





DUBAI | UAE | 22 to 24 APRIL

**DUBAI WORLD TRADE CENTRE** 



MANAGEMENT OF CONSTIPATION

ASAD DAJANI MD, DSM, FACG, JBD, FRCP (GLASG) 24 APRIL 2024

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**DUBAI WORLD TRADE CENTRE** 



A focus on the use of Laxatives

EFMS

Asad Dajani MD, DSM, FACG, JBD, FRCP (Glasg)

Organized by





# Disclosures and agenda





### **Disclosures**



A member of advisory boards for Sanofi (UAE, Global), Abbott, Janssen, Lunatus, Synergy, Takeda, Novonordisk, Julphar.

A lecturer for Sanofi (UAE, Global), Abbott, Janssen, Lunatus, Synergy, Dr Falk, Takeda, Spimaco, Julphar, Tabouk, Novonordisk







- The Speaker and Abbott Laboratories has a Professional service agreement in place as of 19/4/2024 and this presentation is made based on the said Agreement in place between the Speaker and Abbott Laboratories.
- This presentation is not meant to promote any healthcare or pharmaceutical product or services.



# Agenda

- Why constipation?
- Pathophysiology of constipation
- Clinical picture and diagnosis
- Treatment concepts (Focus on Laxatives)
- Conclusion and guidelines



# Constipation is an aggravating, but not life-threatening or debilitating, complaint

• The most common digestive complaint and a health burden with negative impact on quality of life.





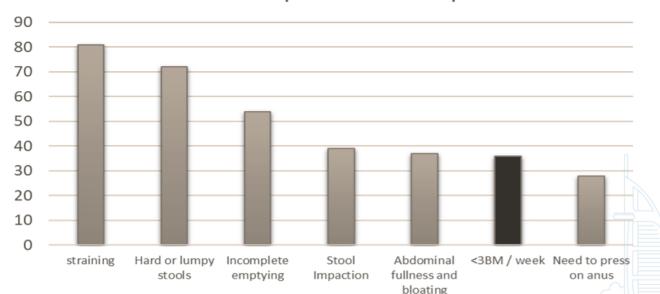
#### Constipation as defined in real world clinical practice.

#### Patient's perspective:

Patients usually define constipation by symptoms (e.g. straining, bloating and hard stools)



#### Patient description of constipation



Healthcare providers often use **bowel movement (BM) frequency** (e.g. < 3 BM per week)

Herz et al. Fam Pract 1996;13(2): 156-9

Cash BD & Chey WD. Pharmacol Ther 2005;22:1047-1060.



#### **Constipation Occasional or Chronic.**

#### **Occasional**

- if acute or subacute constipation occur in a middle-aged or elderly patients, for Less than 6 weeks.
- should prompt a search for an obstructing colonic lesion.

#### **Chronic**

• if it occurred for at least 12 weeks (in total, not necessarily consecutively) during the previous year.



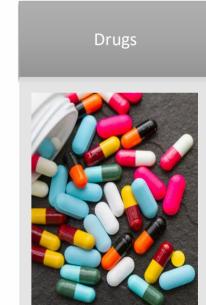
#### **Constipation: Primary or secondary**

#### **Medications**

- -Analgesics:
  - **ONSAIDs**
  - ○Opioids
- -Antihypertensive agents:
  - Diuretics
  - Calcium channel blockers

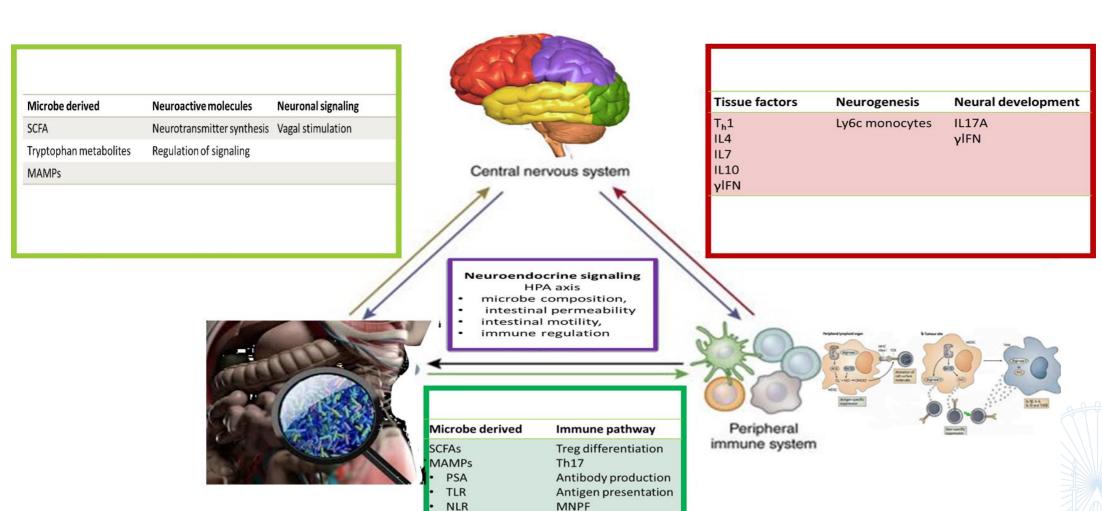


- -Antidepressants
- -Antihistamines
- -Antiparkinson agents
- -Metallic ions





#### **Constipation: A role for dysbiosis.**





# Rome IV: Five distinct categories of Functional GI diseases

- ☐ A functional bowel disorders (FBD) in which recurrent abdominal pain is associated with defecation or a change in bowel habit: constipation, diarrhea, or a mix.
- □ Abdominal bloating/distention.
- □ Symptom onset should occur at least 6 months before diagnosis and symptoms should be present during the last 3 months.

Significant overlap exists between these disorders. These disorders should be thought of as existing on a continuum, rather than as in isolation. A patient with IBS (right) will have symptoms of abdominal pain, in contrast to a patient with FC or FDr, who does not have abdominal pain. Bloating and distention are common symptoms frequently reported by patients with any FBD.



#### **Prevalence of chronic functional Constipation**

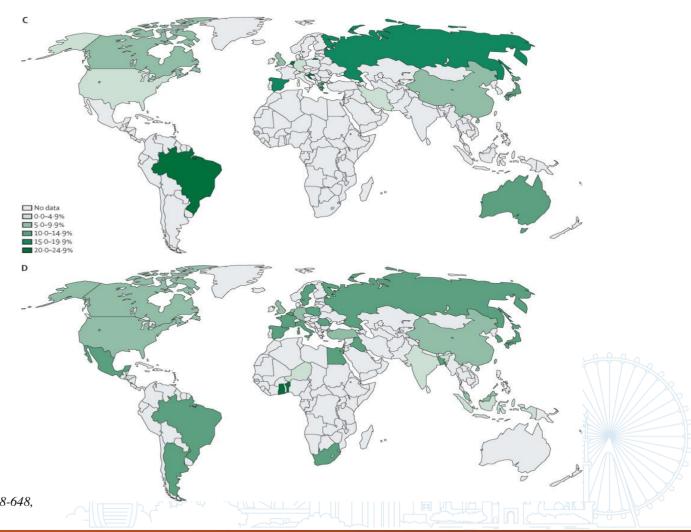
#### Systemic review & Network meta-analysis

8174 citations evaluated, 45 studies fulfilled the eligibility criteria, representing 80 separate populations and comprising 275 260 participants

Criteria	95% CI
Rome I	15·3% (8·1–24·4, <i>I</i> <sup>2</sup> =99·4%)
Rome II	11·2% (7·9–14·9; <i>l</i> ²=99·6%)
Rome III	10·4% (6·5–14·9; <i>l</i> ²=99·8%)
Rome IV	10·1% (8·7–11·6; <i>l</i> ²=98·2%)

#### Functional constipation

- May occur at any age group
- More prevalent in females
- Prevalence is comparable with other chronic diseases





#### Prevalence of Chronic Constipation in pregnancy.



- > The prevalence of constipation in pregnancy ranges from 11% to 44%<sup>1</sup>.
- ➤ Up to 40% of women may suffer symptoms of constipation at some stage during their pregnancy².
- ➤ Evidence suggests that a great number of women experience constipation up to 3 to 6 months postpartum and may even persist to 12 months post-delivery¹.
- $\triangleright$  The prevalence of postpartum constipation was estimated to be 24% at 3 months postpartum<sup>2</sup>.

Trimester	Prevalence of constipation
First trimester	24%
Second trimester	16%
Third trimester	26%
Postpartum	24%

Cesarean section rate in pregnant women with functional constipation (66.97%) is higher than in the other methods of delivery (27.29%)<sup>3</sup>.

<sup>1-</sup>Rungsiprakarn et alCochrane Database Syst Rev.2015Sep 4;(9):CD011448. 10.1002/14651858.CD011448.pub2.

<sup>2-</sup>Bradley et al. Obstet Gynecol. 2007.110(6):1351-1357.

<sup>3-</sup>Shi et al. PLoS ONE. 2015. 10(7): e0133521

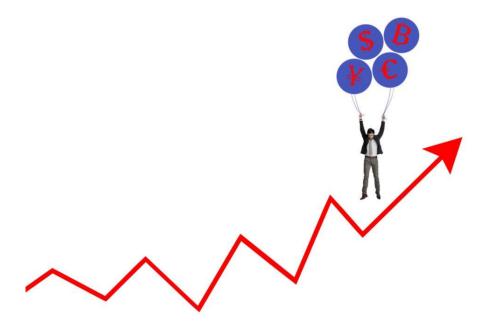
<sup>4.</sup> Turawa et al. Cochrane Database Syst Rev. 2014;(9):CD010273.

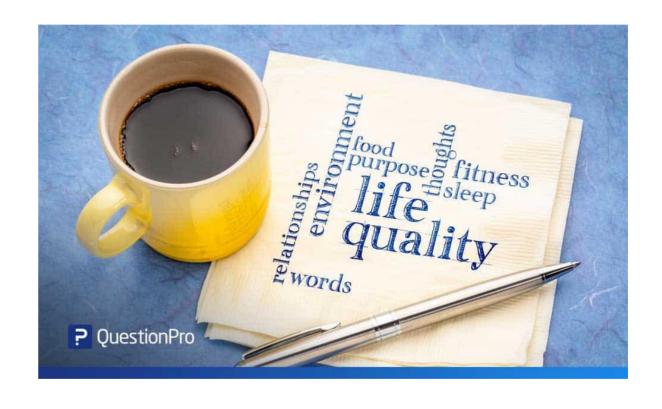


## **Comorbidities associated with Chronic Constipation**

#### **GI** Comorbidities

Gastrointestinal comorbid conditions	Chronic constipation $n = 262$ (%)	Matched controls n = 262 (%)	<i>p</i> -value <sup>a</sup>
Hemorrhoids	59 (23%)	49 (19%)	0.34
Anal fissures	7 (3%)	8 (3%)	1.00
Rectocele	10 (4%)	13 (5%)	0.66
Anal or rectal cancer	3 (1%)	2 (1%)	1.00
Fecal incontinence	2 (1%)	7 (3%)	0.18
Colon cancer	3 (1%)	5 (2%)	0.73
Enterocele/sigmoidocele	3 (1%)	8 (3%)	0.23
Diverticulosis	91(35%)	85 (32%)	0.61
Diverticulitis	13 (5%)	15 (6%)	0.85
Small bowel/colonic stricture/stenosis	2 (1%)	8 (3%)	0.11
IBS <sup>b</sup>	13 (5%)	21(8%)	0.23
Ischemic colitis	4 (2%)	3 (1%)	1.00
Microscopic colitis	0	3 (1%)	0.04
Crohn's disease	1 (0.4%)	1(0.4%)	1.00
Ulcerative colitis	1 (0.4%)	3 (1%)	0.63
Prior cancer	88 (34%)	89 (34%)	1.00
Peptic ulcer disease	4 (1.5%)	3 (1%)	1.00
Anal surgery	6 (2%)	3 (1%)	0.51
Colonic surgery	8 (3%)	11 (4%)	0.65
Cholecystectomy	14 (5%)	15 (6%)	1.00
Abdominal hernia	7 (3%)	4 (2%)	0.55

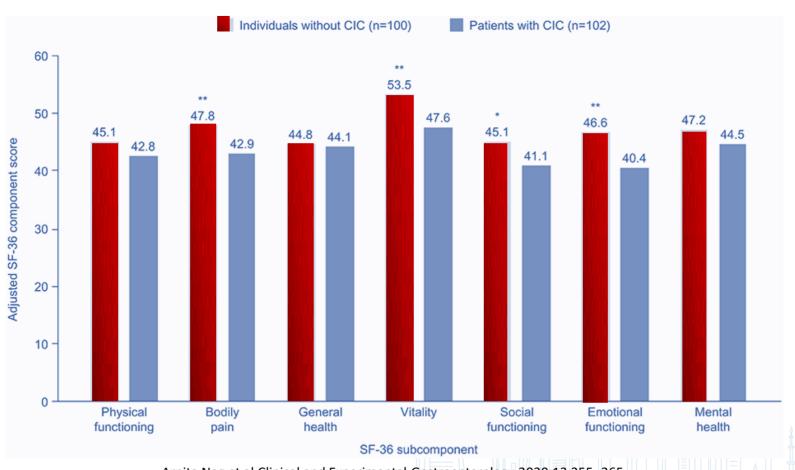




# Quality of life and economic burden.



# Chronic Constipation has a detremintal effect on HRQoL.



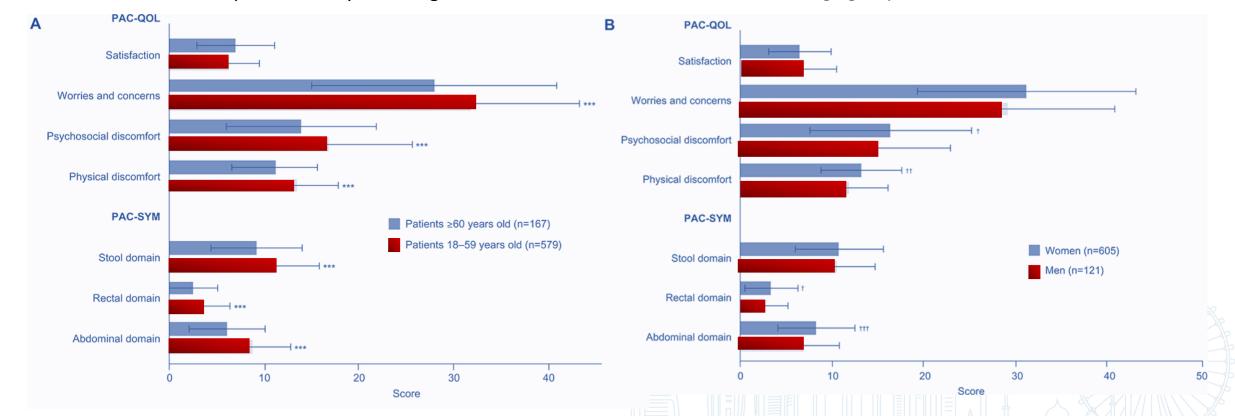
Arpita Nag et al. Clinical and Experimental Gastroenterology 2020:13 255–265



# Chronic Constipation has a detremintal effect on HRQoL.

HRQol for patients 18-59 years was more affected than patients >60 years of age

HRQoL was more affected in Women than in men at all age groups



Arpita Nag et al. Clinical and Experimental Gastroenterology 2020:13 255–265



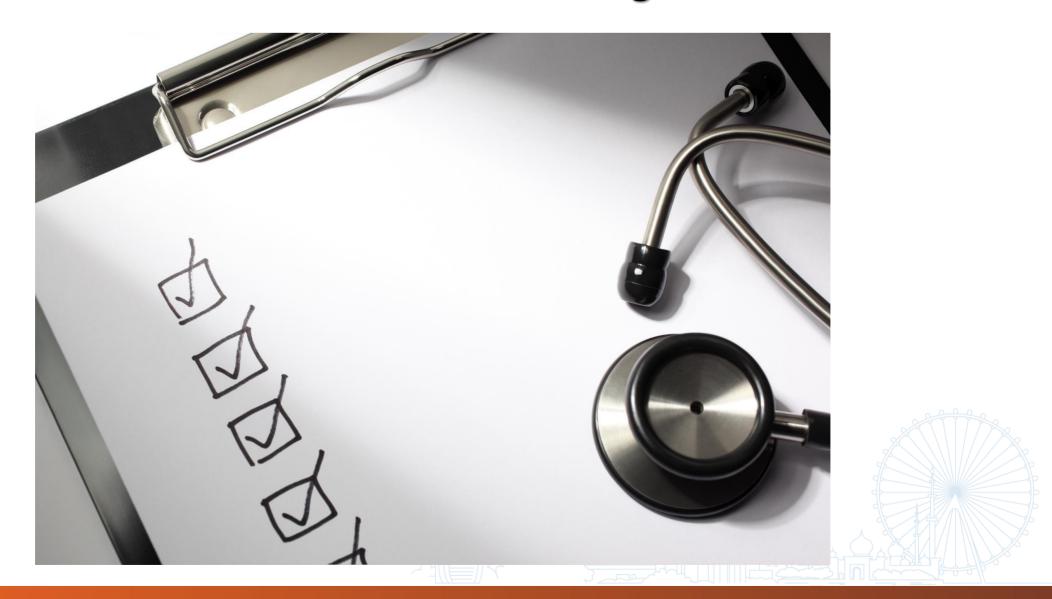
## Chronic Constipation poses a significant health burden.

Direct costs	Indirect costs
Physicians visits (2.7 mio 2001)	OTC treatment
Hospitalization (235 mio 2001)	Restricted activities
Medications (800 mio 2001)	Lost income
Procedures (3000 mio 2001)	Cost of care givers

Dennison C, et al. Pharmacoeconomics 2005;23(5):461–476



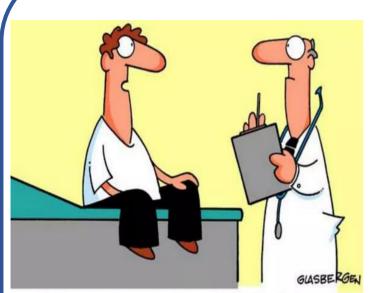
## Clinical evaluation and diagnosis.





### **Chronic Constipation: Clinical evaluation.**

#### Step 1 ... Get detailed medical history



"I already diagnosed myself on the Internet. I'm only here for a second opinion."

- 1. Ask key questions and consider ways of improving patient-physician communication.
- 2. Define patient complaints and consider if they meet the criteria of CC
- 3. Consider the red flags
- Frequency
- Straining or not?
- Time spent in the toilet ?
- Stools shape & consistency?
- Postural or digital manoeuvres to assist defecation?
- complete or incomplete evacuation?
- Use of laxatives ? Response?
- Exclude secondary causes

#### **Step 2** ... rectal examination



"Believe me, this is worse for me than it is for you."



### **Chronic Constipation: The red flags.**

**Unexplained weight loss Rectal bleeding** 

Eamily history of CDC

Constipation may be one of the earliest symptoms when CRC had developed

Rectal masses
Increased cancer markers



#### **Chronic Constipation: Step 3, diagnostic tests.**

- Anorectal manometry
- Colonic transit assessment
- Balloon expulsion test
- Defecography and MR defecography
- Wireless motility capsule test
- Colonic manometry



#### **Chronic Constipation: Balloon expulsion test.**

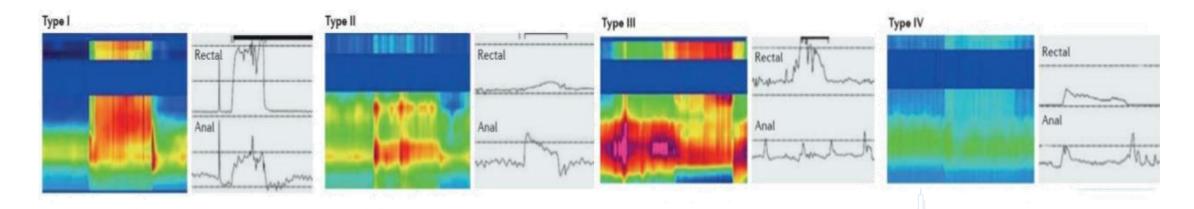
Expulsion time of >2 min is considered abnormal

High specificity (80-90%) & low sensitivity (50%) for Dyssynergia





#### **Chronic Constipation: Manometric patterns.**



Type I: adequate rectal push effort with paradoxical anal sphincter contraction

Type II: inadequate rectal push effort with paradoxical anal sphincter contraction

Type III: adequate rectal push effort but inadequate relaxation (<20%) of anal sphincter pressure

Type IV: inadequate rectal push effort and also inadequate relaxation (<20%) of anal sphincter pressure



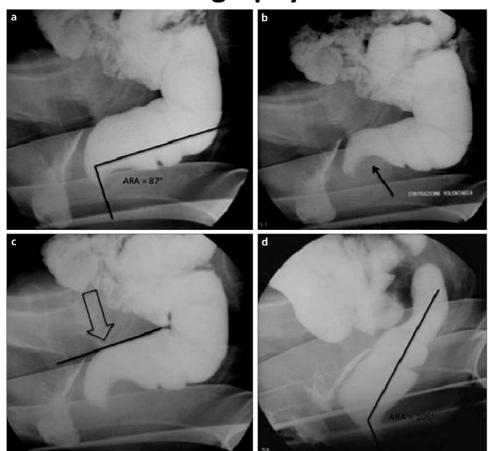
## **Chronic Constipation: Colonic transit time.**





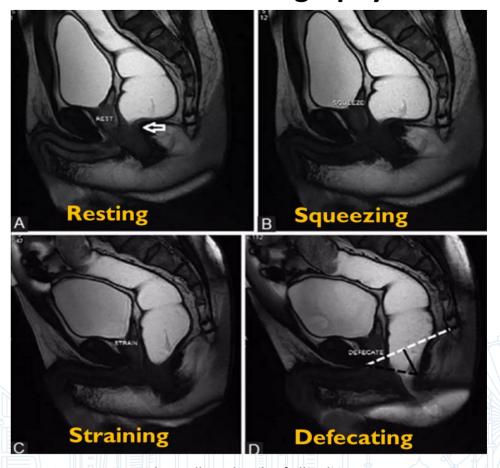
## **Chronic Constipation: Radio-imaging.**

#### **Defecography**



150 ml of barium paste is placed in the rectum & patient is asked to expel the barium in a sitting position on a special commode

#### MR defecography

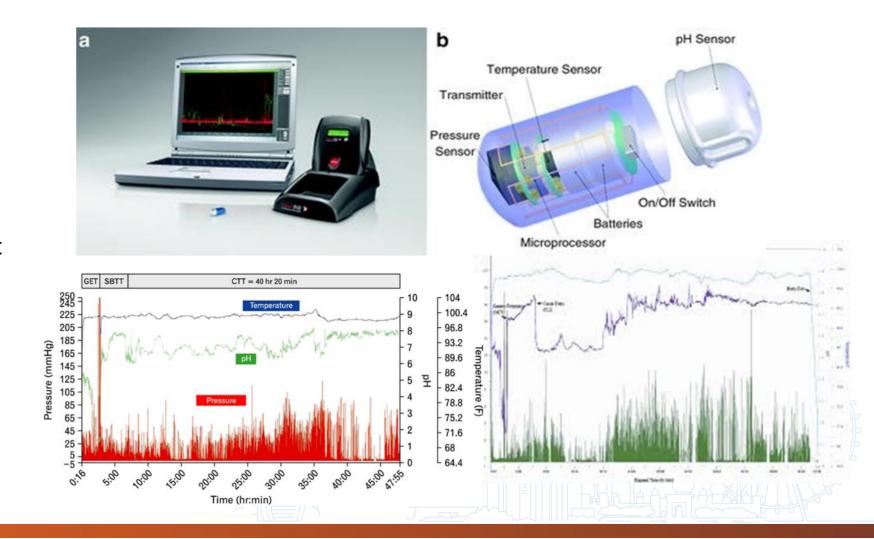


More precise with excellent details of all pelvic organs but more expensive



### Chronic Constipation: Wireless Motility Capsule (WMC).

- FDA approved
- Recommended by the American and European NGMS
- No radiation exposure
- Asses regional and whole gut transit time
- Normal colonic transit time
   <59 hrs</li>
- Whole gut transit <73 hrs</li>





# Chronic Constipation: Treatment Life style modification.

	European GL	French GL	Spanish GL	Korean GL
Lifestyle	Overall lifestyle modifications have a  positive effect  Evidence: Moderate  Recommendation: Strong  Agreement: 100%	Behavioural rules (daily presentation to the toilet, optimal position on the toilet, environmental conditions)  • Positive effect • Expert Recommendation	NA	NA
Diet	Increase of fiber intake has positive effect, especially if combined with fluid increase • Evidence: Low • Recommendation: Weak • Agreement 92%	Increase of fiber intake has positive effect. Dried plums have a better efficacy than psyllium in mild to moderate constipation  • Evidence II, Grade B  • Consumption of foods other than fiber: not positive not negative effect  • Expert Recommendation  • Overeating has positive effect only in undernourished patients  • Expert Recommendation	Consuming high fiber foods has a positive effect • Evidence: Moderate • Recommendation: Strong	Dietary fiber has a positive effect. Evidence: C Recommendation: 2 Experts' agreement: completely agree: 27.6%; mostly agree:72.4% It can be an initial strategy Evidence: C. Recommendation: 2. Experts' agreement: completely agree: 35.7%, mostly agree: 60.7%; partially agree: 3.6%
Increase of fluid intake	Positive effects only in dehydrated patients • Evidence: Low • Recommendation: Strong • Agreement: 100%	Positive effect in dehydrated patients or in those assuming fiber • Expert Recommendation • Positive effect of water rich in magnesium • Level II, Grade B	Positive effect only if associated with fiber supplement • Evidence: Low • Recommendation: Weak	Positive effect in dehydrated patients or when bulking agents are added • Evidence: C • Recommendation: 1 • Experts' agreement completely agree: 37.0%; mostly agree: 55.6%; partially agree: 7.4%
Exercise	Not positive not negative effect on constipation • Evidence: Moderate • Recommendation: Strong • Agreement: 92%	Not positive not negative effect on constipation • Expert Recommendation	Positive effect • Evidence: Low • Recommendation weak	Positive effect. • Evidence: C • Recommendation: 2 • Experts' agreement: completely agree: 7.1%, mostly agree: 67.9%, partially agree: 14.3%, mostly disagree: 10.7%

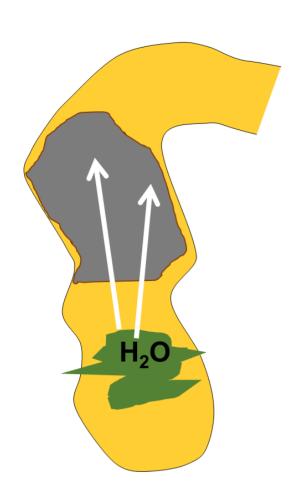


### **Medicinal Treatment.**





#### **Chronic Constipation: Bulking laxatives.**

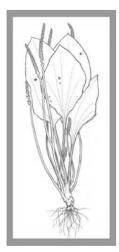


#### Absorbs 40 times its own weight in water

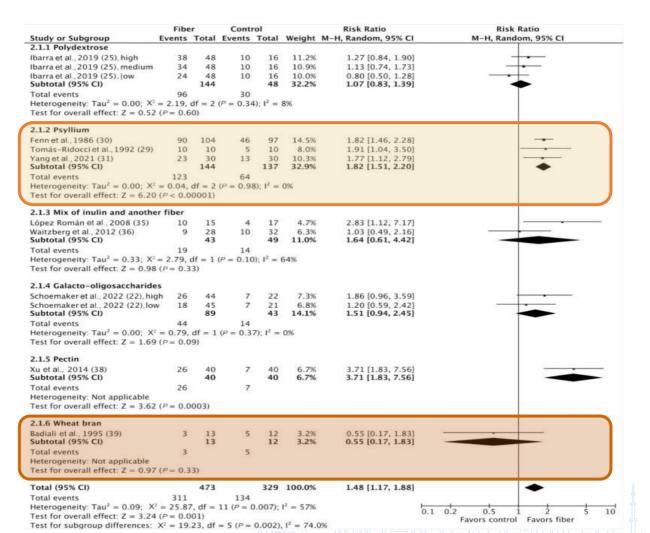
- Should be mixed in a glass of water or juice, stirred and drunk immediately followed by a half to a full glass of water.
- Insufficient fluid intake cause the drug to solidify in GI tract and result in intestinal obstruction
- may not be appropriate for patients who must restrict oral fluid intake (patients with <u>kidney</u> or <u>heart failure</u>).
- Patients with narrowing of the digestive tract (including esophageal stricture, intestinal stricture, or severe adhesions) may be exposed to the risk of blockage of the intestine or the esophagus.
- May cause abdominal distention



### Chronic Constipation: Bulking laxatives, NW meta-analysis.



Plantago Ovata







## **Chronic Constipation: Osmotic laxatives.**



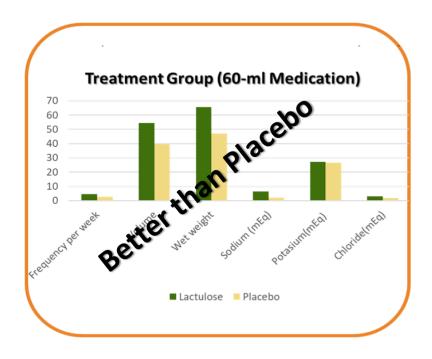
# Chronic Constipation: Osmotic laxatives, Meta-analysis of high quality studies.

	La	Laxatives Placebo					Mean Difference		Mean Difference		
Study or Subgroup	Mean	Mean SD		Mean	SD	Total	Weight	IV, Random, 95% CI	Year	IV, Random, 95% CI	
1.2.1 Osmotic laxativ	es										
Baldonedo 1991	13.56	6.74	16	5.53	3.58	15	5.4%	8.03 [4.26, 11.80]	1991		<b>─</b>
Corazziari 1996	4.8	2.3	25	2.8	1.6	23	17.0%	2.00 [0.89, 3.11]	1996		
DiPalma 2000	4.5	3	80	2.7	1.8	71	18.9%	1.80 [1.02, 2.58]	2000		-0-
DiPalma 2007 Subtotal (95% CI)	7.9	4.5	204 325	5.6	5.5	100 209	16.2% <b>57.5</b> %	2.30 [1.06, 3.54] 2.51 [1.30, 3.71]	2007		•
	7-212/					12					
Test for overall effect:	Z= 3.12 (I	P = 0.0	02)	325		209					
Test for overall effect: Total (95% CI)	Z= 3.12 (I	P = 0.0	02) <b>805</b>	325			100.0%	2.55 [1.53, 3.57]			•



#### **Chronic Constipation: Osmotic laxatives, Lactulose.**

Double-blind drug treatment period comparing lactulose with placebo.



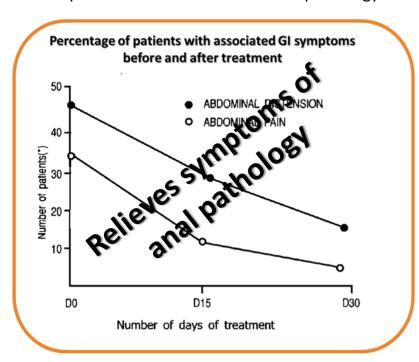
Similar to PEG





#### **Chronic Constipation: Osmotic laxatives, Lactulose.**

Open study conducted on outpatients with constipation associated with an anal pathology



Lactulose is good after of anal sphincter

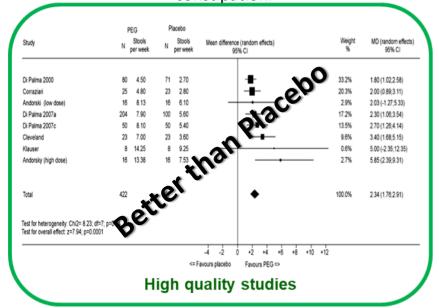
Better than Placebo for postipation

Huang, P., et al.. Journal of Biological Regulators and Homeostatic Agents, 30(2), 523-5



# Chronic Constipation: Osmotic laxatives, Macrogol.

PEG vs Placebo Systematic review and network meta-analysis in adults with non-organic constipation.



Adding electrolytes has added effect

Good for fecal dren

Belsey JD, Geraint M, Dixon TA. Int J Clin Pract. 2010 Jun;64(7):944-55





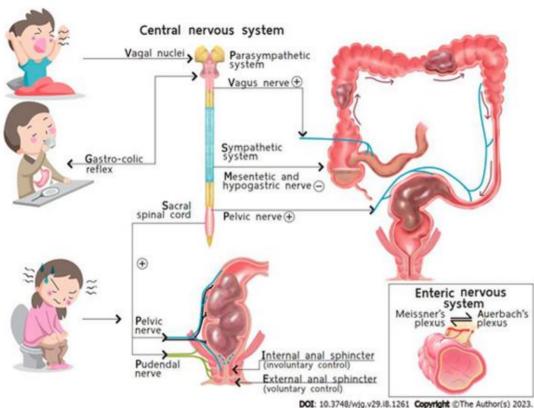
# **Macrogol in Pregnancy and lactation.**



- No effect anticipated during pregnancy, since systemic exposure is negligible with PEG
- PEG 3350/4000 can be taken during pregnancy.
  - No effect aniticipated on the infant, since systemic exposure of the mother to macrogol 4000 is negligible.
- PEG 3350/4000 can be taken during lactation.
- PEG 3350/4000 with Psyllium 3g/or simethicone can also be taken during lactation.



## Macrogol in Pediatric and adolesence.



PEG and lactulose are effective laxatives in children. There is limited evidence of its utilization and safety in infants.



# Macrogol and lactulose: Combined in colonoscopy prep.

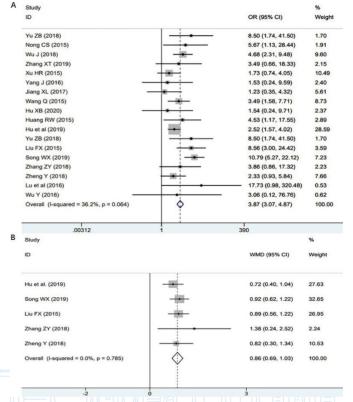
PEG combined with lactulose could improve bowel cleansing effect with fewer adverse reactions, thus serving as a simple, convenient, safe and effective method for bowel preparation.



nificant reduction in the incidence of abdominal distention was observed

pared with PEG alone

Conclusion: PEG combined with lactulose may be a better choice for bowel preparation before colonoscopy co





# **Chronic Constipation: Stimulant laxatives.**

- Types of stimulants
  - Small bowel irritants: castor oil
  - Large bowel irritants: Bisacodyl, Na picosulfate, Senna, Cascara
- Prolonged use can create habituation and drug dependence by damaging the colon's haustral folds.
- Medicine should not be used if:
  - previous <u>allergic reaction</u> to any stimulant laxatives,
  - intestinal obstruction,
  - rectal bleeding,
  - signs of appendicitis.
- To be prescribed with caution for diabetics, hypertensives, and patients with heart disease.

## Effect of Osmotic & stimulant Laxatives on Chronic Constipation: Meta-analysis of high quality studies

	La	xatives	S	PI	acebo			Mean Difference		Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	Year	IV, Random, 95% CI
1.2.1 Osmotic laxative	S									
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DiPalma 2007	7.9	4.5	204	5.6	5.5	100	16.2%	2.30 [1.06, 3.54]	2007	
Subtotal (95% CI)			325			209	57.5%	2.51 [1.30, 3.71]		•
1.2.2 Stimulant laxativ  Mueller-Lissner 2010		0.2	233	1.7	0.14	134	21.3%	1.70 (1.67, 1.73)	2010	
Mueller-Lissner 2010	3.4	0.2	233	1.7	0.14	134	21.3%	1.70 [1.67, 1.73]	2010	
Kamm 2010 Subtotal (95% CI)	5.2	0.27	247 480	1.9	0.34	121 255	21.3% 42.5%	3.30 [3.23, 3.37] 2.50 [0.93, 4.07]	2010	
Heterogeneity: Tau <sup>2</sup> = 1	28: Chi	= 163		f= 1 /P	< 0.00			2.00 (0.00, 4.01)		
Test for overall effect: Z			7.5		0.00	00.7,1	- 130 %			
Total (95% CI)			805			464	100.0%	2.55 [1.53, 3.57]		•
Heterogeneity: Tau <sup>2</sup> = 1	.26; Chi	= 164	2.45, d	f= 5 (P	< 0.00	001); l²	= 100%		H-1	0 -5 0 5 1
Test for overall effect: Z	= 4.92 (	P < 0.0	0001)							Favours placebo Favours laxative:
Test for subgroup differ	rences: (	Chi <sup>2</sup> = (	0.05, df	= 1 (P =	0.81)	$I^2 = 09$	6			arours praceso Tarours raxaure

Ford & Suares. Gut 2011



# **Chronic Constipation: Emolients.**

### **Mechanism of Action**

- Indigestible, minimally absorbed. Coat and allow easier passage
- Inhibit colonic absorption of water increasing weight and deceasing transit time.
- Emollient agents prevent constipation rather than treating long-term constipation they are best used in hospitalized patients following MI or surgery

### **Adverse effects**

- May be absorbed systemically and may lead to foreign body reaction in lymphoid tissue.
- Aspiration may lead to lipoid pneumonia
- Decrease absorption of fat-soluble vitamins (A,D,E,K)
- Mineral oil may leak from anal sphincter if given rectally



Heterogeneity: Tau2 = 0.08; Chi2 = 7.17, df = 3 (P = 0.07); I2 = 58%

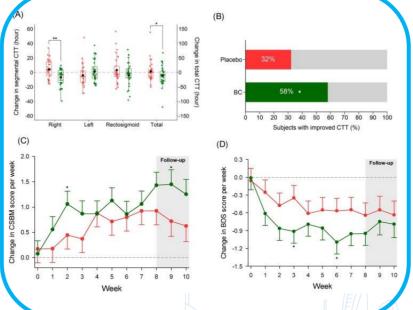
Test for overall effect: Z = 0.73 (P = 0.46)

# **Chronic Constipation: Probiotics.**

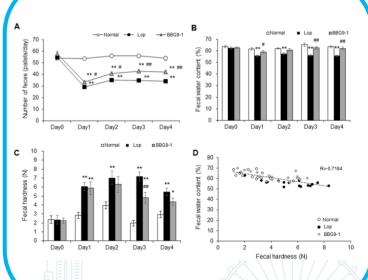
#### Probiotics for IBS-Constipation Meta-analysis and system review

Overall analysis Mean Difference Study or Subgroup Dimidi et al 2018 Fateh et al 2011 Mazivn et al 2013 Mirghafourvand et al 2016 2.00 [-3.33, 7.33] Heterogeneity: Tau2 = 0.03; Chi2 = 4.10, df = 3 (P = 0.25); I2 = 27% Test for overall effect: Z = 1.53 (P = 0.13) Favours probiotics Favours control Overall analysis Std. Mean Difference Dimidi et al 2018 Fateh et al 2011 -0.70 [-1.22, -0.18] Mazlyn et al 2013 -0.18 [-0.59, 0.24] Mirghafourvand et al 2016 0.19 [-0.32, 0.69]

Bacillus coagulans SNZ 1969 improved intestinal motility and constipation perception mediated by microbial alterations in healthy adults



Improvement of loperamide-induced slow transit constipation by *Bifidobacterium* bifidum G9-1 is mediated by the correction of butyrate production



Chengcheng Ziang et al. Clinical nutrition 2020. volume 39, Issue 10, 2960-2969

-0.14 [-0.50, 0.23]

Favours probiotics Favours control

Seunghee Kang et a.Food Research International. Volume 146, August 2021, 110428

Makizaki Y, et al. PLOS ONE 17(4): e0267927.



# Chronic Constipation: Laxatives, in pregnancy and postpartum.



Safe To be used Unsafe with caution Saline osmotic Laxative<sup>1</sup> Lactulose<sup>1</sup> Anthraquinone<sup>1</sup> Macrogol<sup>2</sup> Castor oil<sup>3</sup> Polyethylene Bisacodyl<sup>4</sup> glycol<sup>3</sup> Glycerine<sup>5</sup> Senna<sup>1</sup> Mineral oil3 Docusate sodium<sup>1</sup> Bulking agents 1

<sup>1.</sup> Cullen et al. Best Practice & Research Clinical Gastroenterology 2007;21(5):807–818

<sup>2.</sup> Summary of Product Characteristics. 2015.

<sup>3.</sup> ACG 2007.

<sup>4.</sup> Sanofi.Dulcolax 5 mg Gastro-resistant Tablets. 2017.

<sup>5.</sup> Verghese et al. The Obstetrician & Gynaecologist 2015;17:111–5



# **Chronic Constipation: Beyond front line.**

Medicine	Class	Effect
Procalupride	5HT4 agonist	Accelerate colonic transit, concerns about QT interval
Veleutrag	5HT4 agonist	Clinical trial
Naronapride	highly selective, high-affinity 5-HT(4) receptor agonist	Clinical trial



# Non-medicinal treatment.

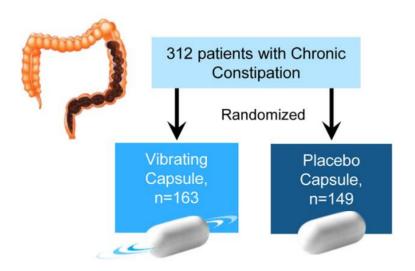




# Chronic Constipation: drug free treatment (vibrant cap).

## **Vibrating Capsule Treatment for Chronic Constipation**

Phase 3, Double Blind, Multicenter, Placebo controlled trial



- Patients ingested one capsule at bedtime daily for 5 days a week
- · Duration of study= 8 weeks

#### **Primary Outcome Measures:**

Increase in one or more or two or more complete spontaneous bowel movements (CSBM) per week over baseline in 6 out of 8 weeks

Gastroenterology



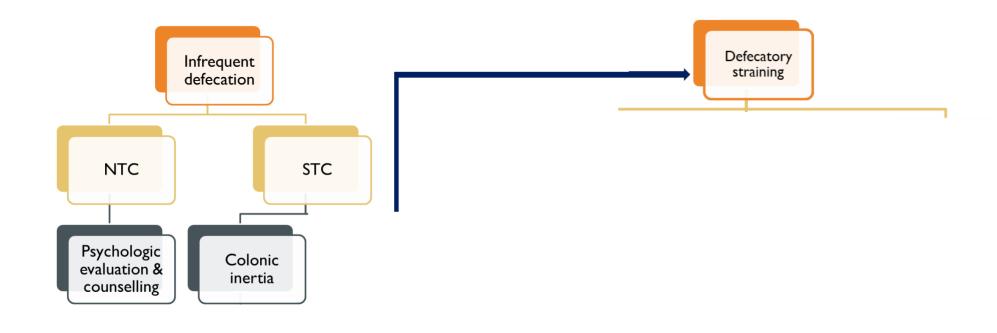
# **Chronic Constipation: drug free treatment (Biofeed back).**

- An instrument-based learning process;
- Using equipment to record or amplify a patient's anorectal activity and then provides feedback to the patient and therapist.
- The goal of anorectal BFT is to:
  - >strengthen the pelvic floor muscles,
  - >retrain rectal sensation,
  - coordinate the activity of abdominal, pelvic floor and sphincter muscles during evacuation





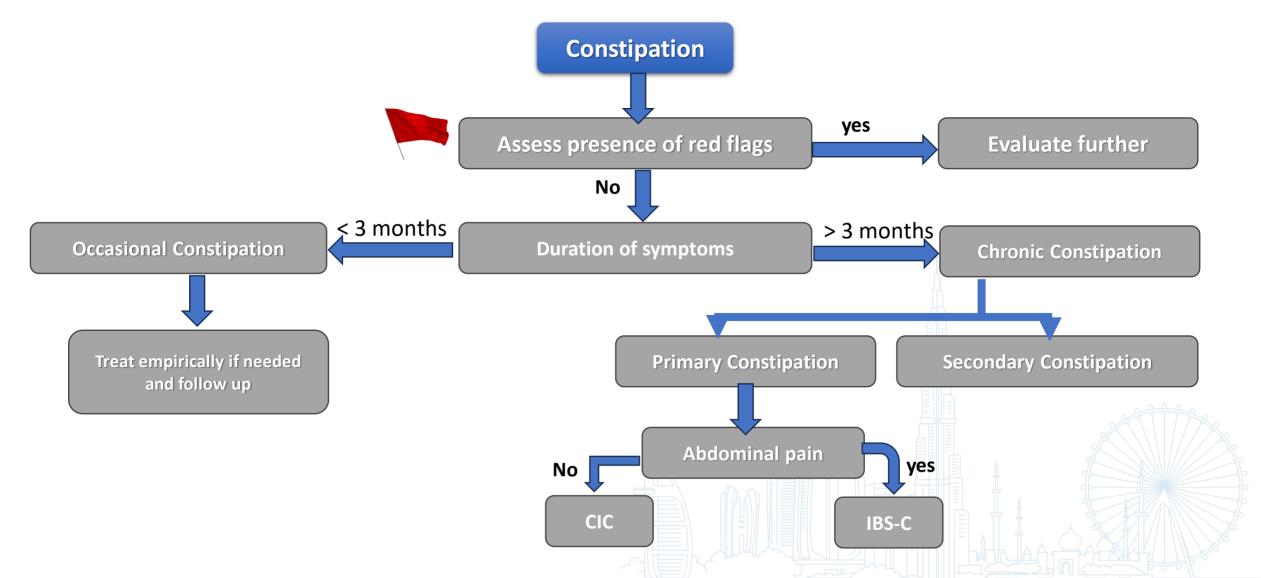
# Chronic Constipation: drug free treatment (Surgery).







# Chronic Constipation: diagnostic algorithm.





# **Chronic Constipation: ACG-AGA guidelines.**



Fiber supplementation with bran, inulin, psyllium, and methylcellulose. The <u>best data exist for psyllium</u> (Increase in global relief and Spontanous Bowel Movement/week).







PEG results in an increase in complete spontaneous bowel moments (CSBMs) per week, compared with placebo (meta-analysis data. First line treatment of osmotic laxatives.



MgO at a lower dose, which may be increased if necessary is an attractive first-line option (good efficacy, tolerable, OTC available, and low cost.



Lactulose may be considered if symptoms of CIC have failed to improve with fiber and OTC laxatives, and individuals do not experience significant bloating or abdominal pain



<u>Sodium Pico Sulphate</u> leads to an increase in CSBMs and SBMs per week and to improved stool consistency (Side effects are common), Bisacodyl is another stimulant



# **Chronic Constipation: ACG-AGA guidelines.**



the combination of efficacy, impact on quality of life, OTC availability, and low cost makes <u>senna a</u> viable first-line option for patients with CIC.







<u>lubiprostone improved stool frequency and consistency as well as abdominal discomfort and bloating</u>. The panel advised that patients with moderate or severe hepatic insufficiency should receive a lower dose, such as 8 mcg twice daily.



<u>linaclotide leads to increases in the number of CSBMs</u> per week and SBMs per week, improves stool consistency, and increases the rates of global relief. (If diarrhea evolves reduce the dose)



<u>plecanatide in adults with CIC</u> is associated with an increase in the number of CSBMs per week and SBMs per week and improves the quality-of-life scores, although it may be associated with diarrhea.



<u>prucalopride</u> was associated with an increased number of CSBMs per week. The most common side effects were headache, abdominal pain, nausea, and diarrhea.



# Chronic Constipation: Rome IV & WGO guidelines.

Pediatrics	Adolescents or children	Adults (according to WGO)
Non-pharmacological treatment for FC consists of education, demystification,	> Two or fewer defecation/W.	Patient history + physical examination
regular dietary advice and in older children toilet training, a reward system and a stool	At least one episode of fecal incontinence/W.	Classify the patient's type of constipation
diary.	H/O retentive posturing or excessive volitional stool retention	Medical approach in uncomplicated normal- transit constipation without alarm symptoms
> Reduce fear and, make the child and parents		
understand the underlying pathophysiological mechanisms and the	➤ H/O painful or hard bowel movements	In treatment-resistant constipation, specialized investigations can often identify a
need to learn how to recognize these in daily life.	Presence of a large fecal mass in the rectum	cause and guide treatment
<ul> <li>Disimpaction should be attempted, followed</li> </ul>	H/O large diameter stools that can obstruct the toilet No other medical condition.	If treatment fails, continue with specialized testing (this may only apply to the "extensive
by maintenance treatment with laxatives.		resources" level)
> PEG is the primary preferred medication for		Treatment of STC* with aggressive laxative
disimpaction		programs



- CC is recognized by the Rome IV criteria as a subtype of functional GI disorders, a commonly encountered disease.
- The diagnosis is based on clinical, laboratory, radiologic and manometric judgement
- The first line of drug treatment are laxatives
- Osmotic laxatives (Lactulose & PEG) when used alone, with electrolytes, other bulking fibers, or anti-flatulents, are safe and can be used for children, the pregnant and lactating mothers, the elderly and in some chronic disorders



