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Pattern of intragastric carbon dioxide concentrations in healthy subjects and patients with functional dyspepsia and gastroparesis: Evidence of greater duodeno-gastric reflux in association with delayed gastric emptying?

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Background

The Atmo gas sensing capsule is a novel, single-use ingestible device that measures several parameters, including carbon dioxide (CO₂). Intra-gastric CO2 "bursts" were observed during a pilot study of regional gut transit in dysmotility.¹ The aetiology of this CO₂ was unclear prompting further evaluation.

Aims

To describe these intragastric bursts of CO₂ in patients with a clinical diagnosis of functional dyspepsia and/or gastroparesis compared with healthy subjects in relation to:

- · Frequency and duration
- Clinical diagnosis
- · Physiological events speed of gastric emptying & ingestion of liquid/food
- · Intragastric pH profile

Methods

Subjects: Clinical diagnosis of functional dyspepsia or gastroparesis and healthy subjects.

Capsule studies: Following an 8-hour overnight fast, subjects ingested a standardised cereal bar followed by the gas-sensing capsule and wireless motility capsule (WMC) in random order with an additional 6-hour fast prior to resuming their habitual diet (without carbonated beverages). Motility- and pH-altering medications were withheld 72h prior and for the study duration. Food/drink consumption was recorded on a patient-held data receiver.

1. Zhou et al., DDW, 2023; 2. Ou et al., Sci Rep, 2016; 3. Cuomo et al, Nutr Metab. Cardiovasc Dis. 2009; 4. Levitt and Bond, Gastroenterol, 1970.

Outcomes measured: CO₂ bursts were described by:

- 1) frequency and size (area-under-the-curve);
- 2) associations with food/drink consumption and intragastric pH changes:
- 3) correlation with clinical diagnosis;
- 4) correlation with gastric emptying time.

Results

Example traces shown in Fig 1.





Fig. 1. Example traces of CO₂ (green) with black arrows showing bursts. Gastric emptying shown in mustard.



	Healthy	paresis	Dyspepsia	p-value
No. (%) subjects with CO2 bursts	2/4 (50%)	16/25 (64%)	5/5 (100%)	0.25 ^b
No. of bursts per participant med. (IQR)	3 (2-3)	2 (1-4)	2 (1 -3)	0.57ª
Proportion related to food/drink	0	13/41 (32%)	2/9 (22%)	0.35 ^b
AUC (flat), %.h med. (IQR)	1.2 (0.17-2.2)	6.1 (0.7-30.3)	1.8 (0.5–4.4)	0.22 ^a
P<0.001		250 - 4 200 - 0) 150 - 150 - 0) 100 - 50 -	P<0.001 ← →	

Fig. 2. Frequency and size of CO₂ bursts according to delaved (blue) or normal (pink) gastric emptying.

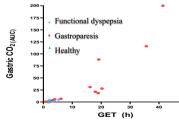


Fig. 3. Association of AUC for CO₂ bursts with gastric emptying time (GET) according to clinical diagnosis.



The greater number and size of bursts in subjects with delayed gastric emptying time remained significant when adjusting for gastric residence time (all p-values <0.01).

Conclusions

Intra-gastric bursts of CO₂ are

- · associated with delayed gastric emptying, not prior clinical diagnosis
- can be quantified by number and size.

So, what are they?

There are four potential sources of intragastric CO₂ production.²⁻⁴

- 1. Chemical reactions with acid/base $H^+ + HCO_3 \rightarrow CO_2 + H_2O_3$ (e.g., duodeno-gastric reflux of bicarbonate-rich fluid)
- 2. Ingestion of CO₂ (e.g., swallowing air, food/drink)
- Alteration in gastric diffusion of CO₂ (e.g., hyperventilation effect) 3.
- 4. Microbial metabolic activity (not significant in non-rumen stomach).



The most likely physiological explanation for these bursts of CO₂ is duodeno-gastric reflux

Implications

Further validation is required in a larger cohort of healthy subjects and those with functional dyspepsia and gastroparesis.

Ability to quantify this phenomenon offers a unique opportunity to evaluate duodeno-gastric reflux

- in a variety of clinical conditions
- · its modulation with therapeutic intervention.

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