

# Best of UEGW 2014 Poster Session on Ulcerative Colitis

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# INCIDENCE AND RISK FACTORS OF C. DIFFICILE INFECTION IN ULCERATIVE COLITIS

## Introduction:

IBD patients esp. UC are at increased susceptibility for Clostridium difficile infection (CDI)

## AIMS & METHODS:

Assess the incidence and risk factors for CDI in UC patients in a tertiary center from Romania.

***Prospective evaluation*** of all UC patients admitted between January 2012 and October 2013 ( 22 Months )

# INCIDENCE AND RISK FACTORS OF C. DIFFICILE INFECTION IN ULCERATIVE COLITIS

**Results:** 70 patients with UC included  
8/70 found to have concomitant CDI ( 11.8%)

## ***Univariate analysis***

Age > 65 years (OR = 1.53, CI= 0.93-16.27; p = 0.048)

Male gender (OR = 1.38, CI= 0.30-14.91; p = 0.032)

Hemoglobin < 9 g/dL (OR = 1.93, CI = 0.19-18.52; p = 0.043)

## ***Multivariate analysis***

*Severe UC disease (OR = 1.22, CI = 0.14-10.5; p = 0.037)*

*Serum albumin < 3 g/dL (OR = 1.86, CI = 1.12- 10.14; p = 0.012)*

## **Conclusion:**

- CDI was detected in ***one of eight patients*** admitted with a UC flare
- ***Severe UC disease and low serum albumin*** were independent risk factors for CDI.

# SYSTEMATIC ANALYSIS OF FACTORS ASSOCIATED WITH PROGRESSION AND REGRESSION OF ULCERATIVE COLITIS IN THE SWISS IBD COHORT STUDY

## INTRODUCTION:

What is the Natural History of Ulcerative Colitis disease location over time?  
Risk factors associated with progression or regression of disease extent.

## AIMS & METHODS:

***Retrospective Study*** to assess disease location over time and to evaluate associated risk factors.

Data from the Swiss IBD cohort study were analyzed.

Patients were recruited from university centers (68%), regional hospitals (14%), and private practices (18%).

Disease locations over time were analyzed and risk factor analysis for a change in disease location was performed using logistic regression modeling.

# SYSTEMATIC ANALYSIS OF FACTORS ASSOCIATED WITH PROGRESSION AND REGRESSION OF ULCERATIVE COLITIS IN THE SWISS IBD COHORT STUDY

## RESULTS:

1,016 UC patients

### *Disease locations at diagnosis:*

199 (19.6%) proctitis  
338 (33.3%) left sided colitis  
381 (37.5%) extensive colitis/pancolitis  
98 (9.6%) unknown

### *During a median of 9 [5-16] years disease duration:*

**Progression** was documented in 145/1016 (14.3%)  
**Regression** in 176/1016 (17.3%)  
**Stable disease** location 624/1016 (61.4%)

# SYSTEMATIC ANALYSIS OF FACTORS ASSOCIATED WITH PROGRESSION AND REGRESSION OF ULCERATIVE COLITIS IN THE SWISS IBD COHORT STUDY

## *Factors associated with disease progression presenting with proctitis or left-sided UC at diagnosis:*

Tx Systemic steroids (OR 2.077, 95% > CI 1.359-3.174,  
Tx Immunomodulators (OR 1.647, 95% > CI 1.119-2.424,  
Tx TNF-antagonist(s) (OR 1.668, 95% > CI 1.077-2.581,  
Tx Calcineurin-inhibitors (OR 3.159, 95% > CI 1.679-5.943)

*p = 0.001*

*p = 0.011*

*p = 0.022*

*p < 0.001*

## *What was not associated with disease progression?*

Gender, age at UC diagnosis, BMI, presence of extraintestinal manifestations, smoking status at diagnosis, positive UC family history, nor 5-ASA treatment

## *How about regression in UC patients with extensive colitis/pancolitis or left-sided colitis at diagnosis?*

No specific factors were found to be associated

# SYSTEMATIC ANALYSIS OF FACTORS ASSOCIATED WITH PROGRESSION AND REGRESSION OF ULCERATIVE COLITIS IN THE SWISS IBD COHORT STUDY

## CONCLUSION:

1) Over a median of 9 years disease duration:

**2/3** UC patients maintained the initial disease location

**1/3** either had a progression / regression of the initial disease location.

2) Treatment with systemic steroids, immunomodulators, TNF-antagonists, or calcineurin-inhibitors was **significantly associated** with disease progression

# THE IMPACT OF MUCOSAL HEALING ON SUBSEQUENT CLINICAL COURSE IN THE MANAGEMENT OF ULCERATIVE COLITIS: A PROSPECTIVE OBSERVATIONAL STUDY

## ■ INTRODUCTION:

Mucosal Healing ( MH) is achieved medically in UC.

What are the Clinical Implications of MH in patients with UC?

## ■ AIMS & METHODS:

- **Prospective** Study from Japan -----112 UC patients:

- Clinical remission with medical treatment (5-ASAs, corticosteroids, leukocytapheresis, immunosuppressants, and/or biologics)
- Endoscopic examination
- MH was defined as a Mayo endoscopic subscore of either:
  - 0 = no lesions
  - 1 = mild activity
- All patients were followed up for > 1 year



# THE IMPACT OF MUCOSAL HEALING ON SUBSEQUENT CLINICAL COURSE IN THE MANAGEMENT OF ULCERATIVE COLITIS: A PROSPECTIVE OBSERVATIONAL STUDY

## RESULTS:

112 patients with UC:

62 (55%) ----Mucosal Healing ( MH)

50 (45%) ----No MH

1-year follow-up:

74 patients (66%) maintained clinical remission

38 patients (34%) relapsed

Clinical remission rate-----  **$p = 0.00001$**

MH -----(52/62, 84%)

No MH -----(22/50, 44%)

5-ASA & Clinical Remission rate -----  **$p = 0.002$**

MH----- (25/32, 78%)

No MH----- (13/32, 41%)

Immunosuppressive drugs and/or biologic agents---  **$p = 0.002$**

MH (27/30, 90%)

No MH (9/18, 50%)

THE IMPACT OF MUCOSAL HEALING ON SUBSEQUENT  
CLINICAL COURSE IN THE MANAGEMENT OF ULCERATIVE  
COLITIS: A PROSPECTIVE OBSERVATIONAL STUDY

CONCLUSION:

Patients who achieve ***clinical remission with MH*** have a ***reduced risk of future clinical relapse*** as compared with those without MH in the management of UC.

# CYCLOSPORIN A IN ACUTE STEROID-REFRACTORY OR DEPENDENT ULCERATIVE COLITIS: A PROSPECTIVE STUDY ON LONG TERM OUTCOME

## INTRODUCTION:

In severe corticosteroid refractory ulcerative colitis (UC), cyclosporine A (CsA) or infliximab (IFX) are advantageous to avoid colectomy, and their effectiveness is comparable (**Laherie et al. Lancet 2012**)

## AIMS & METHODS:

- **Prospective study** on use, efficacy, and safety of CsA in acute UC
- Moderate to severe UC refractory, intolerant to or dependent on corticosteroids and treated with CsA between Jan-2007 and Dec- 2009
- Inclusion criteria:
  - Active ulcerative colitis with a total Mayo score of 6 to 12
  - Moderate-to-severe active disease at colonoscopy
- Patients' outcome and adverse effects were followed-up for 3 years or until colectomy as the major endpoint.
- Clinical and laboratory data were collected at the beginning of CsA therapy and at routine scheduled visits.

# CYCLOSPORIN A IN ACUTE STEROID-REFRACTORY OR DEPENDENT ULCERATIVE COLITIS: A PROSPECTIVE STUDY ON LONG TERM OUTCOME

## RESULTS:

- *Colectomy-free survival at 3 years* was 72.1 % (95% CI 60.9 – 83.4).
- Number of colectomies during the 3-year follow-up period was 17 (27.9 %)
- All colectomised patients had disease duration under 5 years
- *Short disease duration was the only independent risk factor for colectomy* in multivariate regression analysis
- Adverse events were registered in 7 patients:
  - CMV infection
  - Pneumocystis jirovecii infection
  - Clostridium difficile colitis
  - Transaminase elevation
  - Venous thrombosis

# **CYCLOSPORIN A IN ACUTE STEROID-REFRACTORY OR DEPENDENT ULCERATIVE COLITIS: A PROSPECTIVE STUDY ON LONG TERM OUTCOME**

## **CONCLUSION:**

- 1) CsA treatment was successful in 72% of patient who avoided colectomy
- 2) CsA as good option in moderate-to-severe acute corticosteroid refractory or corticosteroid resistant thiopurine naive UC.
- 3) Short disease duration was independent risk factor for colectomy.
- 4) Risk of infection under high immunosuppression must be remembered.

# PREVENTION OF OPPORTUNISTIC INFECTIONS IN PATIENTS ON BIOLOGICAL AGENTS FOR MANAGEMENT OF INFLAMMATORY BOWEL DISEASE

## INTRODUCTION:

- Patients with IBD are at increased risk of infection; this is especially true of the 20% on biological agents.
- **ECCO guidelines** recommend the following vaccines:
  - Influenza (annual)
  - Pneumococcal
  - Hepatitis B
  - Varicella
  - HPV (women under 26)
  - Need to exclude latent TB; local policy is to perform an interferon gamma release assay.
- Within the UK vaccination services are provided by primary care.

# PREVENTION OF OPPORTUNISTIC INFECTIONS IN PATIENTS ON BIOLOGICAL AGENTS FOR MANAGEMENT OF INFLAMMATORY BOWEL DISEASE

## AIMS & METHODS:

- Measures to prevent opportunistic infections in patients prescribed anti-TNFs for IBD at 2 UK Hospitals in 2013 were **compared against** the ECCO (OI) guidelines.
- The following were retrieved from electronic records:
  - Age
  - Sex
  - Anti-TNF prescribed
  - Pneumococcal antibodies
  - Hepatitis B core and surface antibodies
  - Varicella IgG
- Attempts were made to retrieve vaccination history from General Practice.

# PREVENTION OF OPPORTUNISTIC INFECTIONS IN PATIENTS ON BIOLOGICAL AGENTS FOR MANAGEMENT OF INFLAMMATORY BOWEL DISEASE

## RESULTS:

60 patients ---- Infliximab

15 patients -----Adalimumab.

**Influenza: 50% (23/46) patients received vaccination against influenza** within the past year

**Pneumococcus: 55% (47/85) patients demonstrated immunity.** 6% (5/85) were not immune and the remainder were not tested. The vaccination history of 26 patients who were not immune or not tested was retrieved. 27% (7/26) had since been vaccinated.

**Hepatitis B:** No patients were core Ab positive. **Surface Ab levels demonstrated immunity in 7% (6/85). 53% (45/85) were not immune, and the remainder were not tested.** Vaccination history of 44 patients who were not immune or not tested was retrieved. Of these, 25% (11/44) had since been vaccinated.

**HPV:** 4 patients were women under 26 years old. **25% (1/4) had confirmed HPV vaccination.**

**Varicella: 21% (18/85) patients demonstrated immunity to varicella.** 2% 2/85 were not immune.



# PREVENTION OF OPPORTUNISTIC INFECTIONS IN PATIENTS ON BIOLOGICAL AGENTS FOR MANAGEMENT OF INFLAMMATORY BOWEL DISEASE

## CONCLUSION:

1) The standards set out by ECCO to protect patients from opportunistic infection are *not being met.*

2) *Potential service improvements include:*

- Provision of vaccines at clinic
- Improved patient education regarding the importance of vaccination
- A check list to review bloods at first anti-TNF prescription

# RARITY OF ADENOMATOUS POLYPS IN ULCERATIVE COLITIS: IMPLICATIONS FOR COLONIC CARCINOGENESIS

## INTRODUCTION:

- Ample research on dysplasia-carcinoma risk in ulcerative colitis (UC)
- Scant data on ***sporadic adenomas' risk in this population.***

## AIMS & METHODS:

- What is the prevalence of sporadic colon adenoma in UC patients?
- Is there a role of ***chronic immune-driven inflammation*** on adenoma development?
- Number and histology of all polyps detected at colonoscopies of UC patients during 2006-2012 were compared to controls undergoing screening colonoscopy

**Group I:** Patients who were over 50 years-old at the time of the index colonoscopy to reinforce the validity of the comparison to screening colonoscopy controls.

**Group II:** To exclude a potential bias, an additional analysis was performed including all prior colonoscopies undergone by the UC group.

**Group III:** Crohn's disease patients was also evaluated to dissect the role of colonic IBD versus ileal IBD on sporadic adenoma rate.

# RARITY OF ADENOMATOUS POLYPS IN ULCERATIVE COLITIS: IMPLICATIONS FOR COLONIC CARCINOGENESIS

## RESULTS:

206 UC patients vs. 624 controls

### I : Adenomatous polyps

- 13/206 UC colonoscopies
- 162/624 colonoscopies of controls  
(**6.3% vs. 25.9%** respectively, OR 0.19, 95% CI 0.1-0.34, **p < 0.0001**)

### II: Considering all prior colonoscopies

(**14.1% vs. 25.9%** respectively, OR 0.47, 95%CI 0.3- 0.72, **p = 0.0005**)

### III: Among 115 Crohn's patients >50 years old

- Rate of adenomas in Crohn's ileitis patients and controls was similar  
(**p = 0.8**)
- Crohn's disease involving the colon had significantly lower rate of adenomas compared to controls  
(**3.9% vs. 25.9%, p = 0.002**)

# RARITY OF ADENOMATOUS POLYPS IN ULCERATIVE COLITIS: IMPLICATIONS FOR COLONIC CARCINOGENESIS

## CONCLUSION:

- 1) Patients with UC or colonic Crohn's disease ***seldom develop sporadic adenomatous polyps***
- 2) These data provide novel insight into possible mechanisms restricting the adenoma-carcinoma sequence and suggest ***organ-specific immune activation*** may confer protection against development of colonic adenomas.

