

The background of the slide is a blue-tinted photograph of a healthcare professional, likely a nurse or doctor, wearing a stethoscope and holding the hand of a patient. The patient's hand has a medical sensor attached to it. The overall tone is professional and caring.

# Acacia Pharma Group plc

2020 Year End Results Presentation

March 29, 2021

Delivering innovative products to  
enhance surgical patients' recovery

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# Acacia Pharma Group – a transformational year in 2020

## **BARHEMSYS® – FDA approved for PONV, launched late 2020**

- Broad label for prevention and treatment of Post-Operative Nausea and Vomiting (PONV)
- Only agent approved for ‘Rescue’ of ~16m patients p.a. in US with PONV after generic antiemetics fail<sup>1</sup>
- PONV “rescue” is an estimated \$2.7 billion annual total addressable market<sup>2</sup>

## **Commercialization began in 2H 2020 and showing good progress**

- Strong sales, marketing, medical affairs, commercial operations teams in place
- Sales team deployed against ~900 initial targeted hospital accounts since mid-October
- Drug shortages and surgery backlog creating pent-up demand (heightened by Covid)

## **BYFAVO™ – In-licensed, FDA approved for procedural sedation, launched Jan 2021**

- Indicated for procedural sedation in adults – launch benefitting from shared value proposition
- Key target: 40m procedures a year in US, including 25m GI procedures<sup>3</sup>
- Estimated >\$1.5 billion annual total addressable market<sup>4</sup>

## **Strong corporate progress**

- Balance sheet strengthened with debt and equity financings
- Gary Gemignani appointed CFO
- Named BEL Small Company of the Year for second consecutive year

<sup>1</sup> Calculations based on available procedural data, applied Compound Annual Growth Rate and quantitative market research responses as follows: National Hospital Discharge Survey, 2006; National Survey of Ambulatory Surgery, 2006 (as revised in 2009); Source Healthcare; NCHS 2005; Life Science Strategy Group, LLC Market Research; Apfel et al., 2004. <sup>2</sup> Based on the calculations in (1) multiplied by the number of doses per patient at a WAC price of \$85 per 10mg dose. <sup>3</sup> iData Research, US Market Report Procedure Numbers for Gastrointestinal Endoscopic Devices February 2019; American Society of Anesthesiologists, Practice Guidelines for Moderate Procedural Sedation and Analgesia 2018; and Quantitative Market Research prepared by The Link Group for Cosmo Technologies (March 2019). <sup>4</sup> Based on the calculation in (4) multiplied by the number of doses per patient at a WAC price of \$39 per dose.

# We have been able to make significant progress in a very challenging operational environment caused by the global pandemic

“ The OR accounts for up to 65% of hospital profit margin, so this missing volume is cutting deeply into cash flow and net income. ”  
– Becker’s Hospital Review<sup>1</sup>

## COVID-19 impact on hospitals and surgical centers

- Non-essential surgeries cancelled creating a significant backlog
- Physical access to hospital decision-makers even more restricted

## Hospital profits have suffered and need to be restored quickly

- Surgeries and procedures are major contributors to hospital profits
- Providers need to dramatically increase throughput to regain lost profits

## We believe our products and team are ideally positioned to help

- BARHEMSYS and BYFAVO can help improve patient throughput – both are now even more relevant and of greater interest to customers due to COVID
- Our strong relationships are helping us gain access to key decision-makers

<sup>1</sup> How to rebuild surgical revenue after COVID-19, **Becker’s Hospital Review**, accessed via <https://www.beckershospitalreview.com/strategy/how-to-rebuild-surgical-revenue-after-covid-19-even-if-you-just-lost-60-of-your-or-volume.html>





# **BARHEMSYS®**

(amisulpride for injection)

**The first and only FDA-approved  
product for PONV rescue treatment<sup>1</sup>**

<sup>1</sup> FDA labels for other recommended treatments do not include treatment after failed prophylaxis. Treatment agents recommended by Society for Ambulatory Anesthesiology Consensus Guidelines (2014). Habib et al (2019): no agent has previously been shown in a prospective trial to be more effective than a placebo for treating PONV for patients who have failed prophylaxis.



# BARHEMSYS® and the PONV Opportunity

## BARHEMSYS Addresses the major unmet need in PONV

- BARHEMSYS is the **only FDA-approved drug for PONV rescue** after failed prophylaxis<sup>1</sup>
- Dopamine D<sub>2</sub>/D<sub>3</sub> antagonist with **broad, differentiated label**
- Offers significant **economic savings** to hospital vs current standard of care

## Large US market opportunity in PONV<sup>2</sup>

- Estimated ~65m eligible surgeries annually, ~49m patients receive preventative antiemetics<sup>4</sup>
- Estimated ~16m patients still develop PONV and need rescue treatment<sup>3</sup>
- **Total addressable PONV rescue market estimated at ~\$2.7 billion/year<sup>4</sup>**

## Concentrated market, addressable by small direct sales force

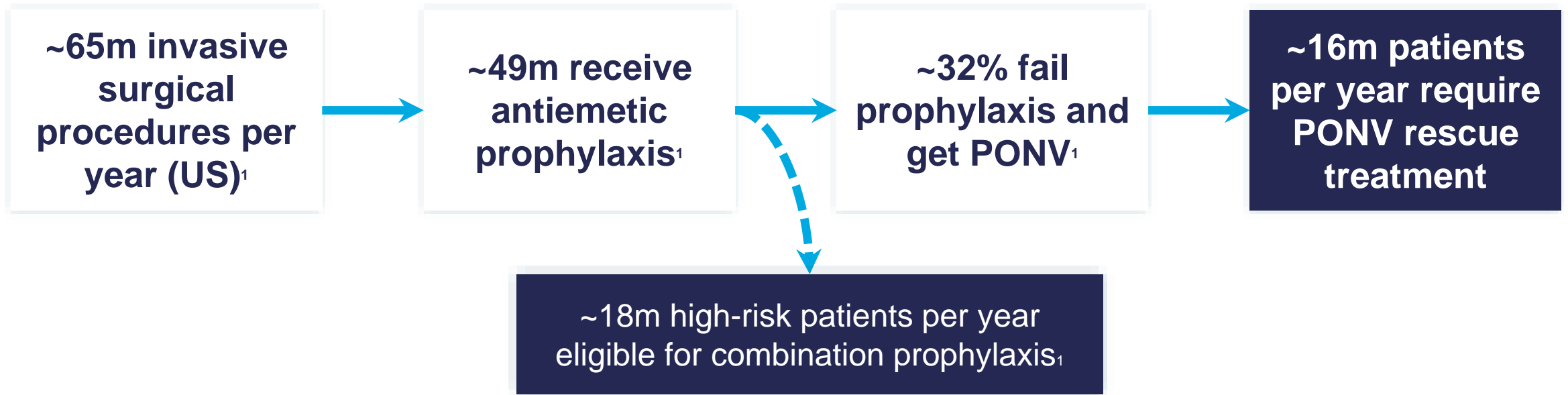
- Estimated 80% of surgeries carried out in ~1,200 hospitals<sup>5</sup>
- 30 sales territories address accounts with greatest immediate opportunity
- **Sales team began customer engagement in mid-October**

## High gross profit, secure supply chain and worldwide rights

- **Cost of goods ~10% of sale price**
- Substantial product inventory to minimize supply risk
- **Worldwide rights and exploring out-licensing opportunities in OUS markets**

<sup>1</sup> FDA labels for other recommended treatments do not include treatment after failed prophylaxis, <sup>2</sup> This is the belief of the Company. <sup>3</sup> Company market size estimates based upon: 2006 National Hospital Discharge Survey; National Survey of Ambulatory Surgery, 2006 (as revised in 2009); LSSG Quantitative Research November 2014; Apfel, NEJM 2004. Calculations based on available procedural data, applied Compound Annual Growth Rate and quantitative market research responses. <sup>4</sup> Based upon WAC price of \$85 per 10mg rescue dose with, on average, 2 rescue doses per patient, and the above estimates. <sup>5</sup> Symphony Health, Source Non Retail, August 2017 - July 2018.

# Targeting PONV Rescue Market



**Total estimated addressable market in PONV rescue  $\approx$  \$2.7B per year**

Secondary market in combination prophylaxis in highest-risk patients, estimated to be worth \$765M per year<sup>2</sup>

1 Company market size estimates based upon: 2006 National Hospital Discharge Survey; National Survey of Ambulatory Surgery, 2006 (as revised in 2009); LSSG Quantitative Research November 2014; Apfel et al, 2004. Calculations based on available procedural data, applied Compound Annual Growth Rate and quantitative market research responses. 2 Based on WAC price of \$85 per 10 mg rescue dose and average 2 rescue doses per patient; \$42.50 per 5 mg prophylaxis dose.

# BARHEMSYS<sup>®</sup> is the Only FDA-Approved Product for PONV Rescue

“ When PONV prophylaxis has failed, patients should receive antiemetic treatment from a different pharmacological class to the PONV prophylaxis ”  
– Consensus Guidelines<sup>1</sup>

Antiemetic	Can't redose	Efficacy issues	Safety issues	Current share of rescue patients <sup>4</sup>
Ondansetron	X <sub>1</sub>			69%
Dexamethasone	X <sub>1</sub>	X <sub>2</sub>		19%
Metoclopramide		X <sub>1</sub>	X <sub>1</sub>	19%
Promethazine			X <sub>1</sub>	11%
<b>BARHEMSYS</b>	✓ <sub>3</sub>	✓ <sub>3</sub>	✓ <sub>3</sub>	<b>INTENT TO PRESCRIBE<sup>4</sup></b> <b>61%</b>



# BARHEMSYS® – Compelling Clinical and Commercial Proposition

## Only FDA-approved product for PONV rescue<sup>1</sup>

- Only drug proven in randomized clinical trial to work in PONV rescue<sup>2</sup>
- Excellent safety profile demonstrated in clinical studies
- Also demonstrated to be effective for prevention

## Throughput and health economic benefits

- 35-minute reduction in PACU stay
- 6-hour reduction in hospital stay
- Offers significant economic savings to hospital vs current standard of care

## Convenient, easy to use, high margin product

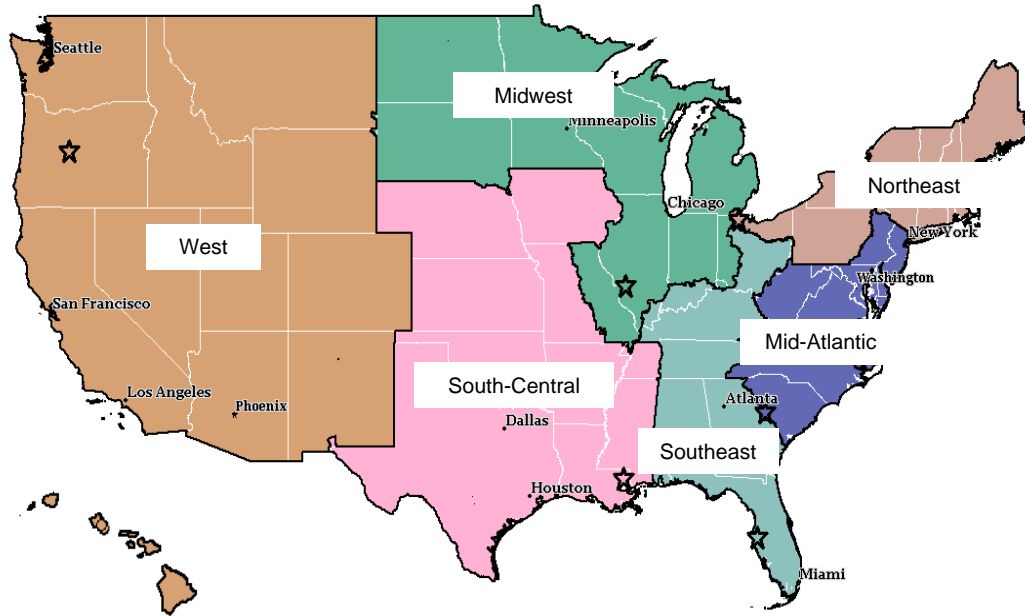
- 5-year room temperature shelf-life
- Fits in auto-dispensing (Pyxis™) machines
- Cost of goods ~10% of sale price

## Helps post-COVID pressure to get through surgical backlog

- Shorter time in PACU (recovery room) allows increased surgical throughput
- Better efficacy and safety means better recovery and patient experience



# Highly Experienced Commercial Team is Driving Formulary Adoption



Team has direct experience successfully launching OFIRMEV into same market to same key customers

## Field Force Design

- 1 SVP of Field Commercial Teams
- 1 National Accounts Group Leader
- 1 National MSL Group Leader
- 6 Sales Regions each with:
  - 1 RBD
  - 1 MSL
  - 1 National Account Director
  - 5 Hospital Territory Managers

### Commercial Leadership Team

**28+**

Years avg industry

**60+**

Launches

### Sales Leadership Team

**22**

Years avg industry

**18+**

Years hospital

### National Accounts Team

**24**

Years avg industry

**21+**

Years hospital

### Medical Science Liaison Team

**22**

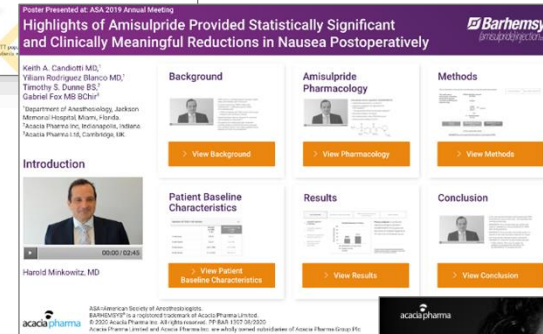
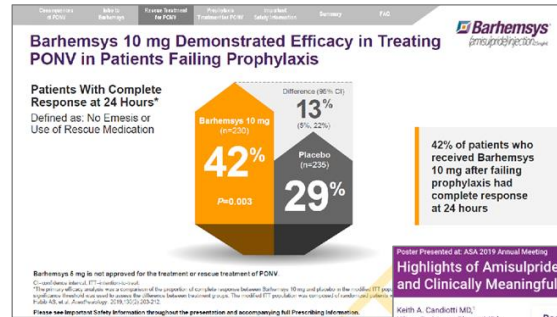
Years avg industry

**10+**

Years as MSL

# BARHEMSYS – Robust Commercial Engagement Creating Traction

- Robust WorseThanPain/ Disease state campaign highlighting unmet need in PONV (worsethanpain.com)
- Medical congress engagements: ~400 symposia attendees and 550 leads generated
- Peer-to-peer programs: 60+ programs with ~450 attendees (live, virtual and on demand) since launch
- All promotional materials available both physically and digitally to meet customers needs
- Digital brand engagements in the most trusted anesthesia publications with 89% readership (51K members)



# Formulary Reviews of BARHEMSYS Began 2H 2020

## Post-COVID pressures improve our access to key decision-makers

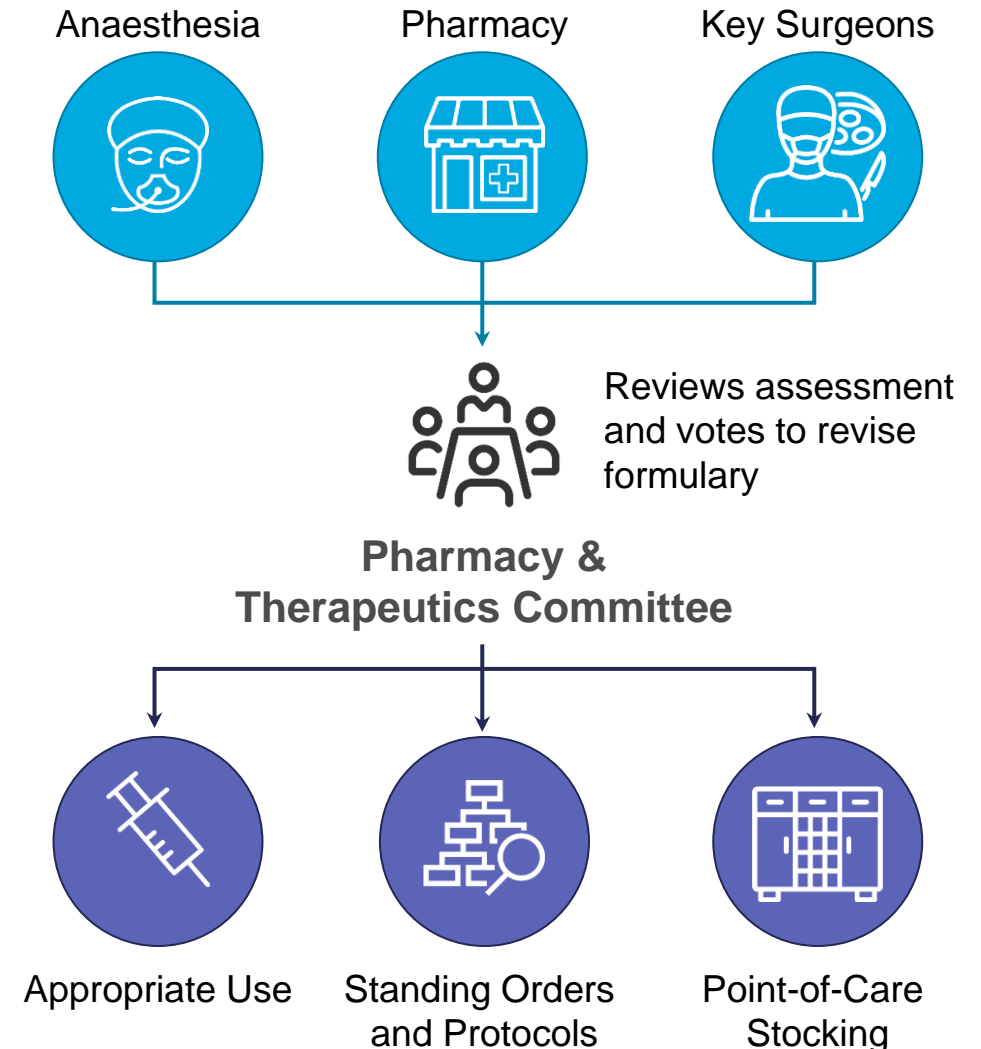
- Surgery backlogs are major issues for US hospitals
- Value of BARHEMSYS and BYFAVO in improving throughput facilitates our access to key decision-makers

## Demonstrate unmet need, appropriate use & health economic benefit

- Highlight current unmet needs with existing products
- Educate on efficacy and safety data for our products
- BARHEMSYS and BYFAVO can provide economic savings to hospital vs current standard of care

## Formulary adoption and pull-through

- P&T Committee approval typically takes 9-12 months on average
- Protocols, standing orders and point-of-care supplies drive sales pull-through





# Formulary Progress to Date on BARHEMSYS



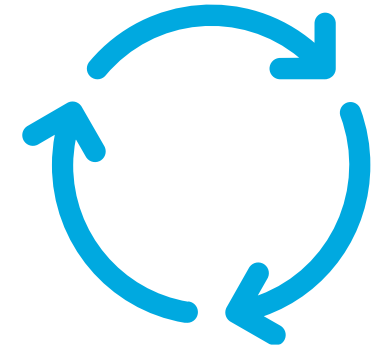
**120 accounts on  
formulary  
>85% win rate**



**141 additional  
accounts have  
scheduled a date  
for formulary  
review**



**>70 accounts  
have already  
ordered the  
product**



**38% of ordering  
accounts have  
placed repeat  
orders**



**BYFAVO™**

(remimazolam) for injection

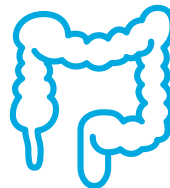
Rapid onset/offset procedural  
sedative with favorable safety profile

# Procedural Sedation Market Opportunity



**~40 million<sup>1</sup>**

procedures each year  
requiring sedation



**~25 million**

GI procedures performed  
each year<sup>2</sup>



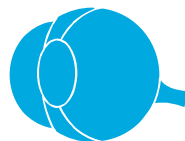
**>80%**

GI procedures have sedation administered  
by an anesthesia provider<sup>3</sup>



**>6 million**

Interventional  
Radiology<sup>4</sup>



**~4 million**

Ophthalmic  
Procedures<sup>5</sup>



**~1 million**

Bronchoscopy<sup>6</sup>



**~1.5 million**

Cosmetic/  
Plastic Surgery<sup>7</sup>

**Total addressable market in procedural sedation >\$1.5B/year<sup>8</sup>**

1 Calculations based on available procedural data, applied Compound Annual Growth Rate and quantitative market research responses. 40 million includes other opportunities: CC (MHA National) EP (dicardiology), Dental (areadentist), Plastic Surgery (American Society of Plastic Surgeons), ECT (MHA National). 2 iData Research, US Market Report Procedure Numbers for Gastrointestinal Endoscopies Nov 2016; CDC website. 3 Quantitative Market Research prepared by The Link Group for Cosmo Technologies (March 2019). 4 Report on interventional Radiology November/December 2007. 5 American Medical Association 2011. 6 iData Bronchoscopy 2019 report. 7 American Society of Plastic Surgeons 2018. 8 Based on the calculation in iData Research, US Market Report Procedure Numbers for Gastrointestinal Endoscopic Devices February 2019; American Society of Anesthesiologists, Practice Guidelines for Moderate Procedural Sedation and Analgesia 2018; and Quantitative Market Research prepared by The Link Group for Cosmo Technologies (March 2019) multiplied by the number of doses per patient at a WAC price of \$39 per dose.

# BYFAVO Addresses Unmet Need in Procedural Sedation

## Propofol

*fast acting but  
significant safety issues<sup>1,2</sup>*

- Rapid onset and offset anesthetic with narrow therapeutic index<sup>1</sup>
- **Dose-related cardiorespiratory depression**, pain at injection site<sup>1</sup>
- Non-linear dosing effects due to individual variability<sup>4</sup>
- **Needs continuous monitoring by anesthesiologist, no reversal agent<sup>2</sup>**
- Lipid formulation susceptible to bacterial contamination<sup>4</sup>

## Midazolam

*better safety profile but  
longer onset and recovery<sup>1,2</sup>*

- Benzodiazepine sedative, reversible by flumazenil<sup>1</sup>
- **Slower onset and offset<sup>2,3</sup>**
- Metabolized by cytochrome system; individual variability affects sedation<sup>1</sup>
- Active metabolite can accumulate and cause prolonged sedation<sup>2</sup>
- **Risk of respiratory depression<sup>1</sup>**

## BYFAVO

*fast acting AND favorable  
safety profile<sup>1,2</sup>*

- **Rapid onset/offset<sup>1,2,3</sup>** benzodiazepine
- Rapid biotransformation into inactive metabolites via non-specific tissue esterases – not dependent on liver enzymes<sup>1</sup>
- **Predictable behavior, no pharmacokinetic drug interactions<sup>5</sup>**
- **Reliable sedation, reliable safety profile<sup>1</sup>**
- Reversible by flumazenil<sup>1</sup>



# BYFAVO™ – Compelling Clinical and Commercial Proposition

## Approved with a broad label

- Indicated for **procedural sedation in adults** in procedures lasting 30 mins or less
- Substantial clinical data package shows **compelling efficacy and safety** in colonoscopies and bronchoscopies, including least fit patients

## Throughput and health economic benefits

- Rapid onset/offset enables shorter procedure times and greater patient throughput for hospitals and surgical centers compared to other recommended treatments

## Commercial synergy with BARHEMSYS

- Target prescribers: **anesthesiologists and proceduralists in hospitals and ambulatory surgery centers**

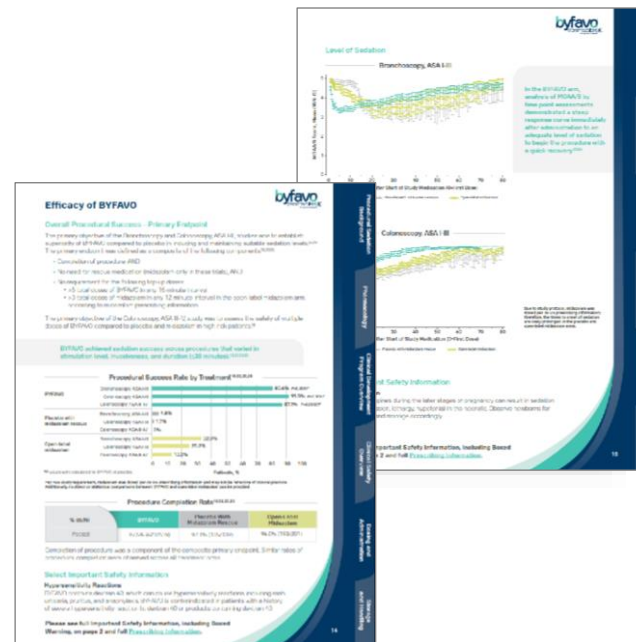
## Helps post-COVID pressure to alleviate procedural backlog

- **Shorter procedure times allow increased procedural volumes**
- Both midazolam and propofol currently on FDA drug shortages list



# BYFAVO – Launching the First New Sedative in the US for 20 years

- On demand promotional programming available, featuring anesthesia, GI, and pulmonary investigators
- Full congress plan across Anesthesia, Pharmacy, and including key GI and Pulm congresses
- All promotional materials available in both digital and physical format
- Non-personal promotional efforts, including direct mail are being initiated
  - Direct email, banner ads, paid search and expanded web site



**byfavo**  
Ezetimibe

**A Short-Acting Sedative for Procedures 30 Minutes or Less**

An Intelligible Sedative

Byfavo is a short-acting benzodiazepine indicated for the induction and maintenance of procedural sedation in adults undergoing procedures lasting 30 minutes or less.

**Preparation and Administration**

Once removed from packaging, protect vials from light. To reconstitute BYFAVO for injection:

- Add 8.2 mL sterile 0.9% NaCl injection, USP to the vial, direct the stream of solution toward the wall of the vial.
- Gently swirl the vial (do not shake) until the contents are fully dissolved.

The reconstituted product will deliver a final concentration of 2.5 mg/mL. Reconstituted BYFAVO can be stored in the vial for up to 8 hours under controlled room temperature at 20°C to 25°C (68°F to 77°F). Discard any unused portion.

**Indication**

BYFAVO is a benzodiazepine indicated for the induction and maintenance of procedural sedation in adults undergoing procedures lasting 30 minutes or less.

**Important Safety Information**

**WARNING: PERSONNEL AND EQUIPMENT FOR MONITORING AND RESUSCITATION AND RISK OF CONCOMITANT USE WITH OPIOID ANALGESICS**

Please see full Boxed Warning on the next page.

**Dosing and Administration**

**Recommended Dosage for Procedural Sedation**

	For Adult Patients	For ASA II and IV Patients
<b>Induction</b>	Administer 5 mg intravenously over a 1 minute time period.	Administer 2.5 mg to 5 mg intravenously over 1 minute based on the general condition of the patient.
<b>Maintenance (as needed)*</b>	Administer 2.5 mg intravenously over 15 seconds.	Administer 1.25 mg to 2.5 mg intravenously over 15 seconds.

\*At least 2 minutes must elapse prior to administration of any supplemental dose.

**Preparation and Administration**

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**Administration With Other Fluids**

BYFAVO has been shown to be compatible with the following fluids:

- 0.9% NaCl injection, USP
- 5% Dextrose injection, USP
- 20% Dextrose injection, USP
- 5% Dextrose and 0.45% NaCl injection, USP
- Ringer's Solution

**BYFAVO is incompatible with:**

- Lactated Ringer's Solution
- Acetated Ringer's Solution

Do not mix BYFAVO with other drugs or fluids prior to administration. BYFAVO compatibility with other agents has not been adequately evaluated.



# Formulary Progress to Date on BYFAVO



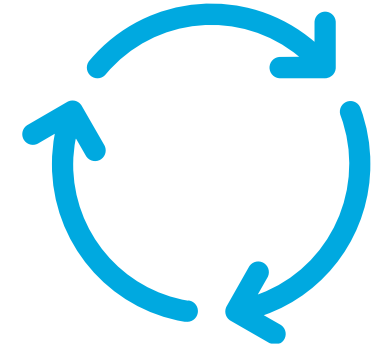
**7 accounts on  
formulary in eight  
weeks of launch**



**36 additional  
accounts have  
scheduled a date  
for formulary  
review**



**8 accounts have  
already ordered  
the product**



**38% of ordering  
accounts have  
placed repeat  
orders**

# Financials



## Loss after tax for the year ended 31 December 2020 of \$33.5m (2019: \$22.8m)

- The operating loss increased by \$8.5m to \$30.9m (2019: \$22.4m), reflecting the investment in our US commercial infrastructure and product launch preparations
- R&D expenses \$0.1m (2019: \$3.9m) with the reduction reflecting lower R&D activities on completion of BARHEMSYS clinical program, together with a \$1.4m credit on reversing certain inventory provisions on the approval of BARHEMSYS
- Sales and marketing expenses \$19.4m (2019: \$14.0m) reflecting increased activities leading up to the planned launch of BARHEMSYS and BYFAVO
- General and administrative expenses \$11.6m (2019: \$4.4m) with 2020 costs higher as a result of fundraising activities, personnel costs and amortisation of intangibles

# Non-GAAP Reconciliation of Operating Expenses and Loss

	2020				2019			
\$millions	Research and development expenses	Sales and marketing expenses	General and administrative expenses	Total	Research and development expenses	Sales and marketing expenses	General and administrative expenses	Total
OPEX As reported	\$ (0.1)	\$ (19.4)	\$ (11.6)	\$ (31.1)	\$ (3.9)	\$ (14.0)	\$ (4.4)	\$ (22.4)
Inventory provision reversal	(1.4)	-	-	(1.4)	1.4	-	-	1.4
Share based compensation	0.0	1.3	1.4	2.7	0.1	1.6	0.7	2.4
Amortization*	-	-	3.1	3.1	-	-	-	-
Adjusted OPEX	\$ (1.5)	\$ (18.1)	\$ (7.1)	\$ (26.7)	\$ (2.4)	\$ (12.4)	\$ (3.7)	\$ (18.6)
Operating Loss as Reported				\$ (30.9)				\$ (22.4)
Adjusted Operating Loss				\$ (26.5)				\$ (18.6)

\* Relates to amortization of BYFAVO intangible asset

## Cash Runway Extended Through Q2 2022

- **Cash and cash equivalents as of 31 December 2020 of \$46.7m (2019: \$17.0m)**
  - €20m (~\$22m) equity investment from Cosmo as part of BYFAVO license agreement
  - €25m (~\$31m) loan from Cosmo related to BYFAVO approval
  - €25m (~\$29m) equity financing in August 2020
- **Additional equity financing in February 2021 with gross proceeds of €27m (~\$33m)**

# Summary



# In Summary

Our products address **unmet medical needs** in therapeutic areas with **large total addressable market opportunities** (~\$2.7B in PONV rescue, >\$1.5B in procedural sedation)

The team made **tremendous progress in 2020** despite a very challenging operating environment due to the global pandemic

We have seen a **great response from customers in the early stages of launch adding our products to formulary** with positive customer feedback on both products so far

While the environment remains very dynamic, **we believe we should be able to benefit from a relaxation of restrictions and a rebound in procedural volumes in the second half of this year**

We believe we have **the right team, with the right experience** to continue to drive the successful commercial launch for both products this year

# Q & A