



Azienda Ospedaliero - Universitaria di Parma



Changes in antibiotic sensitivity in children with H. pylori infection over 13 years

Marco Manfredi MD, PhD

Pediatrician & Gastroenterologist
"Pietro Barilla" Children's Hospital
University of Parma, Parma, Italy





World Congress and Exhibition on **Antibiotics**

September 14-16, 2015 Las Vegas, Nevada, USA

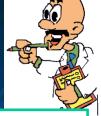






September 14-16, 2015 Las Vegas, Nevada, USA





Hindawi Publishing Corporation Gastroenterology Research and Practice Volume 2015, Article ID 717349, 5 pages https://dx.doi.org/10.1155/2015/717349



Research Article

Primary Antimicrobial Susceptibility Changes in Children with Helicobacter pylori Infection over 13 Years in Northern Italy

Marco Manfredi, ¹ Pierpacifico Gismondi, ¹ Valentina Maffini, ¹ Barbara Bizzarri, ² Fabiola Fornaroli, ² Carmen Madia, ¹ Antonino Salerno, ¹ A. Marta Cangelosi, ¹ and Gian Luigi de'Angelis^{1,2}

no conflict of interest





September 14-16, 2015 Las Vegas, Nevada, USA





Helicobacter pylori infection

40%-50% developed countries

GI diseases

80% developing countries

Chronic Gastritis





Gastric Cancer

Eradication



Risk of developing GC

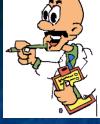
European, Asian, American Guidelines





September 14-16, 2015 Las Vegas, Nevada, USA





Many efforts have been made in trying

Antibiotic Resistances

Several difficulties

ercome

Triple therapy



the main eradication therapy

Widespread use/abuse antibiotics (respiratory tract infections)







September 14-16, 2015 Las Vegas, Nevada, USA





Effectiveness of triple therapy



mainly due to primary resistance towards antibiotics



clarithromycin over the past 10 years

9.8%

17.5%

Maastricht



clarithromycin

in empirical eradication therapy



testing susceptibility for this antibiotic

before using it if its resistance prevalence is higher than 15-20%







September 14-16, 2015 Las Vegas, Nevada, USA







ESPGHAN/NASPGHAN guidelines



antibiotic susceptibility testing for clarithromycin

before starting clarithromycin-based triple therapy

in areas/populations with a known high resistance rate (>20%)

In Italy the clarithromycin resistance is around 25% both in adults and children

Megraud et al, Gut 2013







September 14-16, 2015 Las Vegas, Nevada, USA





Culture

common & specific

invasive diagnostic method

antimicrobial susceptibility of *H. pylori*

Thijs JC et al, Am J Gastroenterol 1996

sensitivity



variable (60–90%)



several methodological factors

time from sampling to processing

biopsy site



incubation conditions

Marco Manfredi MD, PhD

of Inflammatory and Infectious Diseases

culture medium





September 14-16, 2015 Las Vegas, Nevada, USA





culture of a single antral biopsy



the detection of H. pylori in > 90%

for optimal results

at least

One biopsy

antrum

Two biopsies

corpus

Dixon MF et al, Am J Surg pathol, 1996

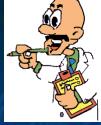
to ensure a correct diagnosis





September 14-16, 2015 Las Vegas, Nevada, USA





in *H. pylori* eradication the most common antibiotics used

amoxicillin

clarithromycin

metronidazole

in children

ESPGHAN-NASPGHAN Guidelines, 2



Looking for the best eradication therapy

if the antimicrobial susceptibility is not available



we should ask to patients.

which



how many antibiotics



they used before

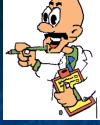
Marco Manfredi MD, PhD





September 14-16, 2015 Las Vegas, Nevada, USA





We wanted to evaluate the variations in primary antibiotic susceptibility over last 13 years in children with *H. pylori* infection in Parma, northern Italy comparing with our previous results obtained in 2001

Street ME, .. Manfredi M, et al, Arch Dis Child 2002

JANUARY 2011							
MON	TUES	WED	THURS	FRI	SAT		
31					1		
3	4	5	6	7	8		
10	11	12	13	14	15		
17	18	19	20	21	22		
24	25	26	27	28	29		
	31 3 10 17	MON TUES 31 3 4 10 11 17 18	MON TUES WED 31 3 4 5 10 11 12 17 18 19	MON TUES WED THURS 31 3 4 5 6 10 11 12 13 17 18 19 20	MON TUES WED THURS FRI 31 3 4 5 6 7 10 11 12 13 14 17 18 19 20 21		



DECEMBER 2012								
MON	TUE	WED	THU	FRI	SAT	SUN		
					1	2		
3	4	5	6	7	8	9		
10	11	12	13	14	15	16		
17	18	19	20	21	22	23		
24	25	26	27	28	29	30		
31								

we diagnosed by endoscopy and histological examination

66 naïve children with *H. pylori* infection





September 14-16, 2015 Las Vegas, Nevada, USA



A 17 11 15 15 15 16		730534557168		
Antibiotic	MIC interpretative values(μg/ml)	Resistance Rate (years 1998-99)	Resistance Rate (year 2011-12)	s p Value
Metronidazole	>16	35 (56%)	15 (33%)	0.014
Clarithromycin	>4	10 (16%)	12 (26%)	0.142
Ampicillin	>4	2 (3%)	0	ns
Tetracyclines	>16	1 (2%)	0	ns
Metronidazole + clarithromycin		5 (8%)	3 (7%)	ns
Total		63/75 (84%)	46/66 (70%)	0.079

Table 3. Differences in antibiotic resistances.

Manfredi M. et al, Gastr Res Pract 2015





September 14-16, 2015 Las Vegas, Nevada, USA





The eradication of *H. pylori* infection represents an enormous challenge in gastroenterology

H. pylori is not so easily attachable

importance

local prevalence of antibiotic resistances

Better therapy

mainly if the antimicrobial susceptibility



therefore we must use an empirical eradication treatment

Megraud F et al, Gut 2013





September 14-16, 2015 Las Vegas, Nevada, USA





Culture

 \rightarrow

Useful, but disadvantages

Difficulty

Sensitivity

High costs

Our study shows a decline of culture development of *H. pylori* over the last 13 years: from 84% in 1998/99 to 70% in 2011/12 with no statistically significant value

Three biospy samples RUT positive

Culture can be negative 62% culture pos

Porowska B et al Aliment Pharmacol Ther 2012









On the contrary

Other italian researchers obtained a very high culture development rate of 94% in adults

Saracino IM et al, J Gastrointestin Liver Dis 2012

main cause of culture failure Chronic PPI intake

biopsy specimens should also be taken from the gastric body

Culture development

Intrinsic difficulty





September 14-16, 2015 Las Vegas, Nevada, USA







clarythromicin .

In empirical therapy

In areas with R > 20%

A good alternative choices

Bismuth-containing quadruple therapy

(B+A+C+M)

Sequential therapy

Horvath A et al, Aliment Paharmacol Ther 2012 Francavilla R et al, GUT 2008





September 14-16, 2015 Las Vegas, Nevada, USA





Conclusion 1

H. pylori culture

could now be substituted by molecular methods

FISH

biopsies

PCR

do not require strict conditions of biopsy specimens transport

can be used to detect the point mutation associated to clarythromicin-resistance





September 14-16, 2015 Las Vegas, Nevada, USA





Conclusions 2

Before recommending *H. pylori* eradication therapy

we should know

either

the antibiotic susceptibility of patient

or

the local distribution of antibiotic resistance rates

to have higher successful probabilities

Helicobacter pylori

Detection Methods, Diseases and Health Implications



Marco Manfredi Gian Luigi de'Angelis Editors



www.novapublishers.com

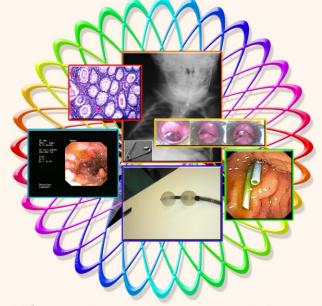
N o v a

В

o m e d

> c a

Marco Manfredi • Gian Luigi de'Angelis



Gastrointestinal Endoscopy in Children

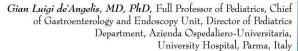


Gastrointestinal Endoscopy in Children





Marco Manfredi, MD, PhD, Pediatrician and Gastroenterologist, "Pietro Barilla" Children's Hospital, Azienda Ospedaliero-Universitaria, University Hospital, Parma, Italy











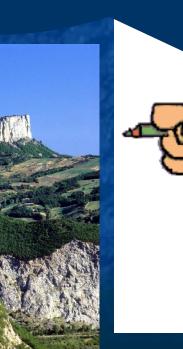


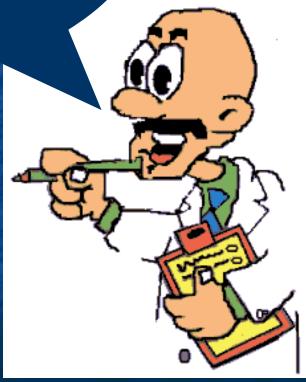


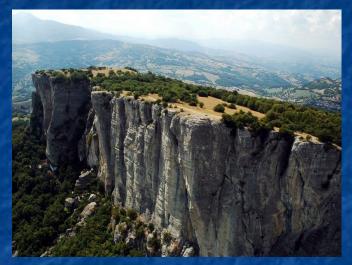
September 14-16, 2015 Las Vegas, Nevada, USA



Thank you for your attention













September 14-16, 2015 Las Vegas, Nevada, USA









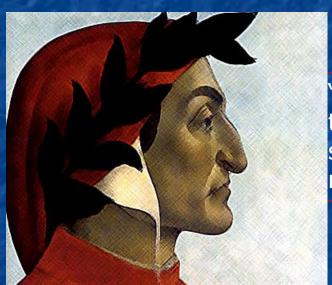


September 14-16, 2015 Las Vegas, Nevada, USA





Dante Alighieri drew inspiration for the mountain of Purgatory



"...You can walk at Sanleo, near Urbino, and descend to Noli, near Savone: you can climb Mount Bismantova, south of Reggio, up to the summit, on foot: but here a man had to fly..."

> Dante Alighieri Divina Commedia Purgatory Canto, IV, 26







September 14-16, 2015 Las Vegas, Nevada, USA



Thank you for your attention

