
UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
Washington, D.C. 20549

FORM 6-K

REPORT OF FOREIGN PRIVATE ISSUER PURSUANT TO RULE 13a-16 OR 15d-16 UNDER THE SECURITIES EXCHANGE ACT OF 1934

For the month of January, 2024.

Commission File Number: 001-40530

GH Research PLC
(Exact name of registrant as specified in its charter)

Joshua Dawson House
Dawson Street
Dublin 2
D02 RY95
Ireland
(Address of principal executive office)

Indicate by check mark whether the registrant files or will file annual reports under cover of Form 20-F or Form 40-F:

Form 20-F

Form 40-F

INFORMATION CONTAINED IN THIS REPORT ON FORM 6-K

GH Research PLC (the "Company") will hold one-on-one investor meetings during the 42nd Annual J.P. Morgan Healthcare Conference, which is scheduled to take place from January 8-11, 2024, in San Francisco, California.

On January 5, 2024, the Company made available an updated investor presentation on its website. A copy of the investor presentation is attached hereto as Exhibit 99.1.

The fact that this presentation is being made available and furnished herewith should not be deemed an admission as to the materiality of any information contained in the materials. The information contained in the presentation is being provided as of January 5, 2024, and the Company does not undertake any obligation to update the presentation in the future or to update forward-looking statements to reflect subsequent actual results.

Exhibit No.	Description
99.1	Corporate Presentation for January 2024

SIGNATURE

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned, thereunto duly authorized.

Date: January 5, 2024

GH Research PLC

By: /s/ Julie Ryan
Name: Julie Ryan
Title: Vice President, Finance



Corporate Presentation

GH Research PLC (NASDAQ: GHR)

January 2024

Disclaimer Regarding Forward-Looking Statements



This presentation has been prepared by GH Research PLC ("GH Research") for informational purposes only and not for any other purpose. Nothing contained in this presentation is, or should be construed as, a recommendation, promise or representation by the presenter or GH Research or any director, employee, agent, or adviser of GH Research. This presentation does not purport to be all-inclusive or to contain all of the information you may desire.

This presentation does not constitute an offer to sell or the solicitation of an offer to buy securities, nor shall there be any sale of these securities in any state or jurisdiction in which such offer, solicitation or sale would be unlawful prior to registration or qualification under the securities laws of any such state or jurisdiction.

This presentation contains forward-looking statements, all of which are qualified in their entirety by this cautionary statement. Many of the forward-looking statements contained herein can be identified by the use of forward-looking words such as "may", "anticipate", "believe", "could", "expect", "should", "plan", "intend", "estimate", "will", "potential" and "ongoing", among others, although not all forward-looking statements contain these identifying words.

Any statements contained herein that do not describe historical facts are forward-looking statements that are based on management's expectations and are subject to certain factors, risks and uncertainties that may cause actual results, outcomes, timing and performance to differ materially from those expressed or implied by such statements. These factors, risks and uncertainties include, but are not limited to: the costs and uncertainties associated with GH Research's research and development efforts; the inherent uncertainties associated with the conduct, timing and results of nonclinical and clinical studies of GH Research's product candidates; GH Research's expectations related to the clinical hold on the GH001 IND, including plans and expectations for progressing any nonclinical programs and any other work to lift the clinical hold, the timing required to lift such clinical hold and for discussions with the FDA and the outcomes and resolution of such discussions; GH Research's ability to obtain, maintain, enforce and defend issued patents; the adequacy of GH Research's capital resources, the availability of additional funding and GH Research's cash runway; and other factors, risks and uncertainties described in GH Research's filings with the U.S. Securities and Exchange Commission.

Except as otherwise noted, these forward-looking statements speak only as of the date of this presentation, and GH Research undertakes no obligation to update or revise any of such statements to reflect events or circumstances occurring after this presentation. Because forward-looking statements are inherently subject to risks and uncertainties, some of which cannot be predicted or quantified and some of which are beyond GH Research's control, you should not rely on these forward-looking statements as predictions of future events. The events and circumstances reflected in any such forward-looking statements may not be achieved or occur and actual results could differ materially from those projected in the forward-looking statements. GH Research cautions you not to place undue reliance on the forward-looking statements contained in this presentation.



Seeking Ultra-Rapid, Durable Remissions in Depression

Pipeline



Stage of Development

PROGRAMS	INDICATION	PRECLINICAL	PHASE 1	PHASE 2a	PHASE 2b	PHASE 3	CURRENT STATUS
GH001 <i>Mebufotenin (5-MeO-DMT) for inhalation administration</i>	Treatment-Resistant Depression (TRD)	Complete			Ongoing		Phase 2b RDBPC trial initiated (GH001-TRD-201)
	Bipolar II Disorder* (BDII)	Complete			Ongoing		Phase 2a POC trial initiated (GH001-BD-202)
	Postpartum Depression (PPD)	Complete			Ongoing		Phase 2a POC trial initiated (GH001-PPD-203)
GH002 <i>Mebufotenin (5-MeO-DMT) for i.v. administration</i>	Psychiatric or Neurological Disorder	Complete					Phase 1 HV trial completed (GH002-HV-105)
GH003 <i>Mebufotenin (5-MeO-DMT) for nasal administration</i>	Psychiatric or Neurological Disorder	Ongoing					Pre-clinical development ongoing

Complete

Ongoing

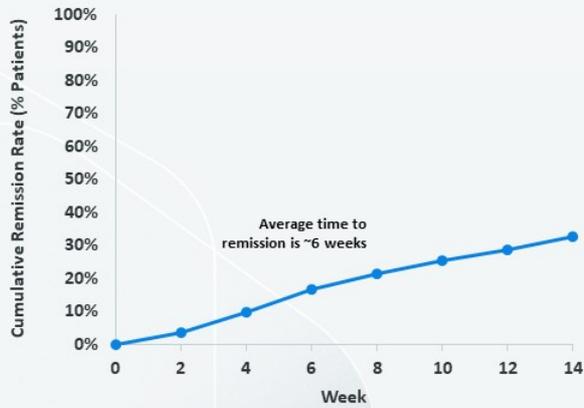
*Bipolar II disorder with a current major depressive episode
 5-MeO-DMT, 3-Methoxy-N,N-Dimethyltryptamine; i.v., intravenous; RDBPC, Randomized, Double-Blind, Placebo-Controlled; POC, Proof-of-Concept; HV, Healthy Volunteer



The Problem for Patients with Depression

Established Therapies are **Slow-Acting**

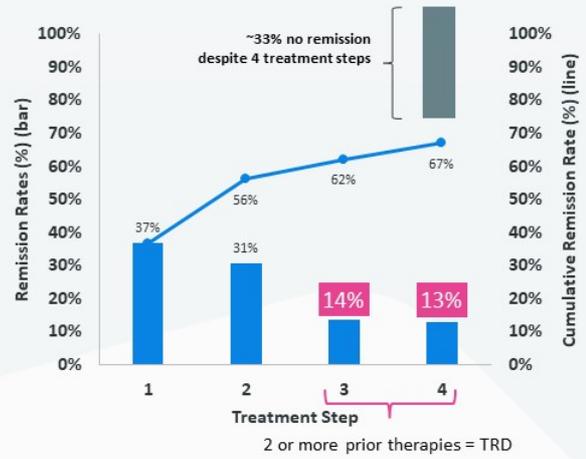
(STAR*D study, Remission Rate Over Time, Treatment Step 1 = Citalopram)



Adapted from Trivedi et al., Am J Psychiatry 2006 and Rush et al., Am J Psychiatry 2006
TRD, Treatment-Resistant Depression

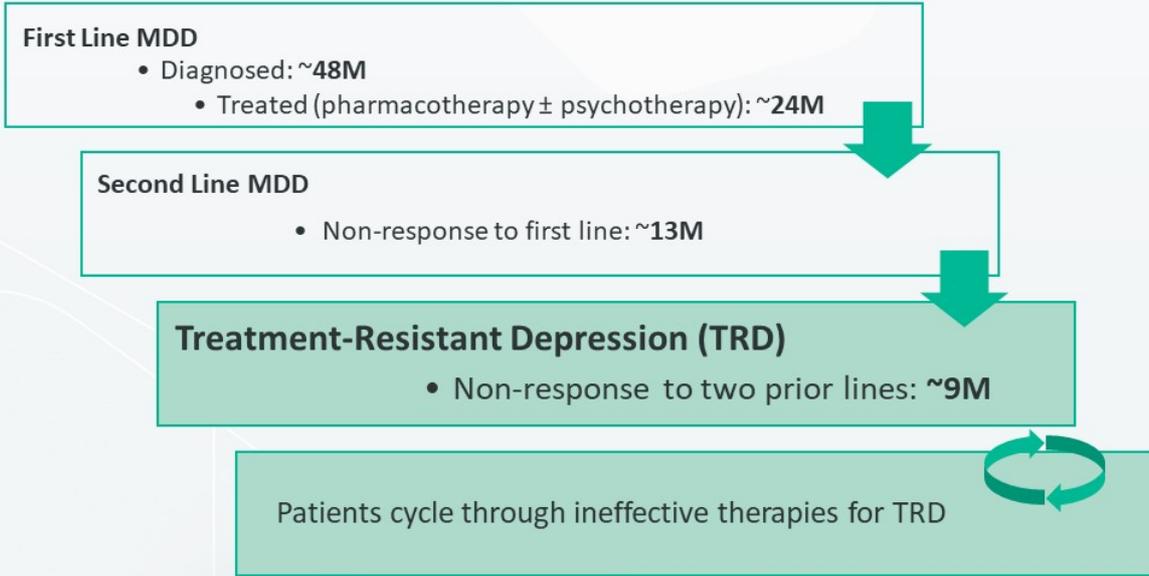
... Remission Rates in TRD < 15%

(STAR*D study, Remission Rates Treatment Steps 1 to 4)



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Large and Open Depression Market in the EU and US



Company estimates based on: <https://www.nimh.nih.gov/health/statistics/major-depression.shtml>; Wittchen et al., The size and burden of mental disorders and other disorders of the brain in Europe 2010. *European Neuropsychopharmacology* (2011); Rush et al., Acute and Long-Term Outcomes in Depressed Outpatients Requiring One or Several Treatment Steps: A STAR*D Report. *Am J Psychiatry* 2006
MDD, Major Depressive Disorder

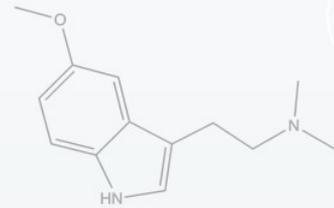
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Mebufotenin (5-MeO-DMT) and GH001



Mebufotenin (5-Methoxy-N,N-Dimethyltryptamine, 5-MeO-DMT)

- Naturally-occurring psychoactive substance from tryptamine class
- **Highly potent** agonist on 5-HT1A and 5-HT2A receptors



Mebufotenin (5-MeO-DMT)

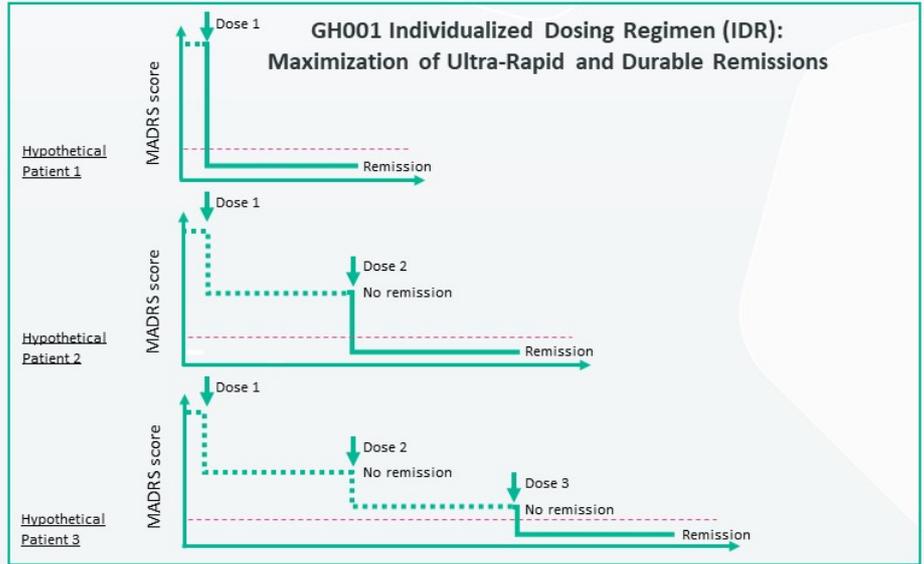
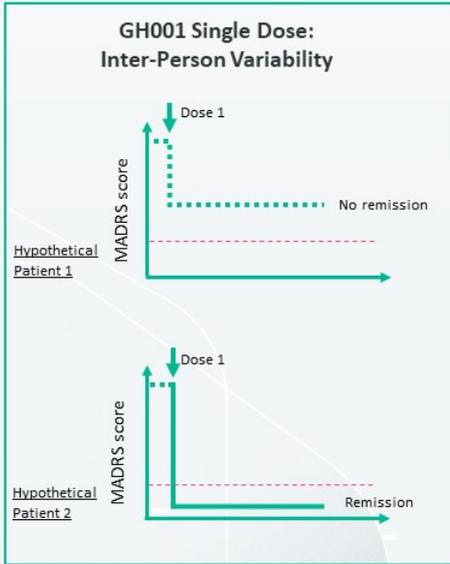
GH001 (Mebufotenin administration via a proprietary pulmonary inhalation approach)

- **Psychoactive effects with ultra-rapid onset** (within seconds) **and short duration** (5 to 30 min)
- **High propensity to induce peak experiences (PE), which may be a surrogate marker for therapeutic effects**
- **Intraday individualized dosing regimen (IDR) for maximization of ultra-rapid and durable remissions**
- **Single visit initial treatment**, with no structured psychotherapy
- Potential for **convenient and infrequent retreatment**

Foundational IP



GH001 – Individualized Dosing Regimen (IDR) for Maximization of Ultra-Rapid and Durable Remissions



MADRS, Montgomery-Åsberg Depression Rating Scale



Phase 1 Trial in Healthy Volunteers GH001-HV-101

(Completed)

Clinicaltrials.gov ID: NCT04640831

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9

Design of Phase 1 Trial in Healthy Volunteers (GH001-HV-101)



HV, Healthy Volunteer; PE, Peak Experience; IDR, Individualized Dosing Regimen

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¹The PE Scale averages answers scored by the subject by marking a visual analogue scale between 0 and 100 for the following three questions:
1. How intense was the experience; 2. To what extent did you lose control;
3. How profound (i.e., deep and significant) was the experience? 10

Part A (Single Dose) and Part B (IDR) – Safety



Study Safety Group review

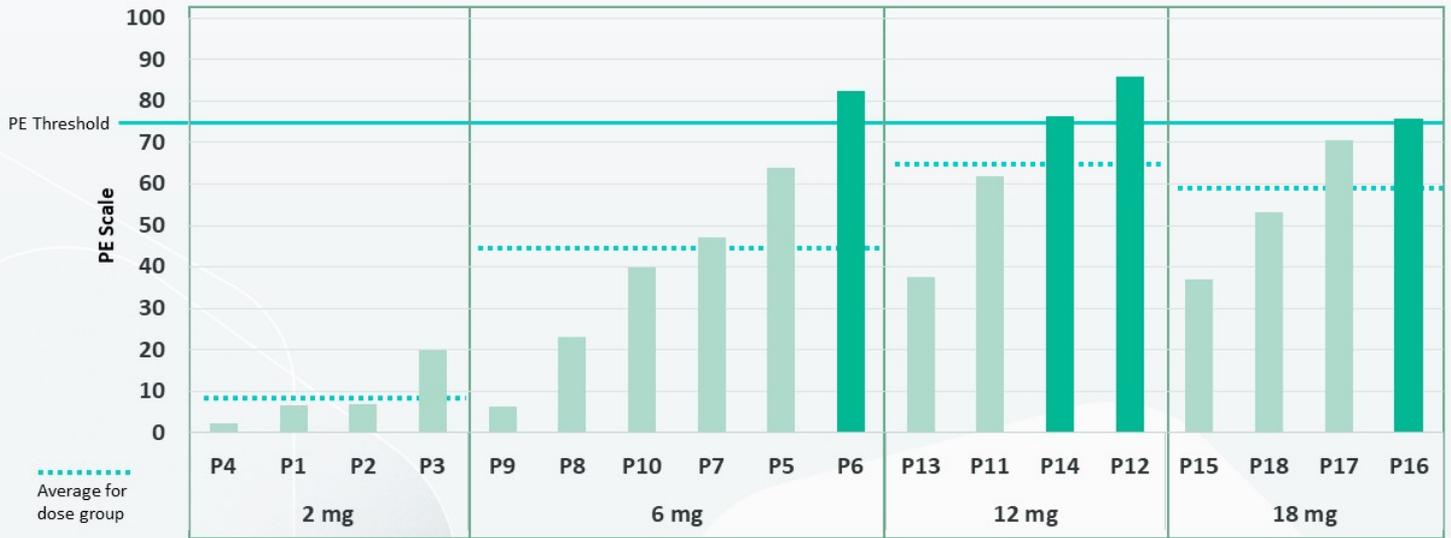
- No SAEs
- All ADRs mild, except two moderate (*)
- All ADRs resolved spontaneously
- Inhalation well-tolerated
- No noteworthy changes in vital parameters, except for temporary, non-clinically relevant increase in heart rate and blood pressure shortly after administration of GH001
- No clinically significant changes in safety laboratory analyses, psychiatric safety assessments or measures of cognitive function

ADRs	Part A (Single Dose)				Part B (IDR)
	2 mg (n=4)	6 mg (n=6)	12 mg (n=4)	18 mg (n=4)	IDR ¹ (n=4)
MedDRA Preferred Term	Number of Events				
Abnormal dreams				1	
Anxiety		1	1		
Clumsiness		1			
Confusional state		1			
Euphoric mood		1			
Fatigue				1	1*
Feeling hot		1			
Flashback				1	
Hallucination				1	
Head discomfort					1
Headache		2		1	1
Heart rate increased			1*		
Hyperacusis				1	
Insomnia				1	
Mental fatigue				1	
Nausea	2	1		1	2
Vision blurred	1				

SAE, Serious Adverse Event; ADR, Adverse Drug Reaction, an adverse event with a relationship code to the investigational product of definite, probable, or possible, or where code is missing; IDR, Individualized Dosing Regimen

*6 mg (n=1); 6-12 mg (n=2); 6-12-18 mg (n=1)

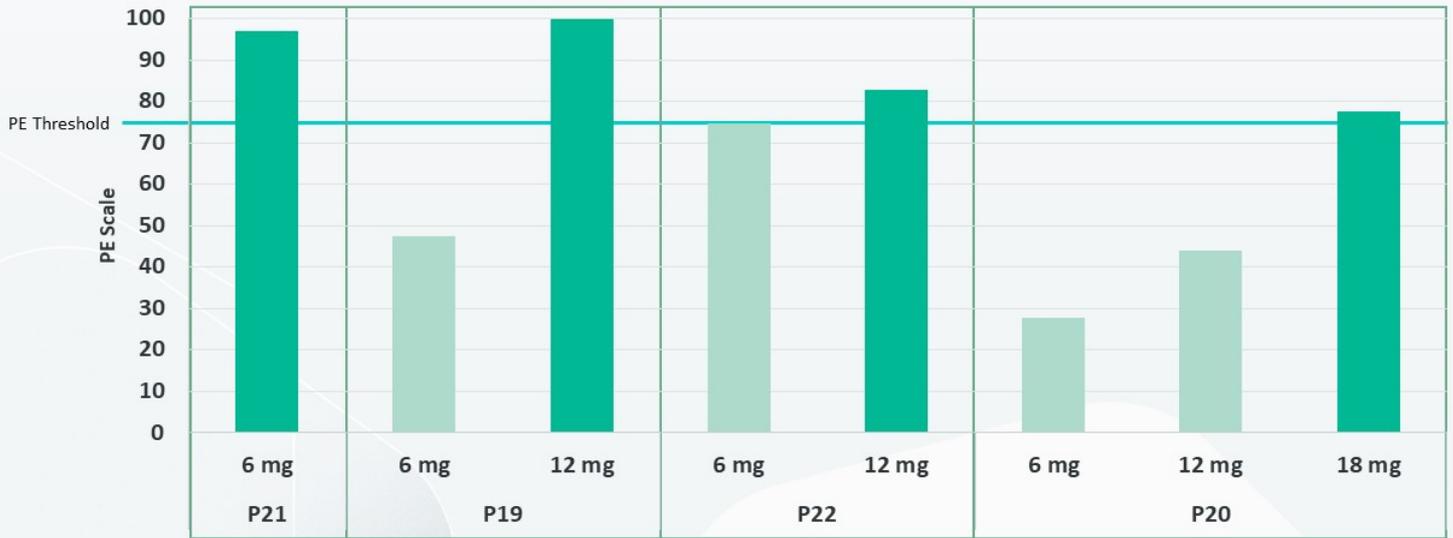
Part A – Peak Experience (PE) Dose-Effects and Inter-Person Variability



PE, Peak Experience

Part B – Peak Experience (PE)

Effect of Intraday Individualized Dosing Regimen (IDR)



PE, Peak Experience



Phase 1/2 Trial in Treatment-Resistant Depression GH001-TRD-102

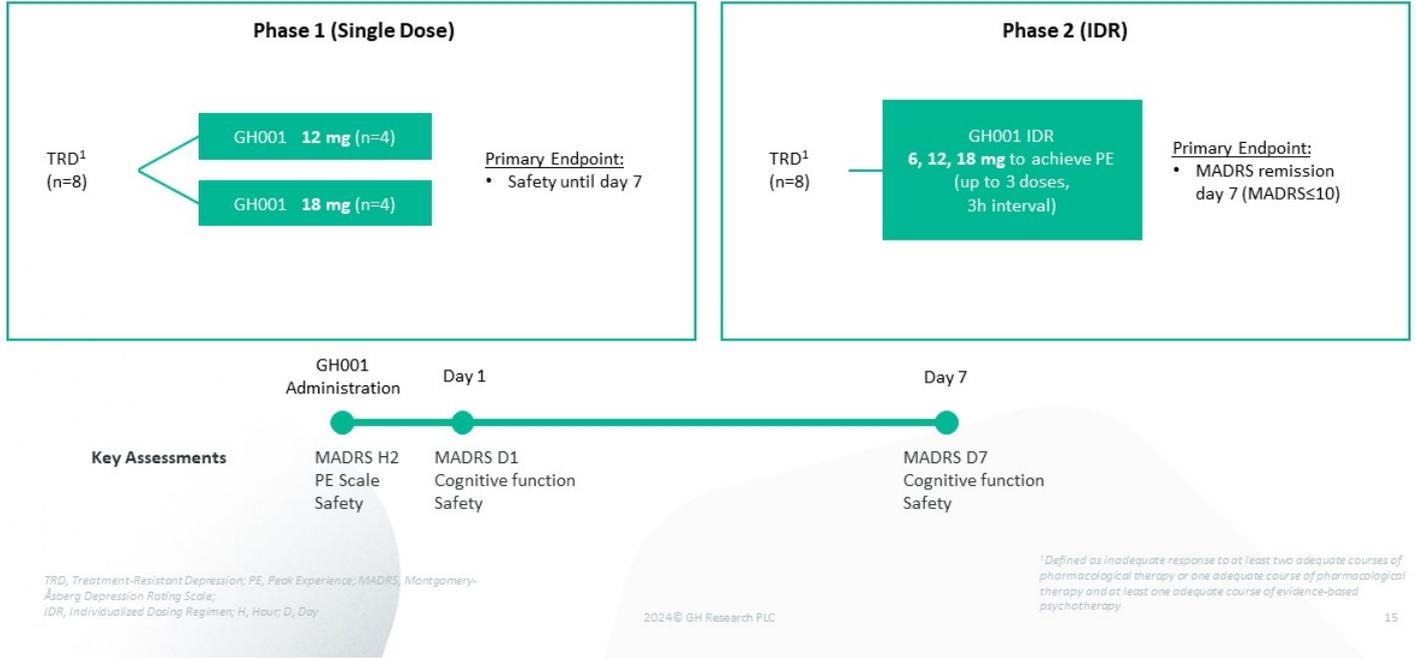
(Completed)

Clinicaltrials.gov ID: NCT04698603

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14

Design of Phase 1/2 Trial in TRD (GH001-TRD-102)



Phase 1 (Single Dose) and Phase 2 (IDR) – Safety



Study Safety Group review

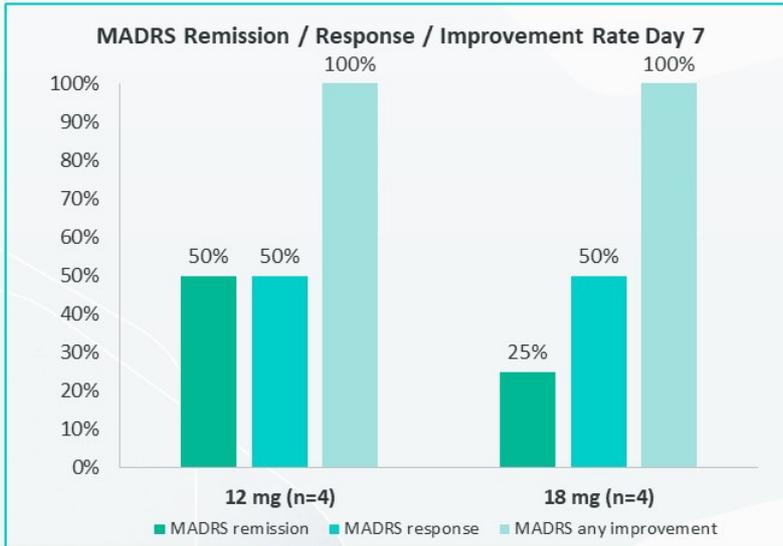
- No SAEs
- All ADRs mild, except three moderate (*)
- All ADRs resolved spontaneously
- Inhalation well-tolerated
- No noteworthy changes in vital parameters, except for temporary, non-clinically relevant increase in heart rate and blood pressure shortly after administration of GH001
- No clinically significant changes in safety laboratory analyses, psychiatric safety assessments or measures of cognitive function
- No safety signal relating to suicidal ideation or suicidal behavior, based on C-SSRS and MADRS subscore item “suicidal thoughts”

ADRs	Phase 1 (Single Dose)		Phase 2 (IDR)
	12 mg (n=4)	18 mg (n=4)	IDR ¹ (n=8)
MedDRA Preferred Term	Number of Events		
Abdominal discomfort			1
Anxiety			2
Depressive symptom			1*
Dizziness	1		
Feeling abnormal	1	1	
Flashback	1	1	2
Headache	2	1	3
Muscle discomfort			1
Muscle spasms		1	
Nausea			2*
Paresthesia			1
Sensory disturbance			3

SAE, Serious Adverse Event; ADR, Adverse Drug Reaction, an adverse event with a relationship code to the investigational product of definite, probable, or possible, or where code is missing; IDR, Individualized Dosing Regimen; C-SSRS, Columbia-Suicide Severity Rating Scale; MADRS, Montgomery-Åsberg Depression Rating Scale

¹6-12 mg (n=6); 6-12-18 mg (n=2)

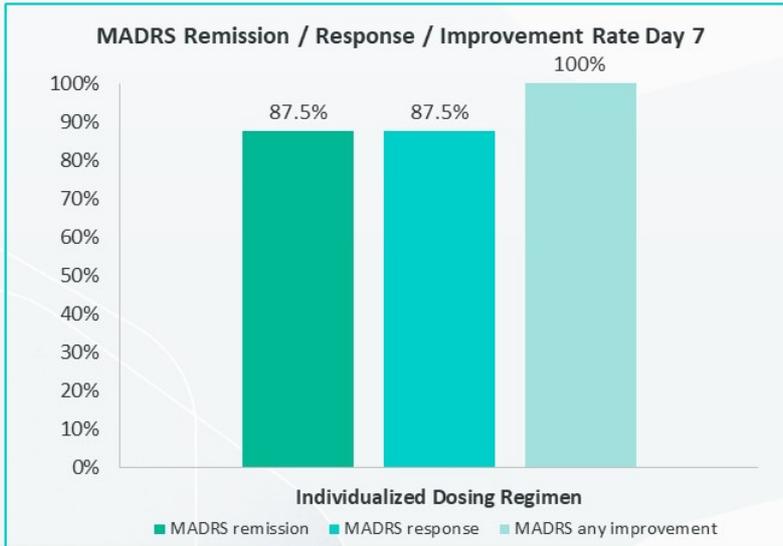
Phase 1 (Single Dose) – Efficacy (MADRS)



- 2 of 4 (50%) in the 12 mg group and 1 of 4 (25%) in the 18 mg group had a MADRS remission at day 7
- 2 of 8 patients had a PE and both had a MADRS remission at day 7

PE, Peak Experience; MADRS, Montgomery-Åsberg Depression Rating Scale
MADRS remission = MADRS of ≤10; MADRS response = Reduction of ≥50% from baseline in MADRS; MADRS any improvement = any reduction from baseline in MADRS

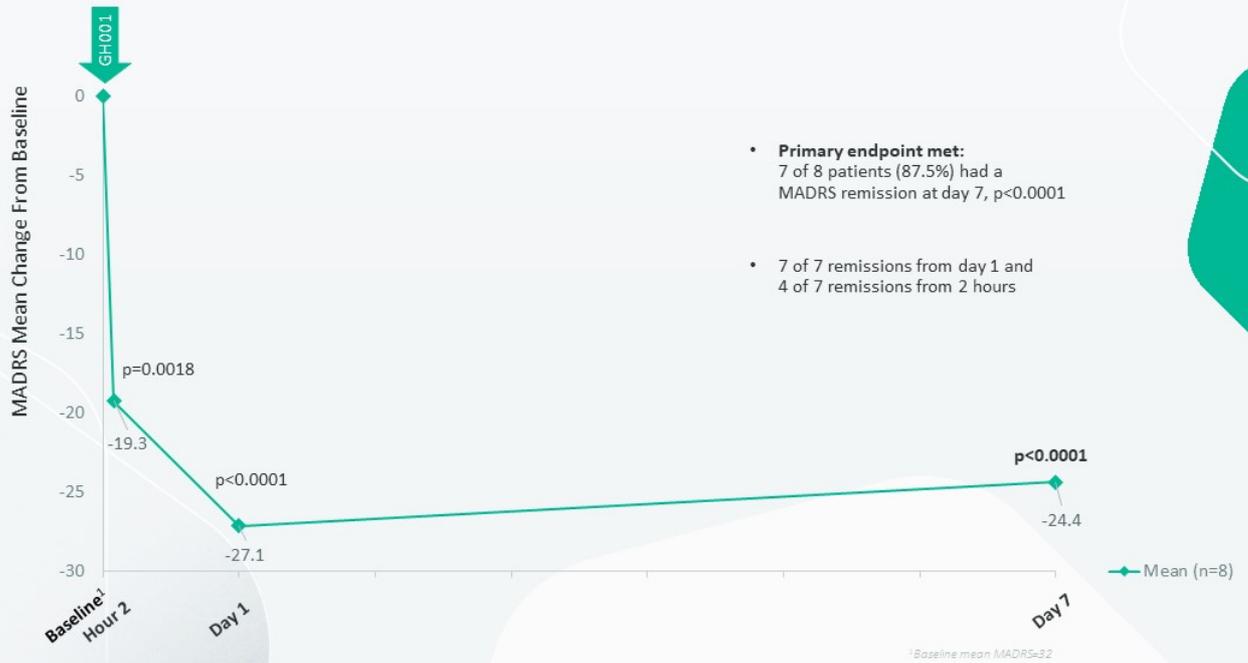
Phase 2 (IDR) – Efficacy (MADRS)



- **Primary endpoint met:**
7 of 8 patients (87.5%) had a MADRS remission at day 7, $p < 0.0001$
- 7 of 8 patients had a PE and 6 of those had a MADRS remission at day 7

PE, Peak Experience; MADRS, Montgomery-Åsberg Depression Rating Scale
MADRS remission = MADRS of ≤ 10 ; MADRS response = Reduction of 250% from baseline in MADRS; MADRS any improvement = any reduction from baseline in MADRS

Phase 2 (IDR) – Efficacy (MADRS Change from Baseline)



- **Primary endpoint met:**
7 of 8 patients (87.5%) had a MADRS remission at day 7, $p < 0.0001$
- 7 of 7 remissions from day 1 and 4 of 7 remissions from 2 hours

MADRS and PE – Observed Improved Outcome in Phase 2 (IDR) vs Phase 1 (Single Dose)



	Phase 2 (IDR)	Phase 1 (Single Dose) 12 mg	Phase 1 (Single Dose) 18 mg
MADRS Remission Rate Day 7	87.5% (7 of 8)	50% (2 of 4)	25% (1 of 4)
Mean MADRS Change Day 7	-24.4 (-76%)	-21.0 (-65%)	-12.5 (-40%)
Rate of PE	87.5% (7 of 8)	50% (2 of 4)	0% (0 of 4)
Mean PE Score	90.4 (at final dose)	58.2	59.1

PE, Peak Experience; MADRS, Montgomery-Åsberg Depression Rating Scale;
IDR, Individualized Dosing Regimen

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20



Phase 1 Clinical Pharmacology Trial in Healthy Volunteers GH001-HV-103

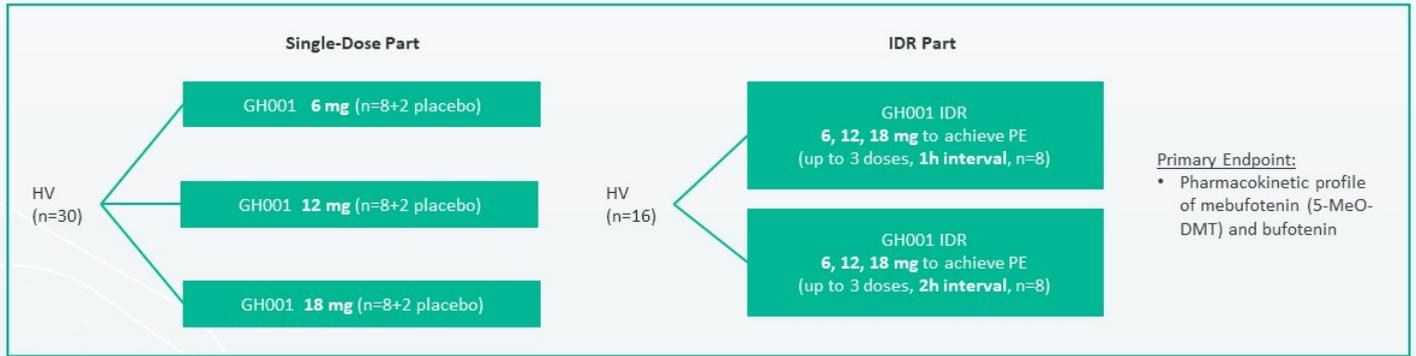
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Clinicaltrials.gov ID: NCT05163891

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21

Design of Phase 1 Clinical Pharmacology Trial in Healthy Volunteers (GH001-HV-103)



Primary Endpoint:
• Pharmacokinetic profile of mebufotenin (5-MeO-DMT) and bufotenin



HV, Healthy Volunteer; PE, Peak Experience; IDR, Individualized Dosing Regimen

Single Dose and IDR – Safety and Further Results



Safety Review

- No SAEs
- All ADRs mild
- All ADRs resolved spontaneously
- Inhalation well-tolerated
- No noteworthy changes in vital parameters, except for temporary, non-clinically relevant increase in heart rate and blood pressure shortly after administration of GH001
- No clinically relevant changes in ECG, safety laboratory analyses, peak flow, cognitive function, psychiatric safety assessments, including the C-SSRS

Further Results

- Pharmacokinetic analyses and psychoactive effect assessments (PE Scale) support that an interval down to 1 hour between individual doses of the IDR is feasible for future clinical trials

ADRs	Single-dose				IDR	
	6 mg (n=8)	12 mg (n=8)	18 mg (n=8)	Placebo (n=6)	1h interval (n=8) ¹	2h interval (n=8) ²
MedDRA Preferred Term	Number of Events					
Abnormal dreams						1
Chest discomfort		1				
Crying			2		2	
Dizziness			1			
Dry mouth	1					
Dyskinesia			1			
Fatigue		1			2	1
Headache	3		1		1	1
Hypoesthesia oral		1				
Paresthesia oral						1
Retching			1			
Somnolence		1				
Tachycardia			2			
Tension						1
Tremor			1			

SAE, Serious Adverse Event; ADR, Adverse Drug Reaction, or ADR, an adverse event with a relationship code to the investigational product of definite, probable, or possible, or where code is missing; IDR, Individualized Dosing Regimen; C-SSRS, Columbia-Suicide Severity Rating Scale; PE, Peak Experience

¹6 mg (n=1), 6-12 mg (n=3); 6-12-18 mg (n=4)
²6-12 mg (n=3); 6-12-18 mg (n=5)



Phase 2b Trial in Treatment-Resistant Depression GH001-TRD-201

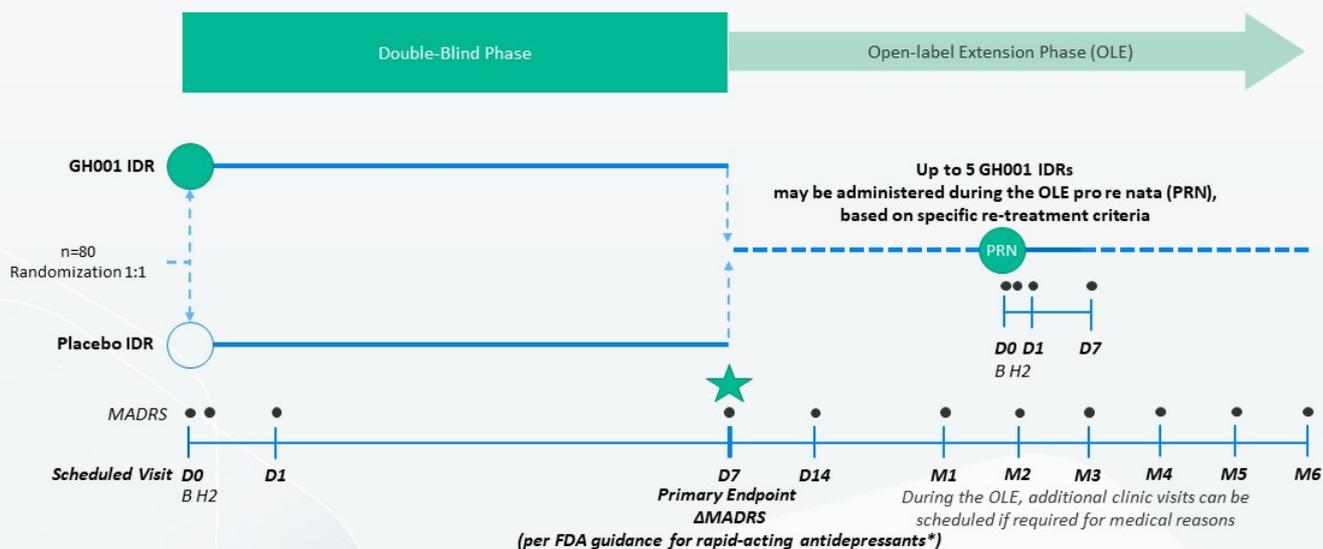
(Initiated)

EudraCT Number: 2022-000574-26

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24

Design of Phase 2b Trial in TRD (GH001-TRD-201)



The bold solid lines indicate the fixed duration of 7 days (± 1 day) after an IDR with visits on D0, D1 and D7. The bold dotted line indicates the variable duration until a potential GH001 IDR in the OLE. The GH001 IDR consists of up to 3 increasing doses (6, 12, 18 mg) and the Placebo IDR consists of up to three placebo doses, to achieve a peak experience, given at a 1H interval. As in previously completed trials, the GH001-TRD-201 trial will be conducted under the supervision of a healthcare provider, but without any planned psychotherapeutic interventions before, during, or after dosing. IDR, Individualized Dosing Regimen; PRN, pro re nata (as needed); B, Baseline; H, Hour; D, Day; M, Month. *FDA draft guidance for industry "Major Depressive Disorder: Developing Drugs for Treatment"

Three-Layer Protection Strategy



LAYER 1: REGULATORY EXCLUSIVITY

FDA: 5 years (+2.5 years paragraph IV stay)
EMA: 10 years (+1 year for new indication)

LAYER 2: PATENTS

Patent families filed relating to mebufotenin (5-MeO-DMT), including:

- Novel uses in various disorders (including inhaled, nasal, buccal, sublingual, i.v., i.m., s.c. routes)
- Novel aerosol compositions of matter
- Novel manufacturing methods and novel salt forms
- Novel device-related aspects

LAYER 3: TECHNICAL

Complex bioequivalence for systemically-acting inhalation/intranasal products with high intra- and inter-subject variability

Board of Directors & Executive Management



Florian Schönharting
MSc
Chairman of the Board, Co-founder



Michael Forer
BA, LLB
Vice-Chairman of the Board



Dermot Hanley
BSc, MBA
Board Member



Duncan Moore
MPhil, PhD
Board Member



Theis Terwey
PD Dr. med.
CEO, Co-founder



Velichka (Villy) Valcheva
MD, MSc
VP, Clinical Research and Medical Affairs



Julie Ryan
ACA, MAcc, BComm
VP, Finance



Aaron Cameron
MSc, MBA
Chief Operating Officer



Magnus Halle
BSc
Managing Director, Ireland, Co-founder



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University of Pennsylvania




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Charité, Berlin




Johannes Ramaekers
Prof. Dr.
Professor, Faculty of Psychology
and Neuroscience of Maastricht University


Anticipated Milestones and Financial Overview



GH001

- Complete double-blind phase of European multi-center, randomized, double-blind, placebo-controlled Phase 2b trial in TRD in Q3 2024, and provide top-line data in Q3 or Q4 2024
- Provide update on U.S. IND clinical hold and planned Phase 1 clinical pharmacology trial with proprietary aerosol delivery device after taking into account the conclusions of expected meeting with FDA in Q1 2024
- Provide update on timeline for completion of Phase 2a trials in PPD and in BDII in Q1 2024

GH002

- Provide top-line data from completed Phase 1 clinical pharmacology trial in healthy volunteers in Q1 2024

GH003

- Complete preclinical development

Financial Overview

- Cash, cash equivalents, other financial assets and marketable securities were \$228.7 million as of September 30, 2023
- We believe existing cash, cash equivalents, other financial assets and marketable securities will be sufficient to fund operating expenses and capital expenditure requirements into 2026



Seeking Ultra-Rapid, Durable Remissions in Depression