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## NVESTOR PRESENTATI

## Corporate Snapshot

DEVELOPING EVIDENCED-BASED SOLUTIONS FOR GASTROINTESTINAL TRACT HEALTH ISSUES

#### **About Anatara**

Anatara Lifesciences Ltd (ASX:ANR) ("Anatara" or "the Company") is focused on the validation and commercialisation of innovative, evidence-based products for human health with significant unmet needs with a focus on gastrointestinal considerations.

The Company has completed Stage 1 of its pivotal clinical trial for irritable bowel syndrome with the patented GaRP product, with interim results released in Q3CY23.

Additionally, there are a number of other indications for the use of GaRP and Anatara is actively assessing other opportunities in the healthcare space.

ASX Code	ANR	
Share Price	\$0.025	023
Shares on Issue	~119.9m	JG 2
Market Capitalisation	~\$2.9m	18 AL
Cash (June 30 2023) + R&D rebate	~\$0.35m + (~\$922k R&D rebate)	

## AN

### Investment Highlights

#### **Novel Technology**

 Gastrointestinal Reprogramming (GaRP) technology aimed at restoring and maintaining the GIT lining, integrity and homeostasis.

#### Pivotal clinical trial – Irritable bowel syndrome

- Primary Endpoint Efficacy of 'GaRP' for irritable bowel syndrome.
- Positive Stage 1 results announced 3Q CY2023, Stage 2 (full results) anticipated 2Q CY2024.

#### **Significant market opportunities**

Digestive Health Market to be valued at **US\$23.4B** in 2030 with a CAGR of **8.1%.**1

#### Significant unmet medical need

- Treatments for IBS/IBD are often not effective in controlling symptoms.
- Only 15-20% of patients are 'very satisfied' with their treatment options.<sup>2</sup>
- GaRP addresses the underlying factors of many GIT disorders, providing multifaceted symptomatic relief while rejuvenating the tract.

#### **Assessing commercial options for GaRP**

- Actively assessing potential commercial partners for GaRP.
- Initiated assessment of whether Stage 1 results will satisfy claims for GaRP to be Listed through the AUST L Nos pathway.

#### **GaRP Technology to provide a pipeline of products**

Potential indications for GaRP include Inflammatory Bowel Disease (IBD), Paediatric indications & Functional Dyspepsia.

#### **Assessing additional corporate opportunities**

Potential to acquire/license innovative assets/programs.

## Key Management & Board of Directors

#### HIGHLY EXPERIENCED BOARD & MANAGEMENT



**Dr David Brookes**Executive Chair

- 30+ years international experience in the health and biotechnology industries.
- Former Chairman of genomics solutions company, RHS Ltd (ASX: RHS); acquired by PerkinElmer Inc (NYSE:PKI).
- Medical Practitioner, Biotechnology Consultant
- MBBS, FACRRM, FAICD.



John Michailidis
Executive Director

- John is an executive with more than 30 years' of commercial pharmaceutical experience.
- His career has ranged from global franchise and regional executive leadership roles with F. Hoffman La Roche to CEO experience with emerging biotechs.
- BSc (Hons) in Genetics from LaTrobe University and exec. business qualifications from Harvard Business School and INSEAD.

Anatara's nimble Board and Management structure provides significant experience and flexibility for potential corporate transactions.



Nicholas Haslam
Non-Executive Director

- Nick is a chartered accountant with 10 years of experience in professional services within M&A and restructuring at PwC.
- His medical experience includes being the CEO and since then ongoing consultant to medical device company Plasma Shield, as well as playing a leading role in scaling up KangaTech, a medical device and injury prevention business with operations now established internationally.



## Gut Health is critical to general wellbeing

#### Gut Brain Axis & Mental Health

The gut and the brain communicate with each other through a complex network of nerves, hormones, and organic messengers.

A healthy gut can have significant benefits:

- Improved Mood: Having a healthy gut can help maintain optimal levels of neurotransmitters (serotonin and dopamine), which can contribute to improved mood.<sup>1</sup>
- Reduced Anxiety and Depression: An imbalanced gut microbiome has been linked to an increased risk of anxiety and depression. A healthy gut can help reduce the risk of these mental health conditions.<sup>2</sup>
- Enhanced Cognitive Function: A healthy gut with a balanced microbiome can produce various molecules, that can support brain health and cognitive function.<sup>3</sup>
- Enhanced Nutrient Absorption: Absorption of essential vitamins and minerals is improved by a healthy gut.<sup>4</sup>

#### Well-being & immune system interactions

Maintaining gastrointestinal integrity and microbiome homeostasis can help assist autoimmune diseases by minimising dysregulation.<sup>11</sup>

A healthy gut can have other significant general benefits:

- Weight Management: Enrichment of bacterial genes and reduced diversity in the gut microbiome has been linked to obesity.<sup>5 6</sup>
- Appetite Control: Microbes in the digestive system produce metabolites that help regulate appetite and body fat stores.<sup>7</sup>
- Reduced Inflammation and Blood Sugar: Imbalances in the gut microbial community are associated with increased inflammation and increased blood sugar levels.<sup>8</sup>
- Improved Nutrient Absorption: Efficient absorption of essential nutrients, which can help optimize metabolism and energy production.<sup>4</sup>
- Regulation of Metabolism: A balanced Microbiota affects the regulation of metabolism which is closely linked to weight management.<sup>9</sup>





## Snapshot: IBS and IBD



11%

IBS affects 11% of the global population <sup>1</sup>

#### **Debilitating Symptoms**

Patients experience symptoms such as pain, bloating and diarrhoea.

#### **Limited Treatments**

Pharmacological options remain limited and often leave patients with poorly controlled symptoms.<sup>3</sup>

#### **Frustrated Patients**

45% of IBS-D patients agreed with the statement "I'm willing to try anything to help manage my IBS".4

1. Clinical Gastroenterology and Hepatology 2012: 10, 712-721. 2. Crohn's and Colitis Australia, 3. Grundmann & Yoon 2010. 4. IBS Global Impact Report 2018 https://badgut.org/wp-content/uploads/IBS-Global-Impact-Report.pdf.

### GASTROINTESTINAL DISORDERS ARE HIGHLY PREVALENT AND POORLY MANAGED



+5 million

IBD affects more than 5 million people globally, with accelerating incidence <sup>2</sup>

#### **Patients seek alternative options**

Up to 50% of IBS/IBD patients use dietary supplements, complementary & alternative medicines. <sup>5 6</sup>

#### **Doctors recommend supportive treatments**

Health-care practitioners increasingly recommend the use of such supportive treatments. <sup>7</sup>

#### **Iberogast recommended**

For example, source of recommended use of Iberogast: Healthcare provider 39.4%. <sup>8</sup>

5. Gastroenterology 2017: 152:415-429, 6. World J. Gastroenterol 2014: 346 – 362. 7. Michelfelder, Lee, & Bading 2010. 8. Yoon, Grundmann, Smith & Mason 2018



## IBS in America Survey

USE OF TREATMENTS FOR IRRITABLE BOWEL SYNDROME AND PATIENT SATISFACTION, BASED ON IBS IN AMERICA SURVEY

#### **Trial Details**

Use and satisfaction of various IBS treatments

3,254

IBS sufferers (Rome III criteria)

302

Physicians and gastroenterologists

#### **OTC** is Preferred Treatment Method

77%

of individuals with IBS have used OTC treatment for IBS symptoms

15%

of individuals with IBS have used prescription medications



The publication

Significant opportunity for an effective, evidenced based OTC product

GaRP

#### **Low Treatment Satisfaction**

15%

of IBS-C sufferers were 'very satisfied' with OTC treatments

20%

of IBS-D sufferers were 'very satisfied' with OTC treatments

#### **Ineffectiveness of Prescription Options**

19%

of IBS-C sufferers were very satisfied with an FDA approved medication

11%

of IBS-D sufferers were very satisfied with an FDA approved medication



## Active Program

#### GARP PROGRESSING TOWARDS COMMERCIALISATION



#### Gastrointestinal RePrograming (GaRP) technology for 'gut health'

Active pivotal clinical trial – Irritable Bowel Syndrome:

- Primary Endpoint Efficacy of 'GaRP' for irritable bowel syndrome.
- 120-140 patient, placebo-controlled, dose-escalating pivotal trial.
- Delivered successful Stage 1 results.
- Stage 2 Final readout 2Q CY2024.

#### **Safety Profile**

• All components GRAS (generally regarded as safe by FDA).

#### **Intellectual Property**

Patent pending formulation with functional coatings of a synergistic combination of components.

#### **Manufacturing**

GMP clinical batches manufactured and released for clinical trial.

#### **Commercialisation via Partnering**

 Anatara is in early stage discussions with potential global partners. The Company is experiencing in-bound interest regarding the clinical validation of GaRP.

#### **Pipeline Indications**

- Evaluating new indications for the GaRP technologies pipeline in particular, inflammatory bowel disease, functional dyspepsia and paediatric indications.
- KOLs engaged.



### Gastrointestinal Health

ANATARA HAS ACTIVE PROGRAMS APPROACHING COMMERCIAL OUTCOMES

#### Gastrointestinal RePrograming (GaRP)

GaRP is a multi-component, multi-coated patented complementary medicine. It is designed to address the underlying factors associated with chronic gastrointestinal conditions such as IBS and IBD, in part by assisting to restore and maintain the integrity of the tract lining and the homeostasis of the microbiome.



						Pre-C	Clinical	Clin	ical		
Program	Product	Indication	Trial Population	Delivery	Discovery	In-Vitro	In-Vivo	Proof of Concept	Pivotal	Registration	Commercial Rights
Human	GaRP	Irritable bowel syndrome	Ages 18-65	Oral Minitabs	<b>√</b>	<b>√</b>	<b>√</b>	3Q CY2023	2Q CY2024	Listed AUSTL Nos pathway	Global



## GaRP

#### ONE PRODUCT, MULTIPLE BENEFITS

#### **Patent Protected**

Scientifically designed to manage and control background IBS symptoms as well as episodic flare ups.

#### **Bromelain – Leading innovator**

Anatara has in-house knowledge of Bromelain protease activity and the associated QA required for clinical benefits.

#### **Everyday Option**

Designed as an everyday option to manage the causes and relieve symptoms of IBS (pain, cramping, gas, bloating, diarrhoea & constipation).

#### **Effective Relief**

Coated components released to the target areas for effective and sustained relief - 2 components release in the small intestine and 3 components have additional coating for delivery in the large intestine (colon).

#### **Natural Components**

Combines natural bromelain extract from pineapple stems in a patent pending formulation with other synergistic coated GRAS components. These components have specialist roles when delivered into specific regions of the GIT. \*must be delivered coated to colon/large intestine for MOA (Method of Action)



#### **Bromelain**

Reduces pro-inflammatory cytokines & serotonin; promotes healing via mucin genes; reduces attachment some bacteria.

#### **Menthol**

Antispasmodic for smooth muscle GITract.

#### **Vitamin D\***

Improves mucosal barrier homeostasis and assists microbiome; down-regulation of pro-inflammatory factors.

#### Threonine\*

Amino acid that stimulates colonic healing & mucin synthesis.

#### **Butyrate\***

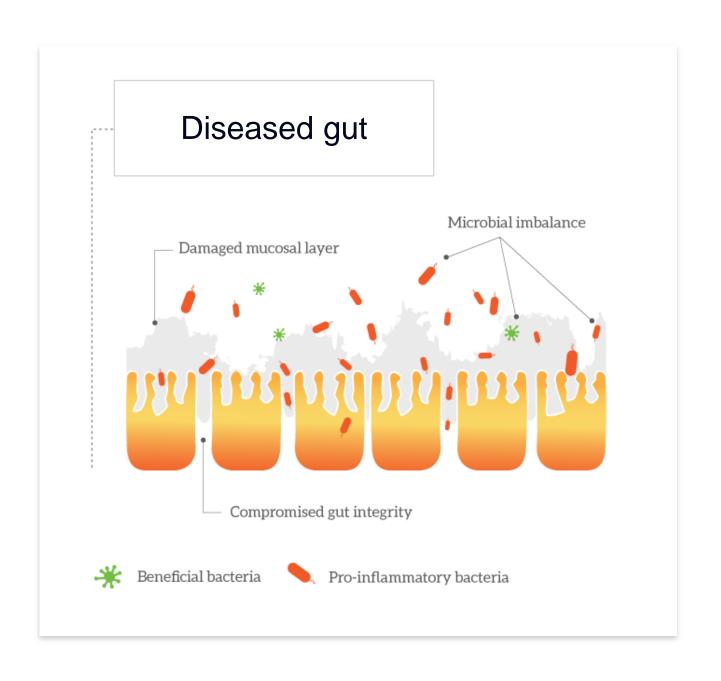
Supports colonocytes as a barrier; reduces pro-inflammatory cytokines & restores homeostasis of microbiome.

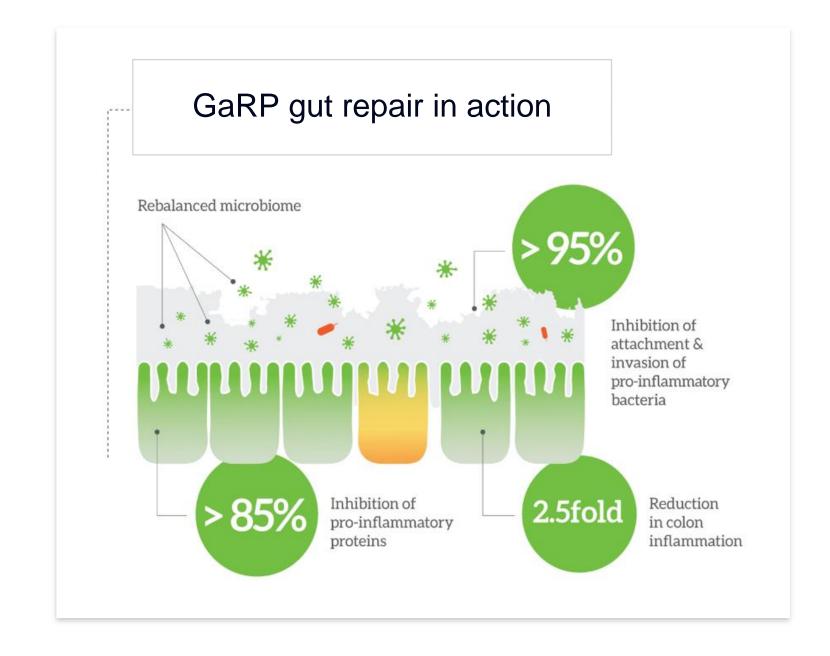


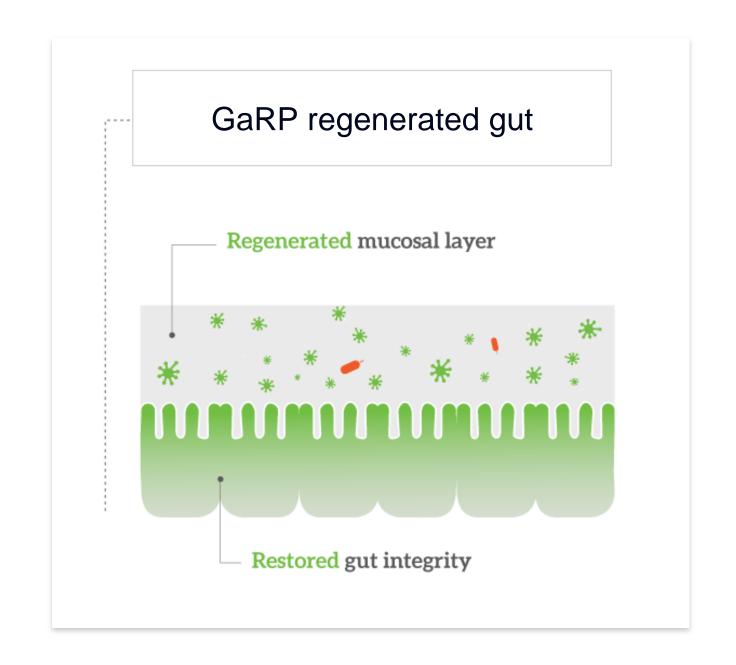


## GaRP - Grounded in scientific evidence

### GARP ADDRESSES THE UNDERLYING FACTORS OF GASTROINTESTINAL DISORDERS







Leaky 'tight junctions' allow pathogens to enter the body causing harm

Increase in mucin gene expression to promote mucosal healing

Restored 'tight junctions' allow the body to resist pathogens and entry of toxins



## Clinical Trial - Irritable Bowel Syndrome

ROBUST CLINICAL TRIAL TO VALIDATE THE EFFICACY OF GARP

61 Patients

Randomised, double blind placebo controlled

Cohort 1

~33% Patients on placebo

Cohort 2

~33% Patients on low dose

Cohort 3

~33% Patients on high dose

3Q 2023

STAGE 2

50-100 Patients (randomised, double-blind placebo-controlled).

Cohort 1

~50% Patients on placebo

Cohort 2

~50% Patients on optimum dose

Final Results Readout

The trial is sufficiently powered to yield statistically significant results vs placebo

2Q 2024



## Clinical Trial - Irritable Bowel Syndrome

ROBUST CLINICAL TRIAL TO VALIDATE THE EFFICACY OF GARP



#### Title

Dose Determination and Efficacy Evaluation of the ReProgramming (GaRP) Gastrointestinal dietary supplement in IBS patients: A Randomized, Double-blind, Placebo controlled virtual clinical trial.

#### **Population**

Males and females 18-65 years of age with irritable bowel syndrome (IBS-SSS score of 175-350 and categorised as IBS on ROME IV criteria), two stages with interim analysis between stages with approx. 60 in Stage 1 and 50-100 anticipated in Stage 2.

#### **Key Milestones & Messages**

Stage 1 Complete – 3Q CY2023 Stage 2 Completion – 2Q CY2024



#### **Endpoints**

- ✓ No Treatment-Related Adverse Events.
- √ Change in IBS-Severity Scoring System between test and placebo groups compared to baseline.
- Safety markers.
- √ Change in IBS quality of life (IBS QoL) points compared to baseline.
- ✓ IBS Adequate Relief (IBS-AR) compared to baseline.
- √ Hospital Anxiety and Depression (HAD) Scale comparing to baseline.

#### **Exploratory Endpoints**

- ✓ Plasma levels of specific inflammatory markers.
- ✓ Use of rescue medication across the study group.
- ✓ Alterations in gut microbiota with respect to diversity, perceived balance and correlation to IBS symptoms including overall wellness.



## Stage 1 Clinical Trial Results

#### **DELIVERED POSITIVE PRIMARY ENDPOINTS**

#### **Stage 1 IBS Clinical Trial**

Number of Patients: Placebo: 20 / Low Dose: 20 / High Dose: 21

#### **Primary Endpoints**

- **Determine optimum Dose:** Low Dose
- ✓ Safety: No serious adverse advents
- ✓ Improvement in Irritable Bowel Severity Scoring System (IBS-SSS)
  - √ 56% reduction after 8 weeks of treatment
  - √ ~20% outperformance over placebo

#### **Secondary Endpoints**

- ✓ Improvement in IBS Quality of Life Score: Low Dose resulted:
  - √ A 14.6pt (8/9 week) & 10.2pt (10/11 week) reduction from baseline
  - √ A 11.4pt (8/9 week) (P=0.10) & 8.3pt (10/11 week) (P=0.29) reduction versus placebo
  - √ A 10pt+ reduction is considered clinically meaningful
- ✓ Improvement in Hospital Anxiety & Depression Score (HADS): Low Dose Resulted in:
  - ✓ Total Score: 4.0pt change from Baseline / 6.0pt Change from Placebo (P<0.001)</p>
  - ✓ Anxiety Score: 3.0pt Change from Baseline / 3.7pt Change From Placebo (P<0.001)</p>
  - ✓ Depression Score: 1pt Change from Baseline / 2.1pt Change From Placebo (P=0.033)

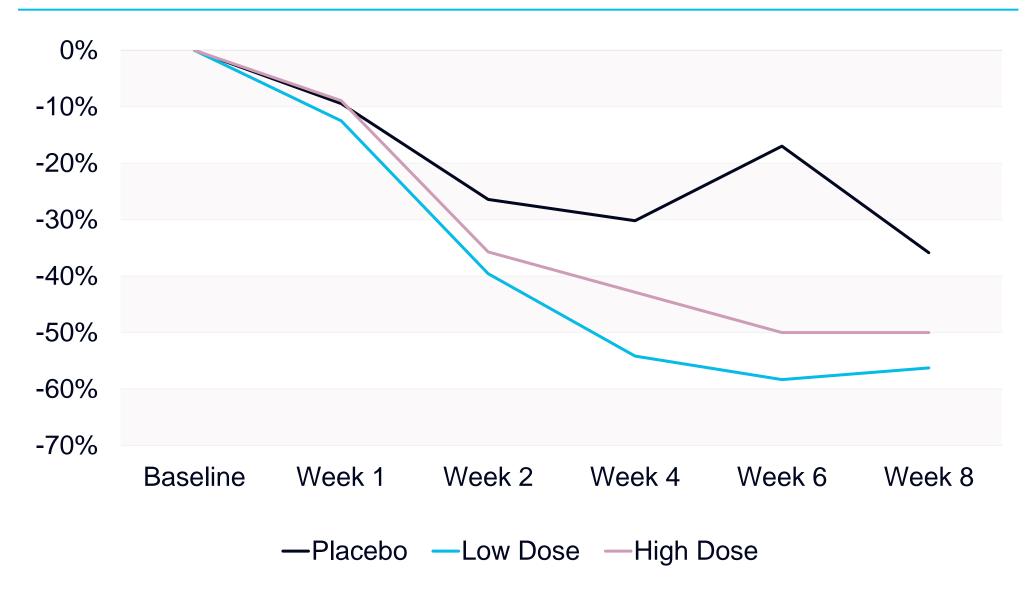
#### **Clinically Meaningfulness & Statistical Significance**

- √ Stage 1 has delivered clinically meaningful results across all endpoints.
- ✓ As is the case with statistical analysis, increasing the population/patients in the trial (as is proposed for stage 2 of the study) is expected to provide statistically significant P values.



	Placebo	Low Dose	High Dose
	n=20	n=20	n=21
Baseline	265	240	280
Week 1	240	210	255
Week 2	195	145	180
Week 4	185	110	160
Week 6	220	100	140
Week 8	170	105	140
Difference in baseline score			
to week 6 score	45	140	140
% Change	-17%	-58%	-50%
Difference in baseline score			
to week 8 score	95	135	140
% Change	-36%	-56%	-50%

#### Median Reduction in IBS-SSS Score - Placebo vs. Low & High Dose



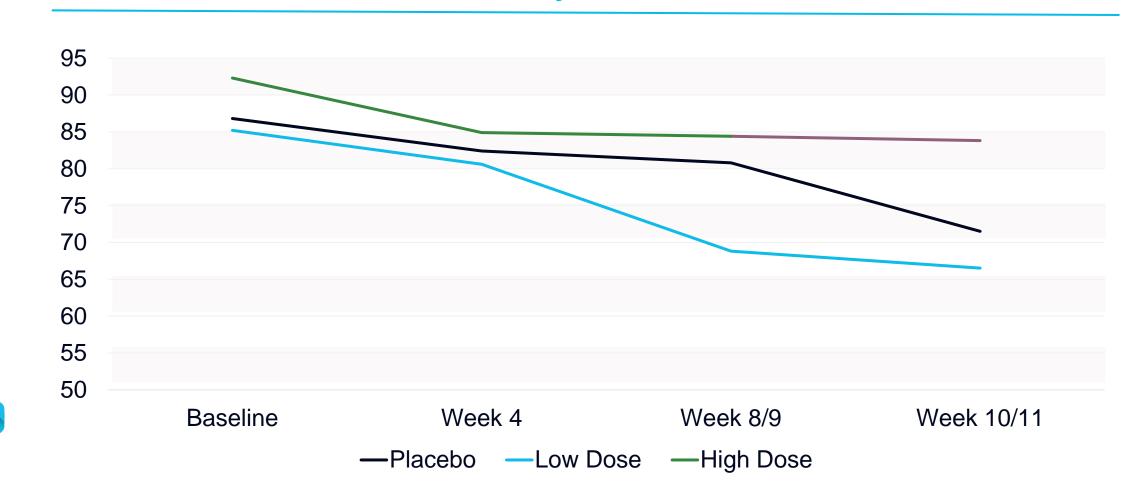
### Stage 1 Clinical Trial Results

#### **DELIVERED POSITIVE SECONDARY ENDPOINTS**

#### **IBS-Quality of Life Score**

	Placebo	Low Dose	High Dose
	N=20	N=20	N=20
Baseline	86.8	85.2	92.3
Week 4	82.4	80.6	84.9
Week 8/9	80.8	68.8	84.4
Week 10/11	71.5	66.5	83.8
Week 8/9 - Change from Baseline	-3.1	-14.6	-11.4
Week 8/9 - Versus Placebo		-11.4 (P=0.10)	-6.2
Week 10/11 - Change from Baseline	-6.0	-10.2	-14.5
Week 10/11 - Versus Placebo		-8.3 (P=0.29)	-8.4

#### **IBS-Quality of Life Score**

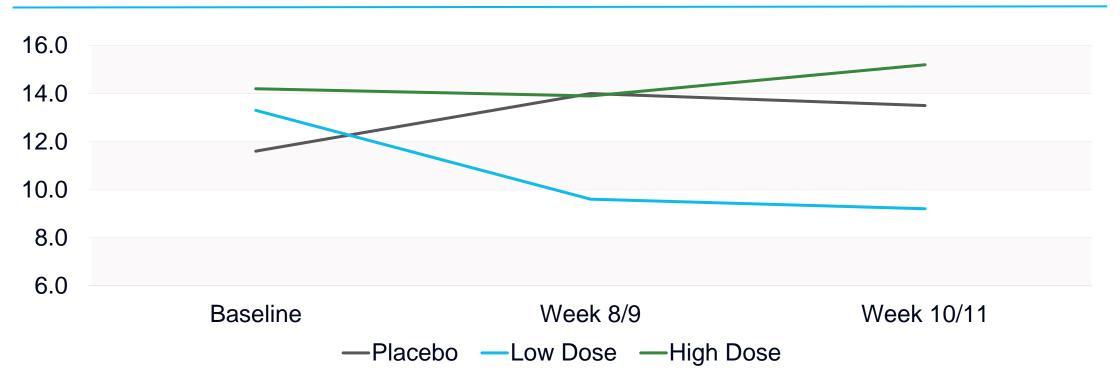


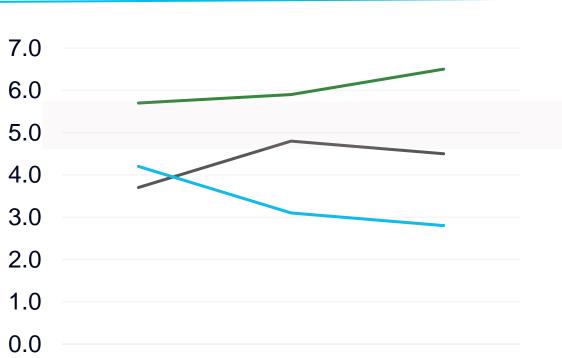


#### **TOTAL SCORE – Hospital Anxiety & Depression Score (HADS)**

Placebo	<b>Low Dose</b>	High Dose
N=20	N=20	N=21
11.6	13.3	14.2
14	9.6	13.9
13.5	9.2	15.2
3.5	-4.0	-0.7
	-6.0 (P<0.001)	-2.4
	N=20 11.6 14 13.5 3.5	N=20       N=20         11.6       13.3         14       9.6         13.5       9.2

#### **TOTAL SCORE – Hospital Anxiety & Depression Score (HADS)**





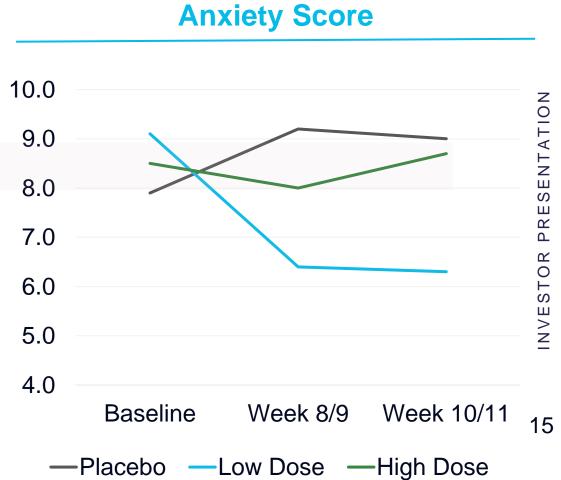
Week 8/9

—Placebo —Low Dose —High Dose

Week 10/11

Baseline

**Depression Score** 

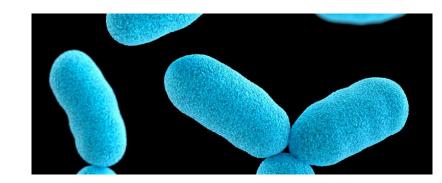


## Gastrointestinal RePrograming (GaRP)

#### A POTENTIAL BREAKTHROUGH FOR GUT HEALTH



Unique knowledge of bromelain's 'fingerprint'



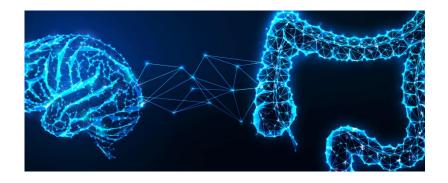
Ingredients are GRAS



Well-characterised proprietary formula



Potential synergistic effects



Gut-Brain Connection

#### FUNCTION WITHIN FORMULATION AND POTENTIAL IMPACT ON SYMPTOMS

	Quality	Inhibitor of	Inhibitor of	Inhibitor of	Inhibitor of	Inhibitor of	Postoros	Influences	Repairs '	leaky gut'	Reduction of	diarrhoea	Modulates visceral	Assists immune	
	Evidence Based Support IBS/IBD	sed attachment / homeostasis of		metabolites of	Reduces gut-wall inflammation	Protection & regeneration of mucosa	Inhibits inflammation & secretory-induced	inflammation & Serotonin		system, brain health & mood	Safety Cautions				
GaRP Formulation	Pre-clinical ++ Pending clinical	<b>√</b>	✓	<b>√</b>	<b>√</b>	✓	✓	<b>√</b>	✓	<b>√</b>	Nil :GRAS				
Iberogast®	✓	X	X	X	✓	✓	✓	Χ	✓	X	Warning: liver				
Probiotics	Controversial- disputed, inconsistent	<b>√</b>	<b>√</b>	✓	<b>√</b>	X	<b>√</b>	Х	✓	✓					
Buscopan®	✓	X	X	X	X	X	X	X	✓	X					



## VESTOR PRESENTATI

## Focused on developing a clinically validated treatment

#### MAJOR GUT HEALTH TREATMENTS LACK CLINICAL EVIDENCE

	Explanation	Evidenced Based Claims	Improvement in IBS-SSS /or Rome II/III	Improving Anxiety & Depression	Improving IBS- QOL Quality of Life	Comments/Key Claims
GaRP Formulation	Multi-component complementary medicine	<b>√</b>	✓	<b>√</b>	<b>√</b>	Clinically meaningful improvement in IBS-SSS score <sup>1</sup> and IBS-Quality of life score (powered for statistical significance in Stage 2) Clinically meaningful and statistically significant improvement in Hospital Anxiety & Depression Score (HADS). <sup>2</sup>
lberogast®	Plant extract formulation	<b>√</b>	✓	X	X	Improvement in Irritable bowel syndrome symptom scores and abdominal pain scale. <sup>3</sup> <b>Product has safety issues.</b>
Align®	Probiotic	<b>√</b>	X	X	✓	Improvement in abdominal pain, Irritable Bowel Syndrome quality of life and composite assessment of IBS symptom relief. <sup>4</sup>
Swisse Daily Digestive Probiotic	Probiotic	X	X	X	X	No clinically meaningful and statistically significant result for probiotic supplementation against placebo for IBS-SSS. <sup>5</sup>
Buscopan®	Spasmolytic (Antispasmodic)	<b>√</b>	X	X	X	Improvement in pain scores among IBS patients, does not address full range of IBS symptons. <sup>6</sup>
Blackmores Digestive Aid	Blend of Enzymes and Herbs	X	X	X	X	Support healthy digestion, no evidence-based claims. <sup>7</sup>
Bromeyal	Enzyme complex (Bromelain)	X	X	X	X	Shown to modulate inflammatory markers in human cells in-vitro, no evidence/claims regarding.8



<sup>2.</sup> GaRP https://anr2.irmau.com/site/pdf/0ae28335-e574-47d2-b5d2-26d26228cc6f/ANR-announces-Positive-Results-from-Stage-1-of-IBS-trial.pdf



<sup>3.</sup> Iberogast https://doi.org/10.1111/j.1365-2036.2004.01859.x

<sup>4.</sup> Align https://pubmed.ncbi.nlm.nih.gov/16863564/

<sup>5.</sup> Swisse https://doi.org/10.3390/jcm12144838

<sup>6.</sup> Buscopan https://pubmed.ncbi.nlm.nih.gov/19337628

<sup>7.</sup> Blackmores Anatara's research did not identify any direct/clinical claims.

<sup>8.</sup> Bromeyal https://doi.org/10.1177/20587384211034686

## Market Opportunity

THE GUT HEALTH MARKET IS SIGNIFICANT, LACKING EFFECTIVE, EVIDENCE-BASED SOLUTIONS

Digestive Health Market to be valued at US\$23.4B in 2030

#### Growth rate of 8.1% to be sustained by:

High prevalence of disease - Obesity, digestive disorders, and lifestylerelated diseases are likely to boost the demand for digestive health products globally.

Poor solutions - There is no clear solution to gut health, resulting in consumers trying a vast array of option.

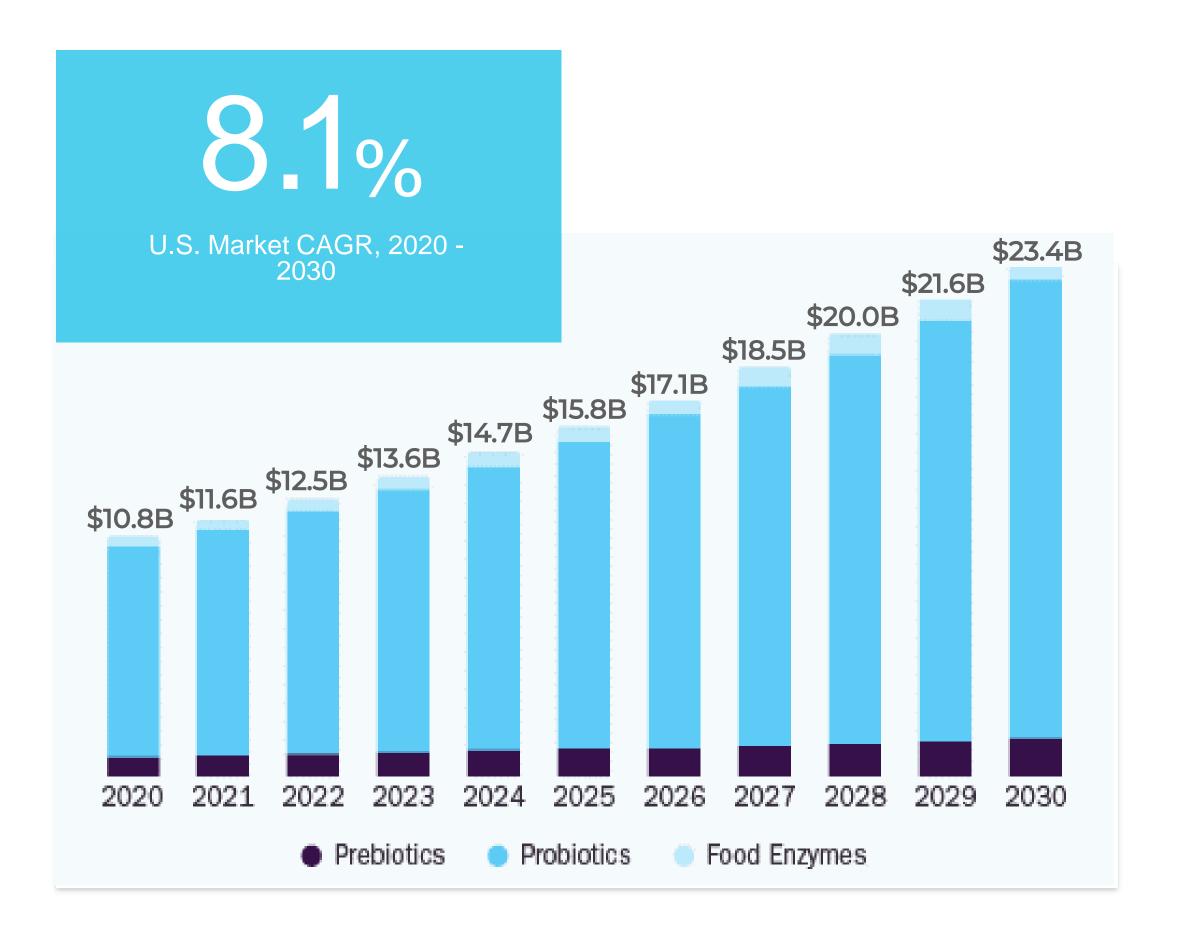
**Strong thematic -** Growing demand for fortifying and nutritional food additives is one of the major factors driving the market.

#### **Growing awareness regarding:**

- 1. Improved health is motivating consumers to pay for better outcomes.
- 2. Maintaining gut health through a holistic approach to healthy living.
- 3. Mainstream interest in product with potential to restore and maintain the gut lining & homeostasis of the microbiome.



U.S. Digestive Health Products Market Size, by ingredient, 2020-2030 (USD Billion)



Source: https://www.grandviewresearch.com/industry-analysis/digestive-health-products-market

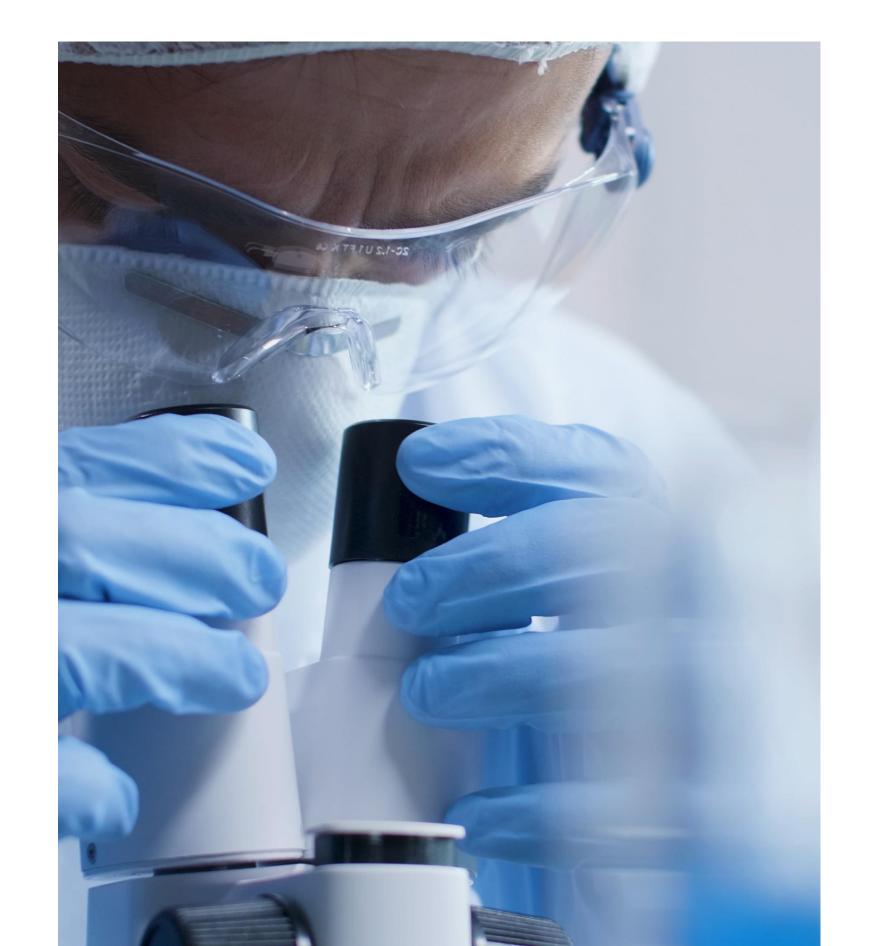


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## Market Opportunity

THE MARKET IS SIGNIFICANT AND LACKING EFFECTIVE, EVIDENCE-BASED SOLUTIONS

Current Large Market/Market Leaders



Probiotic

Align®

Global Sales 2020 \$172M<sup>1</sup>



Proctor & Gamble

Market Capitalisation \$566B



Gastrointestinal Supplements

**Iberogast**®

Germany Sales 2018 €136M<sup>1</sup>



Bayer

Market Capitalisation \$87B



Digestive Supplements Buscopan®

Global Sales 2020 €177M<sup>2</sup>



Sanofi

Market Capitalisation \$206B

sanofi



## Licensing discussions & portfolio diversification

PROGRESSING LICENSING DISCUSSIONS WITH GLOBAL CONSUMER HEALTH COMPANIES FOR GARP



Stage 2 GaRP trial powered for significance

12

Months

Revenue anticipated within 12 months from licensing

#### **Global Partners**

Anatara is experiencing inbound interest from global leaders in the GI field due to the strong evidence based design of the GARP trial. Discussions with global pharma companies are ongoing.

#### **Local Partners**

Partnerships with regional leaders in 'gut' health to leverage local knowledge in registration and marketing of consumer health products and their established market position and infrastructure.

#### **Partnership Opportunity**

Interim Results

Following the positive Stage 1 results, Anatara has reinitiated its discussions with potential commercial partners.

#### **Partnership Opportunity**

Post Trial

Following the anticipated completion of our IBS human trial in 2Q CY2024, Anatara expects to be in a strong position to announce a commercial partnership.



## Advisory Board & DSMB members

ANATARA LIFESCIENCES IS SUPPORTED BY A HIGHLY ACCLAIMED ADVISORY BOARD WITH SIGNIFICANT EXPERIENCE IN GASTROINTESTINAL HEALTH.

Advisory Board members are internationally recognised for their expertise in IBS and IBD with experience ranging from preclinical drug development through to translational research and clinical trials.

The role of the Advisory Board is provide advice on Anatara's research and product development programs in gastrointestinal health.

Members include those pictured; noting the Data Safety Monitoring Board (DSMB) members are Professors Gibson and Rolan and Assoc. Prof. Begun.



Dr Tracey Brown



Associate Professor Rebecca Burgell



Professor Simon Keely



Dr Jeremy Rosenbaum



Associate Professor Jakob Begun



Professor Peter Gibson



Professor Paul Rolan







### Contact Us

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## Significant Corporate Interest

GLOBAL TREND OF RESEARCH & DEVELOPMENT IN GASTROINTESTINAL HEALTH

Several examples of investment and/or collaboration by industry leaders in gastrointestinal health

Company #1	Stock Details	Company #2	Stock Details	Rationale of Deal	Agreement	Further Agreement
Nestlē	SWX: NESN MKT CAP: \$479B AUD	Enterome	Unlisted	Nestle Health Science partnering with French BioTech Enterome to develop and commercialise immunotherapies for food allergies and IBS.	\$67M AUD (Combination of Cash & Equity)	Clinical & Sales Milestone + Royalties on Sales
sanofi	PAR: SAN MKT CAP: \$214B AUD	© caracare	Unlisted	Collaboration to provide a holistic approach to digestive health through combining the gastrointestinal pharmaceutical knowledge of Sanofi with the digital health capabilities of Cara Care.	Agreement to Collaborate	N/A
AstraZeneca	LSE: AZN MKT CAP: \$334B AUD	SERES	NAS: MCRB MKT CAP: \$292M AUD	Collaboration of Seres Therepeutics to research microbiome-based approaches to advance understanding of the efficacy of cancer immunotherapy, utilizing the extensive experience of AstraZenneca in oncology treatment.	Agreement to Collaborate	N/A
SONIC HEALTHCARE	ASX: SHL MKT CAP: A\$14.1Bn	MICROBA	ASX: MAP MKT CAP: A\$94.8M	Sonic Healthcare has completed a strategic investment in Microba to deliver individualised microbiome testing technology through Sonic's existing global pathology network.	\$17.8M Strategic Investment for a 20% stake	\$7.5M option for further investment
ımmur@n	ASX: IMC MKT CAP: A\$18.0m	Ateria Health	Unlisted	Immuron has completed a strategic investment in Ateria Health to expand its offering in the "gut health" market through an IBS drug and expanding their distribution.	\$2.6M Strategic Investment for a 17.5% stake	\$2.55M option for further investment



## Corporate/PE/VC Interest

SIGNIFICANT CORPORATE INTEREST IN THE WELLNESS AND GUT HEALTH SECTORS

#### Corporate Interest in the Wellness space

Acquirer	Stock Details	Target	Stock Details	Rationale of Deal	Agreement
KIRIN	TYO: 2503 MKT CAP: \$19.8B AUD	BLACKMORES°	ASX:BKL	Japanese brewer Kirin acquired Australian vitamins maker Blackmores to diversify away from their extensive alcohol exposure and expand their wellness offering.	A\$1.88B
<u>Biostime</u>	HKG: 1112 MKT CAP: \$1.18B AUD	Swisse	Unlisted	BioStime acquired Australian vitamin maker Swisse Wellness to expand their beauty nutrition offering into new markets and expand their portfolio.	A\$1.67B

#### Private Equity/Venture Capital Interest in Gut Health

Acquirer	Target	Rationale of Deal	Agreement
♣bd-capital	bd-capital SYMPR®VE Venture Capital & Private Equity firm Bd-Capital acquired Symprove, a company that is focused on delivering live and active bacteria to support the gut microbiome.		Undisclosed Sum
MERITECH SEQUOIA LE	H SEQUOIA III Pendulum Pendulum a developer of medical problotics addressing the impalances of the microbiome		US\$54M capital raised for a \$111M total venture investment
middleland	BIOHM	Middleland Capital's VTC Ventures led a US\$7.5M raise for Biohm Health, a microbiome company with products designed to address the roles of bacteria and fungi in gut health.	US\$7.5M capital raise
FIRST Propel VENTURES PARTNERS	Persephone Biosciences	First Bright Ventures & Propel Bio Partners raised US\$15M in a seed financing round for Peresphone Biosciences researching therapeutic approaches for the microbiome including infant health and oncology.	US\$15M seed round
oyster ZBIOTICS®		Oyster Ventures led a seed round of ZBiotics, a probiotic drink that breaks down the byproduct of alcohol responsible for hangovers for US\$2.3M.	US\$2.3M seed round



## Peer Comparison

Company	Stock Code	Company Focus	Market Capitalisation*	Share Price*	Cash Receipts	Net Operating Cash (FY23)
Anatara Lifesciences	ANR:ASX	Anatara is focused on the validation of gastrointestinal products including the clinical validation of GaRP, aimed at restoring and maintaining gut health	<b>\$2.9m</b>	\$0.025	Nil	(\$2.3m)
Nutritional Growth Solutions	NGS:ASX	Nutritional Growth Solutions has developed patented nutritional supplement formulae focusing on the growth development of children	\$2.7m	\$0.011	\$1.5m*	(\$1.6m)*
The Calmer Co International	CCO:ASX	Calmer Co (formerly Fiji Kava) is developing and producing natural solutions for relaxation, anxiety and sleep through a range of functional food and beverages	\$3.2m	\$0.004	\$1.6m	(\$3.3m)
Bod Science	BOD:ASX	Bod Science Limited is developing cannabis derived health & skincare products for general health and wellness, joint/muscle pain, stress/anxiety and sleep.	\$8.9m	\$0.05	\$2.2m	(\$5.7m)
Immuron	IMC:ASX	Immuron produces OTC medication (Travelan & Protectyn) reducing the risk of traveler's diarrhea and gastro-intestinal disorders.	\$18.0m	\$0.079	\$0.6m*	(\$1.0m)*
Bixoyne	BXN:ASX	Bioxyne has developed and globally distributes their patented probiotic strain focusing on immunity and gastrointestinal health and a subsidiary focused on medical cannabis.	\$20.9m	\$0.011	\$3.0m	(\$0.6m)
Vita Life Sciences	VLS:ASX	Vita Life Sciences develops and distributes OTC medicines, as well as complementary and alternative medicines, dietary supplements and health foods	\$89.5m	\$1.66	\$69.0m	\$2.8m*
Microba Life Sciences	MAP:ASX	Microba provides microbiome testing for gastrointestinal diseases and is developing therapeutics in inflammatory bowel disease, cancer Immunotherapy & autoimmune diseases	\$94.7m	\$0.32	\$6.3m	(-\$13.2m)
						*



## Pipeline Programs

DISCUSSING LICENCING POTENTIAL WITH PRODUCERS AND ANIMAL FEED/NUTRITION COMPANIES

#### **Human Health**

- Extensions for GaRP's broad indications for use-trial planning IBD paediatrics (synergistic effect of dose reduction in immune suppressant treatment) & pilot Functional Dyspepsia.
- Actively seeking in-licencing opportunities, M&A assets to broaden portfolio.

#### **Animal health products**

#### **Detach®, ANR-pf and BONIFF**

1.Orally administered, non-antibiotic products. Know how for products with extracts from pineapple stems for potential delivery in water or on feed mix. Unlike antibiotics and zinc oxide, will not contribute to antimicrobial resistance.

#### **Transition away from zinc oxide**

1.Medicinal use of zinc oxide has recently been banned in EU (June 2022 <sup>1</sup>); indications others to follow (recent announcements in Canada <sup>2</sup> and Chile <sup>3</sup>). More than one-third of UK pig farmers were routinely using zinc oxide in pigs after weaning. <sup>4</sup>



- 1. Poultry trials conducted with a leading Australian producer using ANR-pf (poultry) in broilers (chickens for meat production). This most recent trial delivered mixed results that suggested to both parties that a commercial product pathway requires further investigation. (UNE Jan 2021 positive study on efficacy ANR-pf performance broilers subject subclinical NE challenge)
- 2. Murdoch University June 2021 trial report: Pigs fed the BONIFF-SMEC diet performed similarly to pigs fed the SMEC (semi-moist extruded creep transition feed) only diet comprising commercially relevant levels of ZnO and organic acids and phytogenic product.

2. https://www.pigprogress.net/Piglets/Articles/2021/8/Canada-follows-the-EU-with-zinc-oxide-777113E/?utm\_source=tripolis&utm\_medium=email&utm\_term=&utm\_content=&utm\_campaign=pig\_progress 3. https://www.pigprogress.net/World-of-Pigs1/Articles/2021/1/Chiles-meat-sector-deal-for-antibiotics-usage-697251E/?utm\_source=tripolis&utm\_medium=email&utm\_term=&utm\_content=&utm\_campaign=pig\_progress







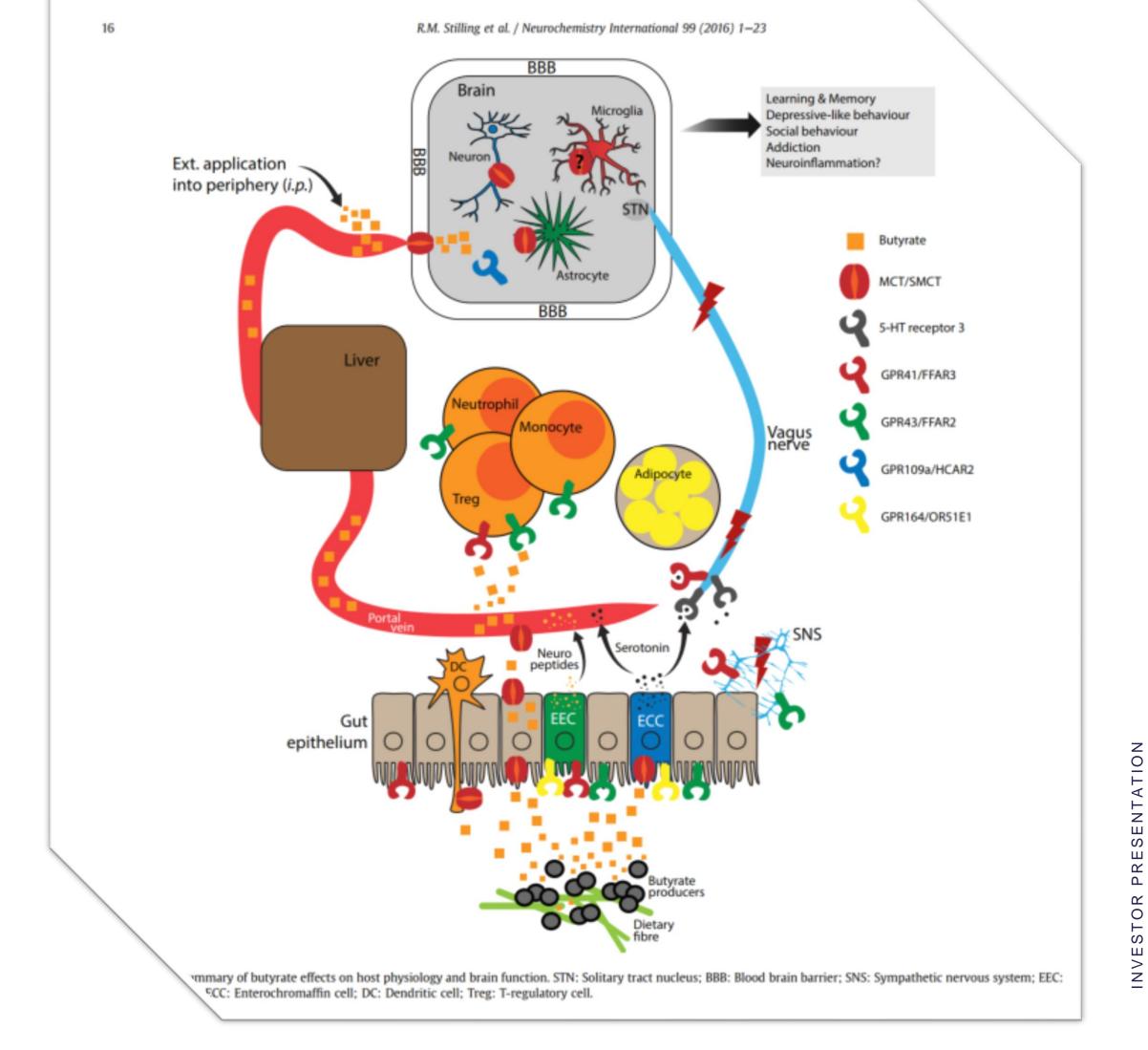
<sup>1.</sup> https://www.ema.europa.eu/en/medicines/veterinary/referrals/zinc-oxide#:~:text=What is zinc oxide?,post-weaning diarrhoea in piglets

## What is the gut-brain connection?

THE BRAIN, FEATURING A RANGE OF GUT MOLECULES

- 1. The gastrointestinal system offers an integrated interface for regulation of various body functions in health and disease.
- 2. The lining of the gut, known as the epithelium, is also the first line of defence against pathogens taken up with the diet. Due to the mutualistic nature of the majority of microbes in the gut, the gut epithelium is also the primary interface for host-microbe crosstalk on all levels of interaction.
- 3. The immune system is trained and regulated by the presence of harmful and beneficial microbes, and products (incl. metabolites).
- 4. Other than the vagus nerve involvement, any gut-brain interaction needs to cross at least two barriers (i.e. the gut epithelium and the blood brain barrier) and permeability through both of these barriers has been shown to be affected by the microbiome.



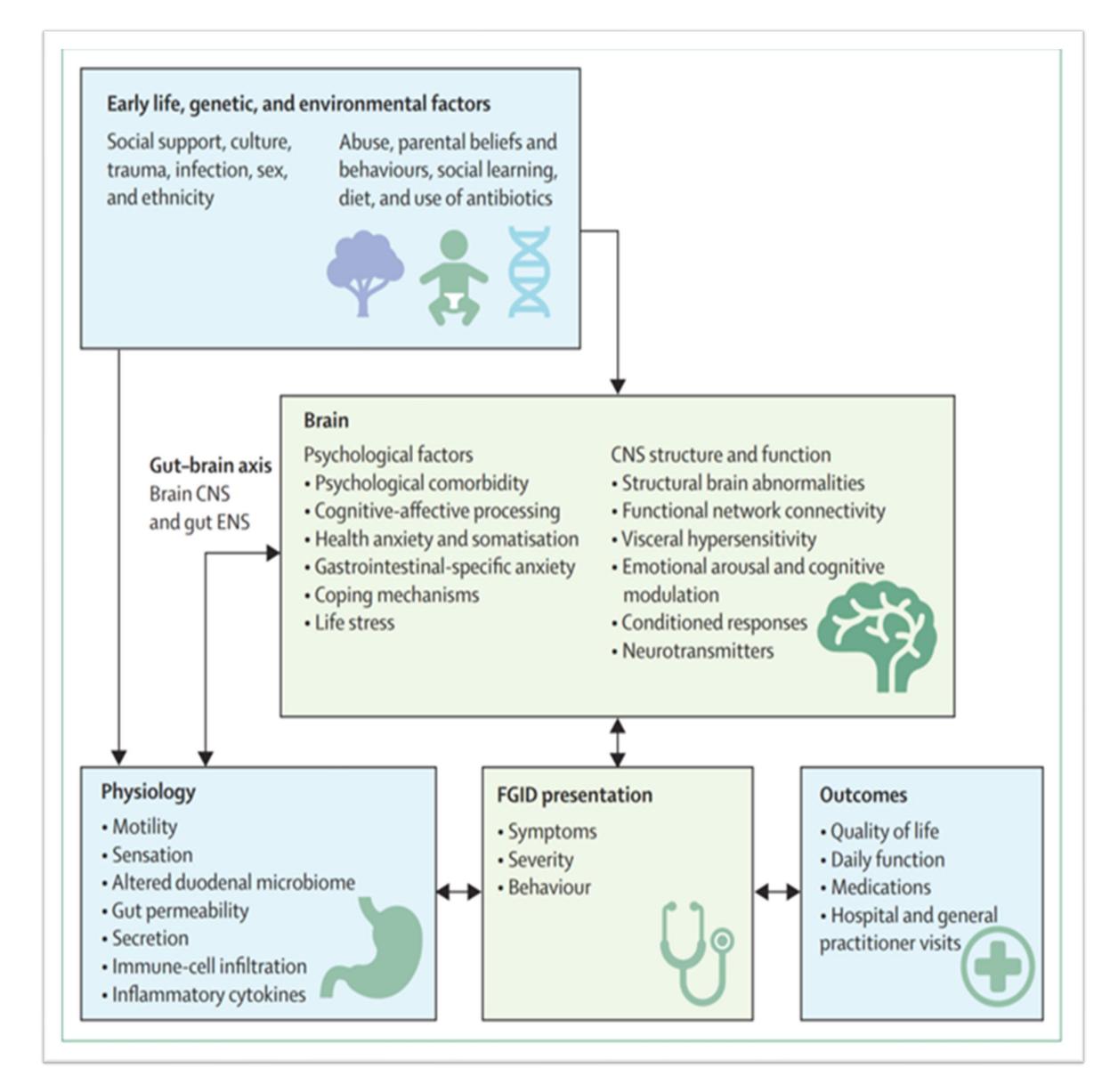




### Pathophysiology

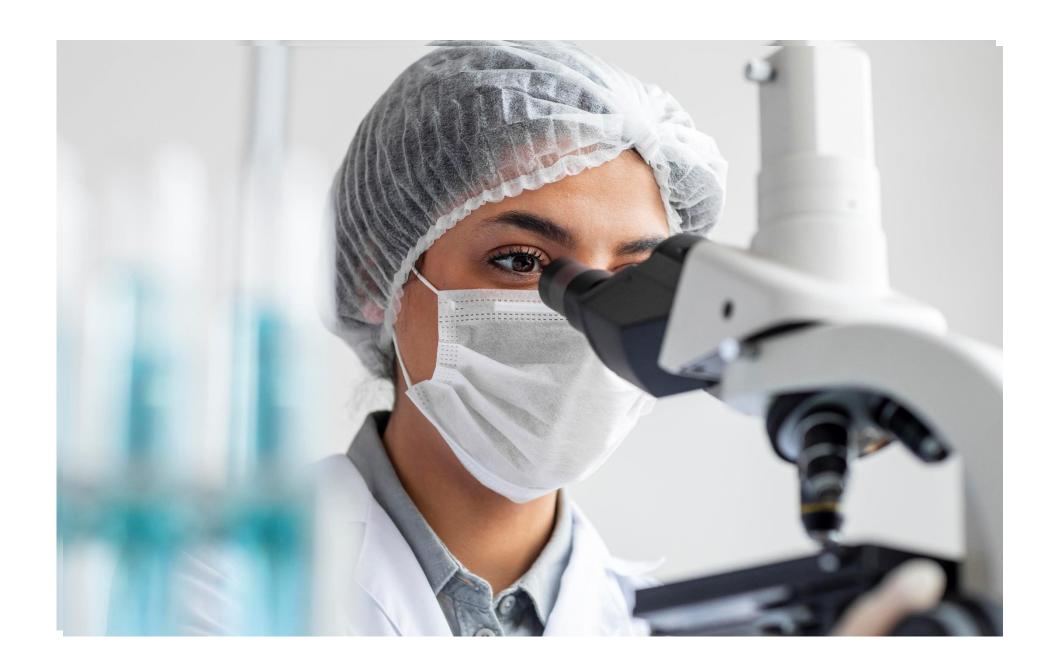
- abnormalities By definition, structural explain Functional Gastrointestinal Disorders (FGIDs) and, on the basis of the biopsychosocial model developed by Engel 34 and adapted by Drossman, 35, 36 they are characterised as complex bidirectional dysregulations of gut-brain interaction, via the gut-brain axis, rather than diseases. FGIDs include IBS, dyspepsia, functional bloating etc.
- Visceral hypersensitivity, abnormal gastrointestinal motility, and psychological disturbances have been recognised to contribute to the pathogenesis of FGIDs for decades, but more recently low-grade intestinal inflammation, increased intestinal permeability, immune activation, and disturbances in the microbiome have been identified, challenging the idea that structural changes are absent entirely.37,38
- The biopsychosocial model articulates illness as holistic and multifactorial, and emphasises the existence of an intimate mind-body connection, facilitated by bidirectional communication between the brain and the gut in FGIDs, which is well accepted.34,35,40
- Emerging data challenge the concept that gut—brain pathways act similarly in all patients with FGIDs. Independent epidemiological studies43-45 suggest that in 50% of cases, FGIDs begin with psychological distress, followed later by gastrointestinal symptoms, whereas in the other 50% of cases gut dysfunction occurs first, and psychological distress follows later.





INVEST

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### Glossary

IBS	Irritable bowel syndrome (IBS) is a condition that affects the colon (also referred to as "large bowel" or "large intestine"). Key symptoms include abdominal pain or discomfort, bloating, and chronic diarrhoea or constipation, or alternating between the two.
IBD	Inflammatory bowel diseases (IBD) are conditions in which the lining of the digestive gastrointestinal tract (GIT) becomes inflamed and damaged. The two common types of inflammatory bowel disease are Ulcerative Colitis and Crohn's Disease. The most common symptoms of inflammatory bowel disease are: diarrhoea, often including blood &/or mucous, abdominal pain, loss of appetite, tiredness, fever, weight loss.
ANR-pf	Anatara's proprietary enriched formulation for poultry in water, designed to allow the full delivery of key additives in a quick flexible dosing method on-farm even when stock illness is a concern.
Bromelain	Bromelain is a mixture of proteases extracted from pineapple stems. Specific proteases within Bromelain have wide activities influencing the GIT(Gastrointestinal tract)including anti-attachment, anti-secretory and reduction of pro-inflammatory factors. Collectively, these proteases provide a broad spectrum of activity from reducing the attachment of some diarrhoea-causing organisms to blocking inflammatory and adverse secretory factors.
BONIFF	Anatara's recently developed in-feed bromelain-based formulation for weaner piglets.
GaRP	Gastrointestinal ReProgramming. Anatara's GaRP product is a multi-component ,coated complementary medicine that has been designed to address underlying factors associated with chronic gastrointestinal conditions, such as IBS and IBD. This products is designed to assist restoration and maintenance of the GIT lining and the homeostasis of the microbiome.
Leaky Gut Syndrome	"Leaky Gut Syndrome" is a proposed condition associated with increased intestinal permeability that allows harmful substances (e.g toxins, bacteria) into the circulation which triggers adverse health effects through systemic inflammation and immune reactions. It is not considered a medical diagnosis and, while a very relevant consideration, it is not clear if it is a symptom or underlying cause of associated chronic diseases.
GMP	Good Manufacturing Practice. GMP describes a set of principles and procedures that when followed helps ensure that therapeutic goods are of high quality.
GRAS	Generally Recognised as Safe. A GRAS ingredient is an ingredient that has undergone safety evaluations by experts and has been proven not to cause harm when used as intended.

