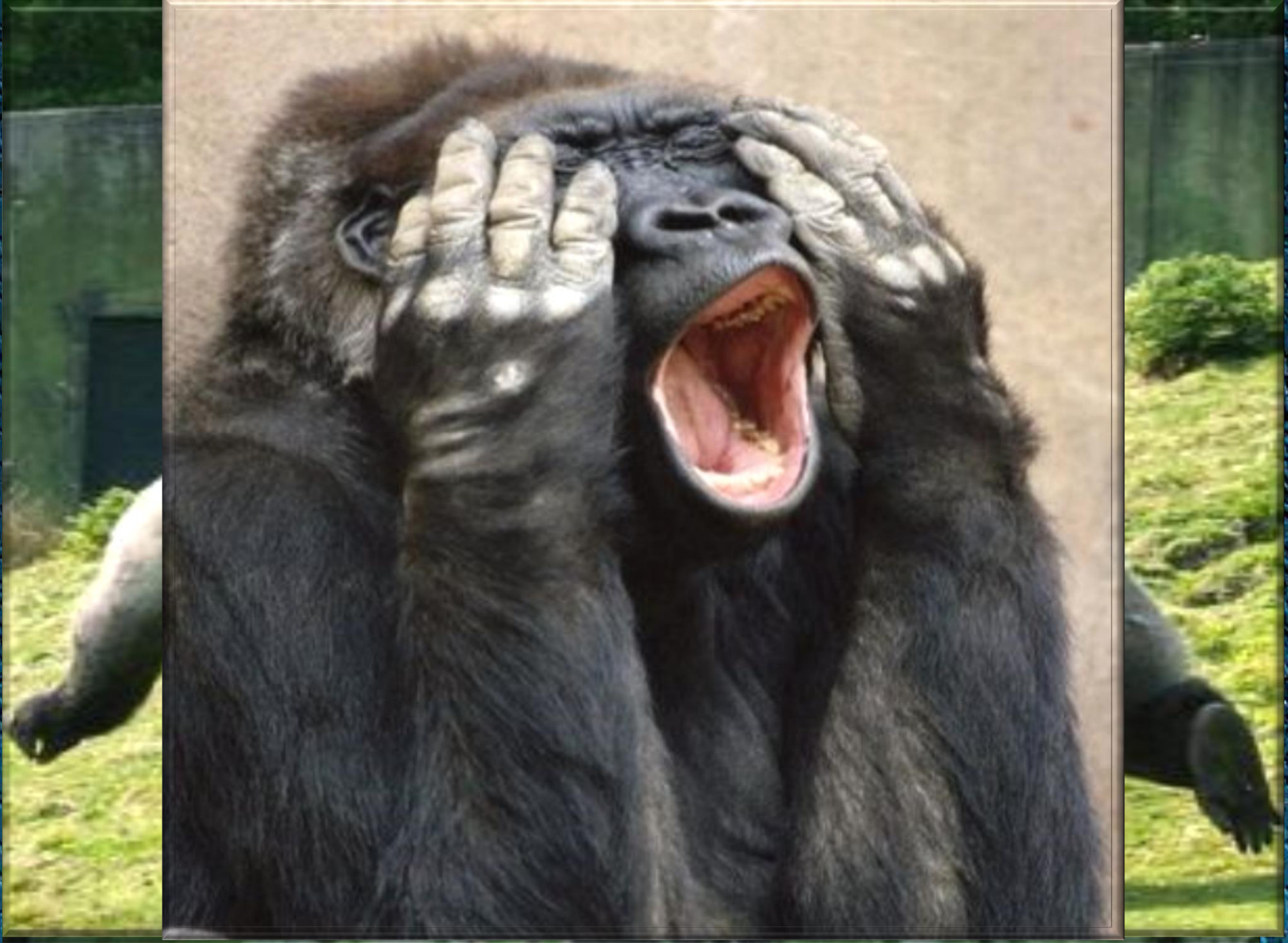


# **THIRD PROBLEM !**

**SCORES OF  
MUCOSAL  
HEALING**









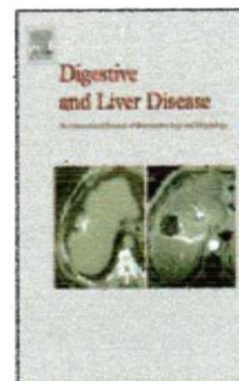


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## Digestive and Liver Disease

journal homepage: [www.elsevier.com/locate/dld](http://www.elsevier.com/locate/dld)



### Review Article

## Definition and evaluation of mucosal healing in clinical practice



Silvia Mazzuoli<sup>a</sup>, Francesco W. Guglielmi<sup>a,\*</sup>, Elisabetta Antonelli<sup>b</sup>, Marianna Salemmme<sup>c</sup>,  
Gabrio Bassotti<sup>b</sup>, Vincenzo Villanacci<sup>c</sup>

<sup>a</sup> Gastroenterology and Artificial Nutrition Department, "San Nicola Pellegrino" Hospital Trani, BT, Italy

<sup>b</sup> Gastroenterology and Hepatology Section, Department of Clinical & Experimental Medicine, University of Perugia, Italy

<sup>c</sup> Department of Pathology, Spedali Civili of Brescia, Italy





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ORIGINAL ARTICLE

# **Disease activity and mucosal healing in inflammatory bowel disease: a new role for histopathology?**

Rish K. Pai<sup>1</sup>  • Karel Geboes<sup>2</sup>



### Table 3 Selected histologic scoring systems in ulcerative colitis

**Truelove and Richards, 1956 [11]** 3-grade scale ranging from no inflammation to severe inflammation

**Saverymuttu index, 1986 [59]** Generates a total score based on enterocyte damage, crypt abnormalities, lamina propria chronic inflammation, lamina propria neutrophilic inflammation

**Riley score, 1991 [60]** Scores 6 items on a 4 point scale (0–4):

1. Neutrophils in lamina propria
2. Crypt abscesses
3. Mucin depletion
4. Surface epithelial integrity
5. Chronic inflammatory cell infiltrate
6. Crypt architectural abnormalities

**Modified Riley, 2005 [61]** Score ranges from 0 to 7

Score 0: Normal biopsy or inactive colitis

Score 1: Scattered individual neutrophils in lamina propria

Score 2: Patchy collections of neutrophils in lamina propria

Score 3: Diffuse neutrophilic infiltrate in lamina propria

Score 4: Neutrophils in epithelium, <25% crypts involved

Score 5: Neutrophils in epithelium, 25 to 75% of crypts involved

Score 6: Neutrophils in epithelium, >75% crypts involved

Score 7: Erosion or ulceration



**Gupta/Harpaz score, 2007 [21]** Activity is graded based on cryptitis and ulcers/erosions

Inactive (no cryptitis)

Mildly active (<50% of crypts involved)

Moderately active (>50% crypts involved)

Severely active (ulcers or erosions)

**Chicago score, 2007 [62]** Score ranges from 0 to 6

Normal (score 0)

Quiescent (score 1)

Increased lamina propria neutrophils without definite intraepithelial neutrophils (score 2)

Cryptitis without crypt abscesses (score 3)

Crypt abscesses in <50% of crypts (score 4)

Crypt abscesses in >50% of crypts or erosions/ulceration (score 5)

**Robarts histopathologic index, 2016**

[26]

Based on the Geboes score. Measures those items that correlate with histologic severity are reproducible and respond to therapies. Calculated score that ranges from 0 to 33

$RHI = 1 \times \text{chronic inflammatory cell infiltrate (0-3)} + 2 \times \text{lamina propria neutrophils (0-3)} + 3 \times \text{neutrophils in epithelium (0-3)} + 5 \times \text{erosions or ulceration (0-3)}$  [combines Geboes subscores 5.1 and 5.2]

**Nancy index, 2016 [27]** 4-point scale taking into account both chronic lamina propria inflammation and active inflammation

Grade 0: No histological significant disease (no or only mild increase in chronic inflammatory cells)

Grade 1: Chronic inflammatory cell infiltrate with no acute inflammatory cell infiltrate

Grade 2: Mildly active disease

Grade 3: Moderately active disease

Grade 4: Severely active disease (ulceration)



**Table 2. Geboes' score for the calculation of histological activity with additional data about basal plasmacytosis.**

**Grade 0 Structural (architectural) changes**

- 0.0 No abnormality**
- 0.1 Mild abnormality**
- 0.2 Mild or moderate diffuse or multifocal abnormalities**
- 0.3 Severe diffuse or multifocal abnormalities**

**Grade 1 Chronic inflammatory infiltrate**

- 1.0 No increase**
- 1.1 Mild but unequivocal increase**
- 1.2 Moderate increase**
- 1.3 Marked increase**

**Grade 2A Eosinophils in the lamina propria**

- 2A.0 No increase**
- 2A.1 Mild but unequivocal increase**
- 2A.2 Moderate increase**
- 2A.3 Marked increase**

**Grade 2B Neutrophils in the lamina propria**

- 2B.0 No increase**
- 2B.1 Mild but unequivocal increase**
- 2B.2 Moderate increase**
- 2B.3 Marked increase**



Table 2. Geboes' score for the calculation of histological activity with additional data about basal plasmacytosis.

Grade 3 Neutrophils in the epithelium

3.0 None

3

3

3

3

3

3

3

3

part of crypt

4.2 Probable: marked attenuation

4.3 Unequivocal crypt destruction

Grade 5 Erosion or ulceration

5.0 No erosion, ulceration, or granulation

**No indications about the number of biopsies and the number of different type of cells to evaluate !!!**

Extra basal plasmacytosis

Absent

Focal basal plasmacytosis

Diffuse basal plasmacytosis



Original

A Sin

Aranzaz

Bart Ler

Marc Fe

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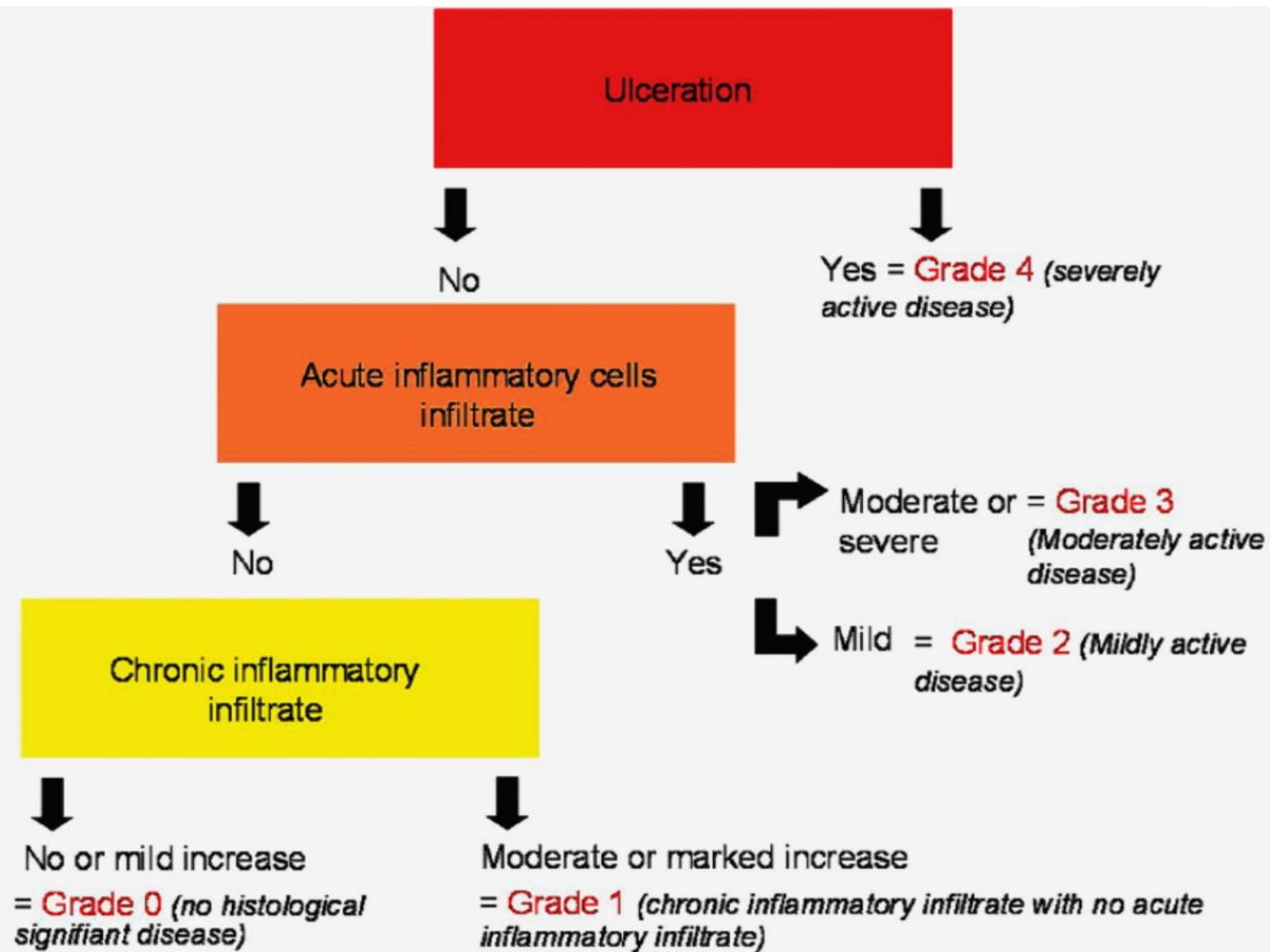
Herestraat 49

**Table 2.** The proposed Simplified Geboes Score.

Grade 0:	0.0 No abnormalities
No inflammatory activity	0.1 Presence of architectural changes
	0.2 Presence of architectural changes and chronic mononuclear cell infiltrate
Grade 1: Basal plasma cells	1.0 No increase
	1.1 Mild increase
	1.2 Marked increase
Grade 2A: Eosinophils in lamina propria	2A.0 No increase
	2A.1 Mild increase
	2A.2 Marked increase
Grade 2B: Neutrophils in lamina propria	2B.0 No increase
	2B.1 Mild increase
	2B.2 Marked increase
Grade 3: Neutrophils in epithelium	3.0 None
	3.1 < 50% crypts involved
	3.2 > 50% crypts involved
Grade 4:	4.0 None
Epithelial injury	4.1 Marked attenuation
[in crypt and surface epithelium]	4.2 Probable crypt destruction: probable erosions
	4.3 Unequivocal crypt destruction: unequivocal erosion
	4.4 Ulcer or granulation tissue



# The Nancy Index





# The Nancy Index

Absence of significant histological disease  
grade 0

Chronic inflammatory infiltrate with no acute inflammatory infiltrate  
grade 1

Mildly active disease  
grade 2

Moderately active disease  
grade 3

Severely active disease  
grade 4



# Robarts Histological Index

## Component

Intercept

Chronic inflammatory infiltrate

0=No increase

1=Mild but unequivocal increase

2=Moderate increase

3=Marked increase

Lamina propria neutrophils

0=None

1=Mild but unequivocal increase

2=Moderate increase

3=Marked increase

Neutrophils in epithelium

0=None

1=<5% crypts involved

2=<50% crypts involved

3=>50% crypts involved

Erosion or ulceration

0=No erosion, ulceration or granulation tissue

1=Recovering epithelium+adjacent inflammation

1=Probable erosion—focally stripped

2=Unequivocal erosion

3=Ulcer or granulation tissue

SE, standard error.



**Table 2** Inflammatory bowel disease histological grading proposal (ECAP system).

Histopathology	Grade/score		
<b>E – Extent of inflammation</b>			
Focal	1		
Multifocal (patchy)	2		
Diffuse	3		
<b>C – Chronicity</b>			
<b>C1 Crypt architectural alteration</b>			
None	0		
Focal alteration	1		
Patchy distortion (<50%)	2		
Diffuse distortion (>50%)	3		
<b>C2 Paneth cell metaplasia</b>			
None	0		
Present	1		
<b>A – Activity of inflammation</b>			
<b>A1 Surface epithelium</b>			
Normal	0		
Reactive changes (mucin depletion/villiform)	1		
Neutrophilic infiltration	2		
Erosion	3		
Ulceration	4		
<b>A2 Neutrophilic cryptitis</b>			
>5%			
<50%			
>50%			
		<b>A3 Crypts abscess</b>	
		None	0
		Present	1
		<b>A4 Crypts destruction</b>	
		None	0
		Crypt destruction	1
		<b>A5 Lamina propria mononuclear cellularity</b>	
		Normal	0
		Mild increase	1
		Moderate increase	2
		Severe increase	3
		<b>A6 Basal plasmacytosis</b>	
		None	0
		Focal	1
		Diffuse	2
		<b>A7 Lamina propria neutrophilic infiltration</b>	
		None	0
		Rare	1
		Scattered	2
		Extensive	3
		<b>P – Plus/others</b>	
		<b>P1 Lamina propria eosinophilic infiltration</b>	
		None	0
		Mild	1
		Moderate	2
		Severe	3
		<b>P2 Lymphoid follicles/aggregates</b>	
		None	0
		Rare	1
		Prominent	2
		<b>Total score</b>	



**EXTREMELY COMPLICATED  
AND SUBJECTIVE !**

**NOT APPLICABLE IN  
ROUTINE DIAGNOSIS !**



# PROPOSAL

SIMPLIFIED SCORE		
Patient	Morphological element	Grade
#1		
	Crypt Abscess	Absent 0
		Present 1
	Active inflammatory infiltrate in lamina propria	Moderate (2) 2
		Severe (3) 3
		Ileum
	Sites involved from disease	Right Colon
		Transverse Colon
		Left Colon
		Rectum

Simple and Quick !!!





**USEFULNESS OF DIFFERENT  
PATHOLOGICAL SCORES TO  
ASSESS MUCOSAL HEALING IN  
INFLAMMATORY BOWEL  
DISEASE:**

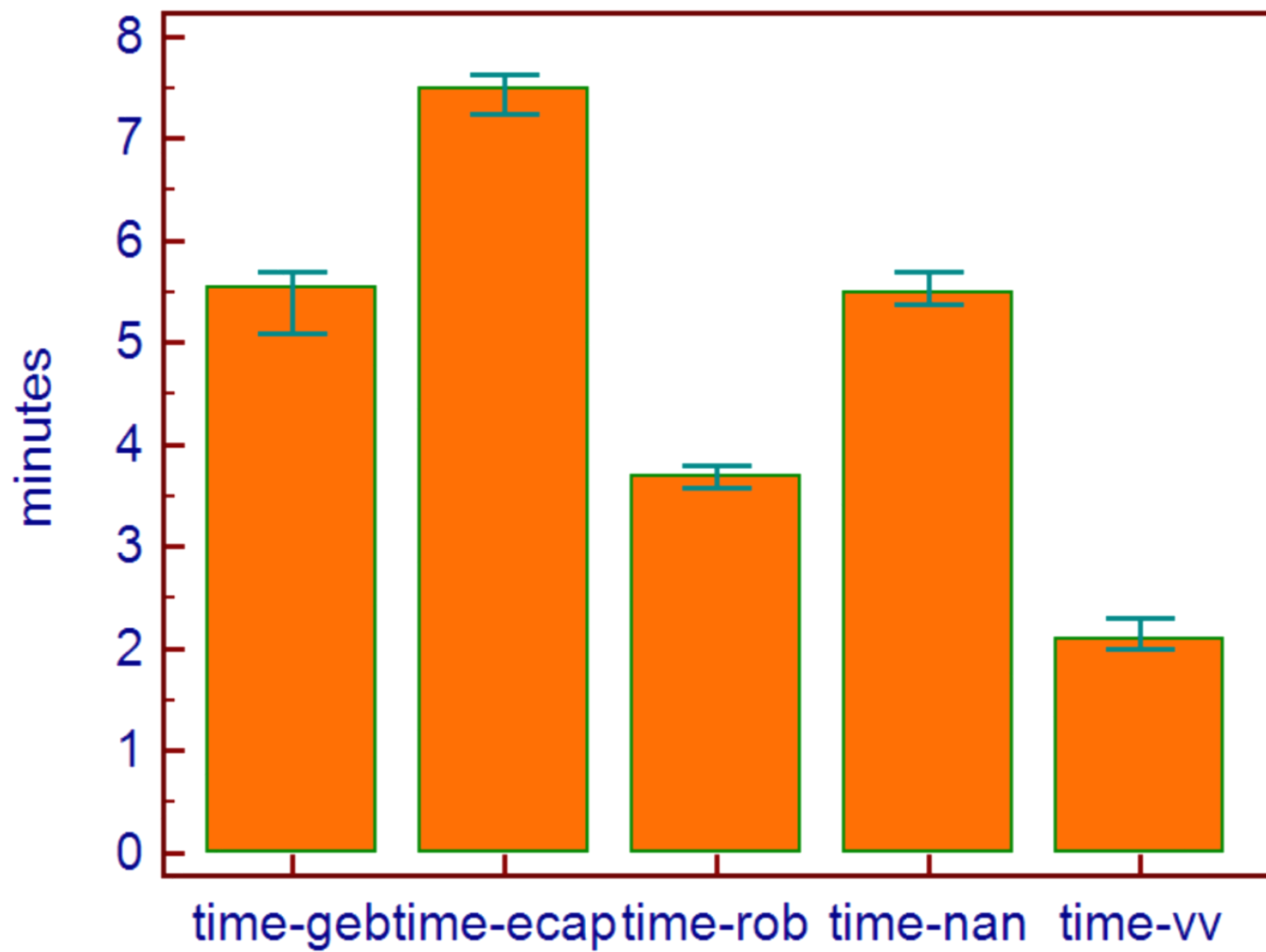
**A REAL LIFE STUDY**

**Scientific Reports 28-July-2017**



**Complete biopsy sampling (at least four samples from the terminal ileum, and at least two samples from cecum, ascending colon, transverse colon, descending colon, sigmoid, and rectum) pre- and post-therapy. Thus, usually having three slides (one for terminal ileum, one from the cecum to the descending, one for the sigmoid and the rectum), a total of 144 slides (72 pre- and 72 post-therapy) was evaluated for a total of 384 biopsies**









Pericolo  
scampato !!



His  
unfulfilled promise

Vincenzo Villanacci, Elisabetta Antonelli, Karel Geboes, C



## CONCLUSIONS





# **Remember.....**

**“...he that increaseth knowledge  
increaseth sorrow ”**

**Ecclesiastes, 1:18**



SIMPLIFIED SCORE		
Patient	Morphological element	Grade
#1		
	Crypt Abscess	Absent
		Present
	Erosions and ulcerations	Absent
		Present
	Active Inflammatory Infiltrate in Lamina Propria	Absent
		Low (1)
		Moderate (2)
		Severe (3)
		Ileum
	Sites involved from disease	Right Colon
		Transverse Colon
		Left Colon
		Rectum

abnormalities.





**DO YOU HAVE ANY QUESTIONS?**