Colonoscopy Preparation

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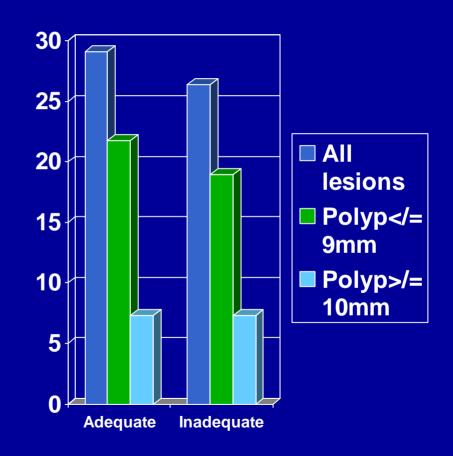
Colon Preparation Purpose

- Improve Visualization:
 - -Remove stool from lumen and wall.
 - -Avoid obscuring of lenses/ camera and light beam.
 - -Minimize "missed lesion rate".
- Remove hydrogen, methane, and other explosive gasses.
- Shorten procedure time.
- Limit Costs.

Prep Quality vs Neoplasia Detection

Harewood et al. Gastrointest Endosc 2003;58:76-9

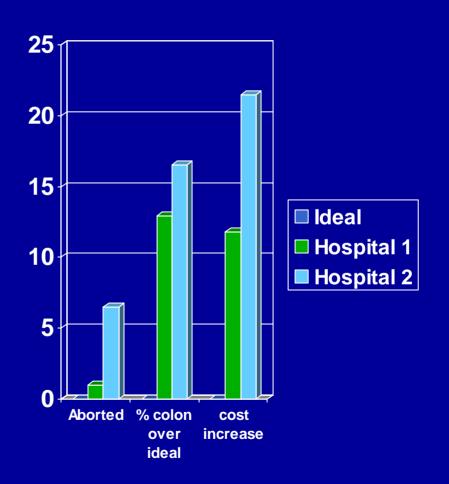
- Clinical Outcomes
 Research Initiative
 database.
- Retrospective
- Subjects: 93004
- 01/01/2000-12/31/2001
- Adequate (76.9%) vs
 Inadequate prep (23.1%)
- Larger lesions detected equally. Some small lesions missed.



Bowel Prep vs Efficiency and Cost

Rex et al. Am J Gastroenter 2002;97:1696-1700

- Compared ideal prep vs observed in:
 - Private Hospital (1) w.13.5% poor prep
 - Public Hospital (2) w.26.5% poor prep
- Observed frequency of
 - aborted exams,
 - exams repeated early
 - cost increase.



Risk of Explosive Gases

- Fatal Colonic Explosion after colon prep with Mannitol (Bigard et al. Gastroenterology 1979;77:1307-10)
- Colon explosion with perforation after APC treatment in unprep colon (enemas only) (Soussan et al. Gastrointest Endosc 2003;57:412-3)
- REMEMBER: Use of Lactulose or Sorbitol can cause flammable gases.

Instruments and Risk of Explosion

• RISKY:

- Monopolar Electrocautery ("hot" snare & "hot" biopsy)
- Bipolar Electrocautery (BICAP)
- Argon Plasma Coag.
- Laser

NOT RISKY

- Biopsy and "cold" snare
- Heater Probe
- Injection hemostasis
- Endo-loop
- Hemo-clip
- Band-ligation

Conclusions

- Poor colonoscopy prep increases risk of missed polyps.
- Poor colonoscopy prep increases cost by increasing the number of aborted exams and early re-examinations.
- Poor colonoscopy prep and use of fermentable carbohydrates (sorbitol, lactulose, mannitol) increase risk of colonic explosion.

Methods for Colon Preparation

- Two day clear-liquid diet + 300 ml Mg
 Citrate @ pm + tap water enemas until clear 2 h before exam.
- One day clear-liquid diet + Senna 150 mg
 @ noon and again at 6 pm.
- Iso-osmotic colon irrigation (Saline lavage or PEG-balanced solutions).
- Hypertonic saline solutions (NaP, Mg Citrate)

Oral Colon Irrigation Solutions

- Work by flushing the colon with large volume of clear fluid.
- Based in principle that small bowel contents have to remain isotonic to plasma, hence, non-absorbable solutes will draw a "mandatory volume" to the cecum to keep isotonicity.
- Suprep, in two 6 oz bottles has: 492 mEq
 Na, 72 mEq K, 26 mmol Mg, 898 mOsm.

Osmolar Loads of Colon-Preparation Solutions

- NuLytely or Golytely = 1160 mOsm/4 L
- NuLytely or Golytely = 870 mOsm/3 L
- Fleet Phosphosoda = 807 mOsm/ 90 mL
- Visicol = 820 mOsm/40 tab
- Visicol = 660 mOsm/32 tab
- Visicol = 575 mOsm/28 tab
- Mg Citrate = 720 mOsm/900 mL
- Mg Citrate = 480 mOsm/600 mL
- Suprep 12 onz = 898 mOsm/ 336 mL

Fluid needs for Iso-tonicity (fluid deficit) in mL

Nulytely or Golytely (any volume)= 0

• Fleet Phosphosoda 90 mL = 2690

• Visicol 40 tabs = 2820

• *Visicol 32 tabs* = 2275

• Visicol 28 tabs = 1980

• Mg Citrate 900 mL = 1582

• Mg Citrate 600 mL = 1055

• Suprep 12 onz (336 mL) = 2760

Volume Presented to Cecum

 NuLytely or GoLytely 4L 	4000 mL
---------------------------------------------	---------

- NuLytely or GoLytely 3 L 3000 mL
- Fleet Phosphosoda 90 mL 2780 mL
- Visicol 40 tablets
 2820 mL
- Visicol 32 tablets
 2275 mL
- Visicol 28 tablets
 1980 mL
- Mg Citrate 3 x 300mL 2480 mL
- Mg Citrate 2 x 300 mL 1655 mL
- Suprep 12 onz (336 mL) 3096 mL

Contraindications of Oral Colon Irrigation Solutions

- Na Phosphate
- GI obstruction
- Gastric retention
- Bowel perforation
- Toxic colitis
- Ileus
- Megacolon
- Ascites
- Heart failure
- Impaired renal function
- Electrolyte imbalance
- Debilitated condition

- PEG-balanced
- GI obstruction
- Gastric retention
- Bowel perforation
- Toxic colitis
- Ileus
- Hypersensitivity

Oral Irrigation Solutions Conclusions

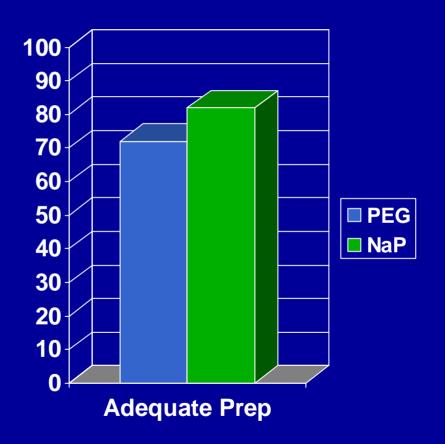
- Iso-osmotic solution need little or no additional fluid ingestion.
- Hypertonic solutions require ingestion of substantial amounts of fluids to avoid dehydration.
- Na Phosphate has additional contraindications in heart failure, kidney disease, electrolyte disorders, ascites, megacolon and inability to drink extra fluids (debilitated).

Single Preparation as in "paper insert"

PEG vs Phosphosoda Meta-analysis

(Hsu et al. Gastrointest Endosc 1998;48:276-82)

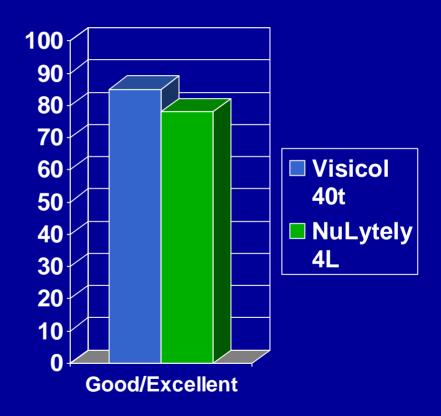
- Meta-analysis of 8 trials, 1286 subjects
- Phosphosoda in splitdose (one or two day)
- PEG solution in single dose



PEG vs Visicol

Kastenberg et al. Gastrointest Endosc 2001;54:705-13

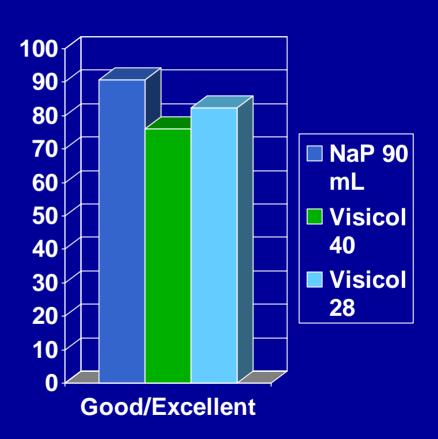
- Prospective & randomized.
- Visicol 20 tab + 1680
 ml water @ 6 pm and
 same repeat in am, vs
- NuLytely 4 L @ 6 pm



Fleet Phosphosoda vs Visicol

Balaban et al. Gastrointest Endosc 2003;57:AB102

- Prospective, randomized.
- Phosphosoda 45 ml@ pm + 45 ml @ am, vs
- Visicol 20 tab @ pm +
 20 tab @ am, vs
- Visicol 14 tab @ pm +
 14 tab @ am



Conclusion

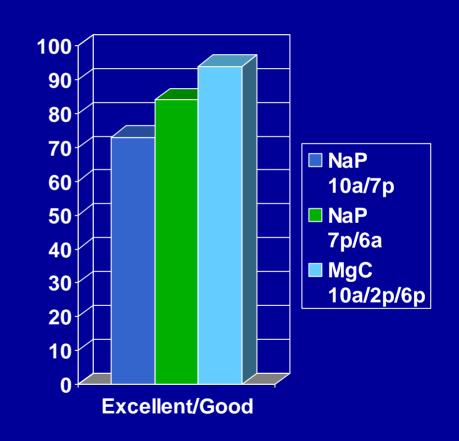
 If used as in "paper insert", colonoscopy prep with liquid Phosphosoda gives better cleansing than with Visicol, and preparation with Visicol gives better cleansing than with PEG-balanced solutions.

Timing vs Quality of Preparation

Timing vs Quality of Prep

Belkelhammer et al Gastrointest Endosc 2002;56:89-94

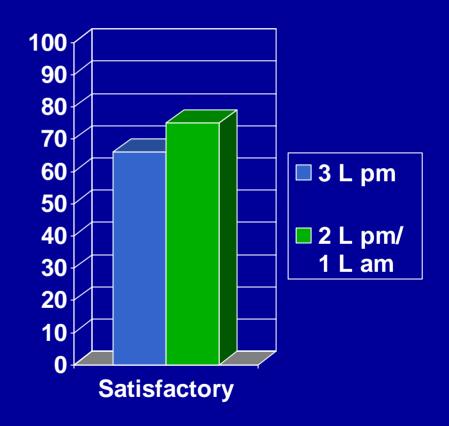
- Prospective, randomized
- All patients on clear liquids + plenty of fluids
- Na P: 45 ml + 45 ml
 -10am/7pm vs
 -7pm/6am)
- Mg Citrate: 300 ml + 300 ml + 300 ml (10am/2pm/6pm)



Timing vs Quality of Prep

ElSayed et al. Gastrointest Endosc 2003;58:36-40

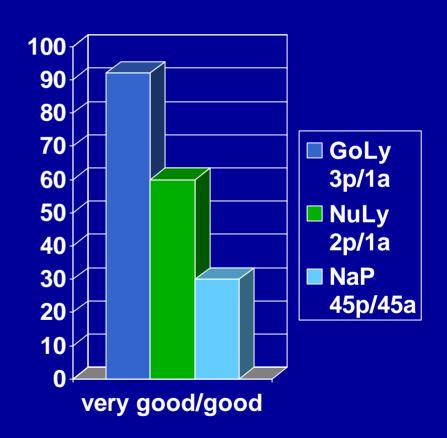
- Prospective, randomized
- Clear liquid diet day before, then NPO
- PEG solution
 -3 L @ 6 pm vs
 -2 L @ 6 pm + 1 L @
 am



Timing vs Quality of Prep

Ell et al. Endoscopy 2003;35:300-304

- Prospective, randomized
- Clear liquid lunch & dinner + plenty fluids
- Golytely 3 L pm + 1 L @ 6 am, vs
 Nulytely 2 L pm + 1 L @ 6 am, vs
 NaP 45 ml pm + 45 ml @ am
- Categorized by worst cleansed segment.

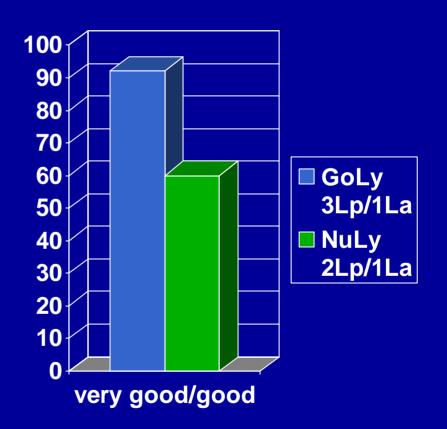


Timing vs Quality Conclusions

- In Phosphosoda Prep: a two-day evening and morning dose is superior to the one-day morning and evening dose.
- In PEG Prep: a two-day evening and morning split-dose is superior to the one-day evening dose.
- A two-day Split dose PEG Prep (3 L or 4 L) is superior to the two-day split-dose Phosphosoda prep.
- A 3-dose, one-day Mg Citrate Prep is superior to the two-day Phosphosoda Prep.

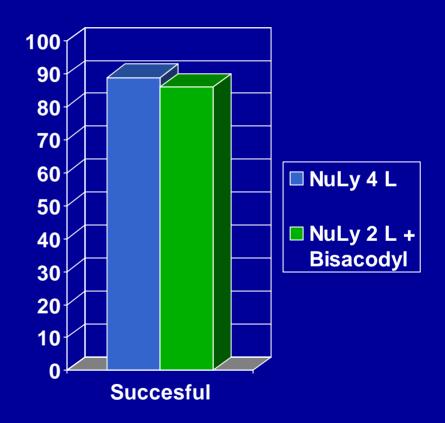
Ell et al. Endoscopy 2003;35:300-304

- Prospective, randomized
- Clear liquid lunch & dinner + plenty fluids
- Golytely 4 L (3 L pm + 1 L @ 6 am), vs
 Nulytely 3 L (2 L pm + 1 L @ 6 am), vs



DiPalma et al. Am J Gastroenterol 2003;98:2187-2191

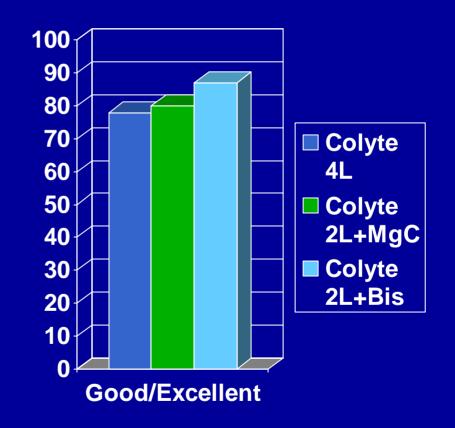
- Prospective, randomized
- Clear liquid lunch & dinner; NPO p MN.
- Nulytely 4 L @ 6pm
- Bisacodyl (Dulcolax)
 20 mg @ noon +
 Nulytely 2L @ 6 pm



4-Liter PEG vs 2-Liter PEG + MgCitrate or Dulcolax

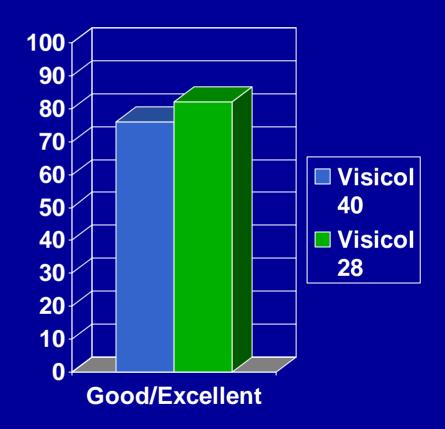
Sharma et al. Gastrointest Endosc 1998;47:167-71

- Prospective and randomized.
- Colyte 4-L one-day@ 4 pm, vs
- MgCitrate 300 ml @
 4pm + 2-L Colyte @
 5 pm, vs
- Bisacodyl 20 mg @
 4 pm + Colyte 2-L @
 4 pm.



Balaban et al. Gastrointest Endosc 2003;57:AB102

- Prospective, randomized.
- Visicol 20 tab @ 6
 pm and 20 tab in am,
 vs
- Visicol 14 tab @ 6
 pm and 14 tab in am.



Volume vs Quality of Prep Conclusions

- PEG 4 Liters is superior to 3 Liters (volume of flow)
- Pre-treatment with Mg Citrate or Bisacodyl, makes 2-Liter of PEG prep equivalent to 4-Liters of PEG alone (force of the flow).
- Visicol 28 tablets is equivalent or better than 40 tablets.

Safety

Safety of Na Phosphate vs PEG Preparations

Hookey et al. Gastrointest Endosc 2002;56:895-902 Beloosesky et al. Arch Int Med 2003;163:803-808

- Frequency of asymptomatic orthostatic hypotension similar to PEG prep.
- Serum K+ mean-drop: 0.4 mEq/L in NaP vs 0.1 with PEG. In 13 to 25%, K+ < LLN.
 In the elderly, 56% had hypo-kalemia.
- Serum Ca⁺⁺ mean drop: 0.05-0.15 mEq/L with NaP vs 0-0.02 with PEG. In 6-11% Ca⁺⁺ <LLN; in 44% iCa⁺⁺ < LLN; none with symptoms. In the elderly, 58% had hypo-calcemia.

Adverse Events of Na Phosphate in Colonoscopy Prep (Adults)

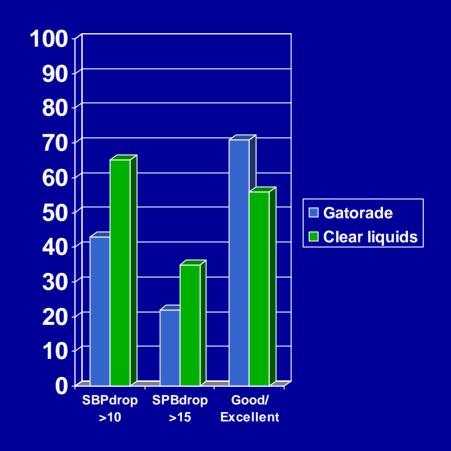
Hookey et al. Gastrointest Endosc 2002;56:895-902

- **Six** hemodynamic and electrolyte disorders with **proper use** (hypovolemia, hypokalemia, hypornatremia, hypocalcemia, ischemic colitis).
- Four seizure due to electrolyte disorders, with proper use.
- Two fatal hyperP & ARF due to high dose.
- Five hemodynamic, electrolyte, and renal disorders due to high dose.
- Increased risk of "phosphate nephropathy" in patients taking ACE inh., ARBs & NSAIDs (Am J Nephrol 2005;16:3389-96).

Gatorade + Phosphosoda

Barclay et al. Gastrointest Endosc 2002;56:633-8

- Prospective, randomized.
- Phosphosoda 45 ml
 @5 pm & 10 pm +
 clear liquids all day.
- Gatorade 3.8 L vs
 Clear liquids 3.8 L.
- Drop in Syst BP
- Quality of Prep.



Phosphosoda and Aphthous Colonic Ulcers

Berkelhammer et al. Gastrointest Endosc 2002;56:89-94

- In 5.5% of patients.
- Small, 1-3 mm, erythematous halo, non-confluent, discrete, over normal mucosa.
- Most common in rectosigmoid.
- Independent of timing of doses (9 vs 11 hours apart)
- Diagnostic confusion in 40% of them.
- Pathology shows mucosal disruption with lymphoid aggregates.

SafetyConclusions

- When used properly in healthy patients, all colon preps are safe.
- Na Phosphate causes hypo-kalemia and hypo-calcemia in more than half of elderly patients.
- Na Phosphate should not be used in patients with significant cardiac, renal, or liver disease, nor in the debilitated. Also avoid them in patients taking ACE inh., ARBs & NSAIDs.
- Gatorade decreases NaP-induced volume-depletion and improves prep quality.
- Aphthous-like lesions can occur with NaP prep, which can cause confusion with IBD.

Some Equivalent Preps

(expected > 90% Good/Excellent)

Breakfast, day before Exam	Noon, day before Exam	6 PM, day before exam	AM of Exam-day (finish 3h before leaving home)
Regular	Clear liquid diet	[PEG 3L or PEG 2 L] + Bisacodyl 20 mg + clear liquids	PEG 1 L
Regular	Clear liquid diet	[PEG 4 L or PEG 2 L] + Bisacodyl 20 mg + clear liquids	Mg Citrate 300 ml + Gatorade 16 oz
Regular	Clear liquid diet	NaP 45 ml + Gatorade 2.85 L (3 Qt = 96 oz or 12 glass)	NaP 45 ml + Gatorade 48 oz (1.5 Qt = 48 oz or 6 glass)
Regular	Clear liquid diet & Mg Citrate 300 ml + Gatorade 48 oz	Mg Citrate 300 ml + Gatorade 48 oz (6 glass)	Mg Citrate 300 ml + Gatorade 16 oz (2 glass)
Regular	Clear liquid diet	Suprep (6 oz + 10 oz water) + clear liquids 88 oz (11 glass)	Suprep (6 oz + 10 oz water) + clear liquids 40 oz (5 glass)

The volume of Gatorade in the 1st day covers needs for insensible losses & urine output

My Conclusions

- Colon Preps work better when they are divided in two-days, with a PM + AM dose.
- Clear-liquid diet should start at least at lunch time on day before exam.
- AM dose in second day, should be ingested completely at least 3 hours before leaving home.
- When taking NaP (90 mL) prep, 3-4 Qt of Gatorade decreases volume depletion.
- Poor preps increase the cost of colonoscopy, and "missed lesion rate".