() BIOTECH

Break Boundaries. Ignite Change.

Nasdaq: IOBT Corporate Presentation May 2025



DISCLAIMER | Forward Looking Statements

Certain information contained in this presentation includes "forward-looking statements" within the meaning of Section 27A of the Securities Act of 1933, as amended, and Section 21E of the Securities Exchange Act of 1934, as amended, related to our business plan, clinical trials and regulatory submissions. We may, in some cases, use terms such as "may," "should," "would," "expects," "plans," "anticipates," "could," "intends," "target," "projects," "contemplates," "believes," "estimates," "predicts," "potential" or "continue" or other words that convey uncertainty of the future events or outcomes to identify these forward-looking statements. Our forwardlooking statements are based on current beliefs and expectations of our management team that involve risks, potential changes in circumstances, assumptions, and uncertainties. Any or all of the forward-looking statements may turn out to be wrong or be affected by inaccurate assumptions we might make or by known or unknown risks and uncertainties. These forward-looking statements are subject to risks and uncertainties including risks related to the execution of our business plan, success and timing of our clinical trials or other studies and the other risks set forth in our filings with the U.S. Securities and Exchange Commission. For all these reasons, actual results and developments could be materially different from those expressed in or implied by our forward-looking statements. You are cautioned not to place undue reliance on these forward-looking statements, which are made only as of the date of this presentation. We undertake no obligation to publicly update such forward-looking statements to reflect subsequent events or circumstances, except as required by law.

BREAK BOUNDARIES. IGNITE CHANGE.

Cylemblo[®] (imsapepimut and etimupepimut, adjuvanted)

THE PATIENT JOURNEY

Disease Presentation

Primary Care Physician or

Cylembio will, if approved, be available off-theshelf to all eligible 1L unresectable or metastatic melanoma patients

Dermatologist Unresectable or Metastatic Melanoma **Diagnosis & Referral** ~15,000 patients²: High unmet medical need still exists due to lack of efficacy and toxicity issues from Newly Diagnosed Melanoma available SOC in ~100,000¹ patients annually in the U.S. 1L Treatment Selection, Insurance and Access (anti-PD1 monotherapy; anti PD1 combination with either anti-CTLA4 or anti LAG3 agents; BRAF inhibitors) **Resectable Melanoma** 3 ~92,000 patients² Patient Checks in at the Infusion Center Surgical candidate? No Yes Product Preparation & Delivery to Infusion Neo/adjuvant treatment and Center **Surgical Resection Product Administration Residual Disease Post-Surgery?** Yes

1. Paulson. JAMA Dermatol. 2019; Rueth. Surg Oncol Clin N Am. 2016; Whiteman. J Invest Derm. 2016; 2012-2022 SEER Melanoma of the Skin Data. 2. Addressable patients in the US-Data on file

(imsapepimut and etimupepimut, adjuvanted)

Expected to be available

Off-the-shelf product

Rapid subcutaneous

administration

across all sites of care

candidate

HIGHLIGHTS | BREAK BOUNDARIES. IGNITE CHANGE.

80%

50%

25.5

STRONG PLATFORM AND PIPELINE

T-win[®] platform delivering investigational, immunemodulatory, off-the-shelf therapeutic cancer vaccines

Pipeline programs

- Cylembio
- IO112
- IO170

Indications (Melanoma, SCCHN, NSCLC), and potentially other difficult to treat cancers

CLINICAL PROOF-OF-CONCEPT ESTABLISHED

ORR*

CRR*

Months

mPFS*

Ph1/2 Clinical proof-ofconcept established

> Breakthrough Therapy Designation for advanced melanoma**

> > Improving clinical effect without adding systemic toxicity

PRODUCT POTENTIALLY ON THE MARKET IN 2026

Ph3 Pivotal Trial

in advanced melanoma, readout of PFS as primary endpoint expected **3Q2025**

2025 Potential BLA Submission to FDA

(imsapepimut and etimupepimut, adjuvanted)

US Brand Name of IO102-IO103

* Results from Phase 1/2 MM1636 Melanoma trial; ** Based on positive Phase 1/2 1L metastatic melanoma data, IO102-IO103, in combination with Merck's anti-PD-1 therapy, KEYTRUDA[®] pembrolizumab), was granted a Breakthrough Therapy Designation for the treatment of advanced melanoma by the US FDA



UNMET NEED IN ADVANCED MELANOMA | IO Biotech aims to address the unmet need for 1L advanced melanoma patients

Melanoma is a growing and common cancer



~331,000

patients newly diagnosed annually (global)

~58,000

patient deaths² annually (global)

23%

5-year survival rate for patients in stage IV³

The global melanoma market is expected to reach >\$13B by 2030⁴



Patients and physicians seek more effective treatment options⁵⁻⁸

<u>...https://gco.iarc.fr/today/en/;</u> 2. https://seer.cancer.gov/statfacts/html/melan.html; 3. Melanoma Research Alliance; 4. Evaluate Pharma 2025;

5. Larkin et al. N Engl J Med 2019;381:1535-1546; 6. Robert et al. Lancet Oncol 2019;20:1239-1251; 7. Tawbi et al. N Engl J Med 2022;386:24-34; 8. Weber et al. Oncologist. 2016;21(10):1230-1240

Phase 1/2 MM1636 Melanoma

MELANOMA | Successful outcomes from Phase 1/2 were published in Nature Medicine and drove continued clinical development for 1L melanoma

Compelling Data Published in Nature Medicine December 09, 2021



Results showing an attractive safety profile

January 2023 Data Cut* as Published in JITC, May 2023



* January 23 update: One patient was re-evaluated and did not have "real progression" but instead pseudo progression Lorentzen et al. J Immunother Cancer 2023;11(5):e006755

** TRAEs were not published in JITC data

 CR, complete response; LDH, lactate dehydrogenase; PD, progressive disease; PD-La, programmer Lorentzen et al. J Immunother Cancer 2020;34(5):eoof575

MELANOMA | Treatment for 1L melanoma is currently in Phase 3, fully enrolled with primary endpoint readout expected in 3Q25



MILESTONES

- Completed enrolment of 407 patients in December 2023
- August 2024: **IDMC recommended trial continue without modifications,** no new safety signals observed, determined investigational arm did not reach superiority on ORR at interim analysis
- Company's confidence in PFS unchanged

NEXT STEPS

- PFS readout expected 3Q25
- If supportive, BLA submission in the US expected in 2025

1L, first-line; AE, adverse event; BLA, Biologics License Application; CNS, central nervous system; CRR, complete response rate; DCR, disease control rate; DOR, duration of response; DRR, durable response rate; ECOG, Eastern Cooperative Oncology Group; IA, interim analysis; IDMC, independent data monitoring committee; mut, mutation; ORR, objective response rate; OS, overall survival; PD-1, programmed cell death protein 1; PFS, progression-free survival; ph, phase; q3w, once every 3 weeks; RECIST, Response Evaluation Criteria in Solid Tumors; SAE, serious adverse event; TTR, time to response; WT, wild-type. ClinicalTrials.gov: NCTo5155254.

MELANOMA | Physician feedback highlights the potential of IO Biotech's vaccine candidate Cylembio[®] (imsapepimut and etimupepimut, adjuvanted)





LAUNCH GOALS | Opportunity to establish Cylembio as a new standard of care for patients with advanced melanoma

1

3

2

PILLARS FOR A SUCCESSFUL LAUNCH

LAUNCH GOALS TO ENSURE RAPID ADOPTION AT LAUNCH



Establish belief in potentially transformative clinical benefit of Cylembio plus pembrolizumab combination

Maximize product value by ensuring all eligible patients have rapid access to therapy

Pragmatic launch model focusing on top melanoma treaters in the academic and community settings



PIPELINE | The T-win[®] platform with 3 product candidates in multiple cancer indications

Product candidates	Line of therapy/ indication	Pre-clinical	Phase 1	Phase 2	Phase 3	Takeaways
Cylembio® IO102-IO103 Targets: IDO1, PD-L1	IOB-013: First Line Advanced Melanoma*					Phase 3 pivotal trial primary endpoint, PFS, expected to read out in 3Q 2025
	 IOB-022: First Line Solid Tumors* Lung (NSCLC) Head & Neck (SCCHN) 					Indication expansion strategy in 1L NSCLC and SCCHN
	IOB-032: Neoadjuvant / Adjuvant Solid Tumors [*] • Melanoma • Head & Neck (SCCHN)					Extension into earlier lines of treatment
IO112 Target: Arginase 1	Solid Tumors Indications TBD 					Next pipeline candidate expected to enter clinical development
IO170 Target: TGF-β1	Solid Tumors Indications TBD 					Early-stage pipeline candidate targeting additional immuno- suppressive mechanisms

Cylembio® (imsapepimut and etimupepimut, adjuvanted)



* In combination with pembrolizumab; NSCLC, non-small cell lung cancer, PFS, progression-free survival; SCCHN, squamous cell carcinoma of the head and neck; IOB-013: ClinicalTrials.gov: NCT05155254; IOB-022: ClinicalTrials.gov: NCT05077709; IOB-032: ClinicalTrials.gov: NCT05280314

UNMET NEED IN ADVANCED NSCLC IO Biotech aims to improve patient outcomes in hard-to-treat cancers

NSCLC corresponds to 85% of lung cancer diagnosis¹⁻³

2nd

most diagnosed cancer worldwide (lung cancer)

~ 2,108,000

patients newly diagnosed annually (qlobal)

~1,544,000 patient deaths annually (global)

28%

5-year survival rate for patients in stage IV

B, billions; IOBT, IO Biotech; mPFS, median progression-free survival; mOS, median overall survival; NSCLC, non-small cell lung cancer

Global NSCLC market is expected to reach ~\$60B 2030³



An unmet need remains⁵⁻⁷ due to complex nature of the disease

of patients fail to **40-60%** respond to first-line treatment options

Current therapies are able to provide a mPFS and mOS of only ~8-10 months and ~2 years, respectively



With **chemotherapy** still major part of the SoC regimens, these treatments are associated with high toxicity

1. GLOBOCAN: https://gco.iarc.fr/today/en/; 2. https://seer.cancer.gov/statfacts/html/melan.html; 3. https://pmc.ncbi.nlm.nih.gov/articles/PMC3864624/; 4. Evaluate Pharma 2025; 5. Reck et al. N Engl J Med 2016;375(19)1823-33; 6. Reck et al. J Clin Oncol 2019;37(7):537-54; 7. Mok et al. Lancet 2019;393;10183:1819-1830

UNMET NEED IN ADVANCED SCCHN | IO Biotech aims to improve patient outcomes in hard-to-treat cancers

SCCHN is a life-threatening disease with poor prognosis ¹⁻²

6th

most diagnosed cancer in worldwide

~890,000

patients newly diagnosed annually (global)

~450,000

patient deaths annually (global)

39% 5-year survival rate for patients in stage IV Global SCCHN market is expected to reach ~\$5B by 2030³



An unmet need remains ⁴ for patients with advanced SCCHN



of patients fail to respond to the available treatment options

Current therapies are able to provide a **mPFS** and **mOS** of only ~5 months and ~12 months, respectively



With **chemotherapy** still major part of the SoC regimens, these treatments are associated with **high toxicity**

B, billions; IOBT, IO Biotech; mPFS, median progression-free survival; mOS, median overall survival; SCCHN, squamous cell carcinoma of the head and neck; SoC, standard of care

SCCHN & NSCLC | Encouraging data in head & neck and lung cancer from ongoing Phase 2 trial





NSCLC | Promising data presented at SITC 2024

Encouraging data for the NSCLC cohort

Best overall response, n (%) RECIST 1.1	N = 31*
ORR (95% CI)*	48 [30-67] confirmed/ 55 [36-73] unconfirmed
Complete Response (CR)	1 (3.2)
Partial Response (PR)	14 (45.2)
Stable Disease (SD)	10 (32.3)
Progressive Disease (PD)	6 (19.4)
Other endpoints	N = 31*
12 months PFS rate, %	48%
Disease control rate (PR + SD), % (95% CI)	81% (63-93)
mPFS, months (95% CI)	8.1

PFS and Confirmed Objective Response



*Efficacy-evaluable patients who received ≥2 cycles of treatment

Of 31 patients, 17 (55%) had a PR (n = 16) or CR (n = 1). Two patients (091 and 035) did not have the PR confirmed. Six patients were not included in the efficacy data set and are therefore not listed on this plot. Of the six patients, one was found ineligible due to squamous histology and five discontinued before completing the second cycle. The reasons for discontinuation were death [n = 2], maculo-papular rash [n = 1], pneumonitis/pulmonary embolism [n = 1], and early progression [n = 1].

Phase 2 IOB-022/KN-D38 Solid Tumor Basket

SCCHN | Primary endpoint met

Encouraging preliminary data for the SCCHN cohort

Best overall response, n (%) RECIST 1.1	N = 18*
ORR (95% CI)*	44.4 [21.5; 69.2]
Partial Response (PR)	8 (44.4)
Stable Disease (SD)	4 (22.2)
Progressive Disease (PD)	6 (33.3)
Other endpoints	N = 18
6 months PFS rate, %	60.6
Disease control rate (PR + SD), n (%)	12 (66.7)
Disease control rate (PR + SD), n (%) mPFS, months (95% CI)	12 (66.7) 6.6 [2.04; 13.14]

Treatment duration, time to response and best overall response per RECIST 1.1



Out of 18 patients, nine (50%) obtained a partial response (PR; light blue triangle). One patient (084) did not have the response confirmed due to death from intercurrent illness. Median duration of response was not yet reached at the time of data cut-off (02-Aug-2024). *HPV-positive/oropharyngeal tumours; *HPV-negative/oropharyngeal tumour.

Riess JW, et al. Presented at ESMO 2024. Poster 1022P.

TOTALITY OF CLINICAL DATA IS COMPELLING | IOBT is building on the body of evidence of Cylembio[®] across multiple tumor types and settings

Starting with **proof of concept** in advanced melanoma...

FIRST-LINE ADVANCED MELANOMA

Results from Phase 1/2 (MM1636): 80% ORR*, 50% CR, 25.5 months PFS

Status: Currently in Phase 3, enrollment complete with 407 patients

Primary Endpoint: PFS readout expected 3Q25

FIRST-LINE RECURRENT/ METASTATIC SCCHN

Results from Phase 2 (ESMO 2024) ORR 44.4%; mPFS 6.6 months Irrespective of HPV status

Benchmark** ORR 23%, mPFS 3.4 months

...expanding the body of evidence across three indications and settings

Status Primary endpoint ORR met with encouraging PFS

FIRST-LINE METASTATIC NSCLC

Results from Phase 2 (SITC 2024) ORR 55% unconfirmed/48% confirmed; mPFS 8.1 months

Benchmark** ORR 39%, mPFS 6.5 months

Status Encouraging ORR and PFS and no progression in nearly half of patients





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Phase 2 IOB-032/KN-D40 Neoadjuvant/adjuvant Basket Trial

CLINICAL TRIALS | Expanding beyond advanced diseases into the neoadjuvant/adjuvant treatment; fully enrolled Phase 2 basket study



CR, complete response; CRT, chemoradiotherapy; DFS, disease-free survival; ECOG, Eastern Cooperative Oncology Group; EFS, event-free survival; MPR, major pathologic response; NR, no response; ORR, objective response rate; p, pathological; PR, partial response; q3w, once every 3 weeks; RT, radiotherapy; SCCHN, squamous cell carcinoma of the head and neck; SoC, standard of care. ClinicalTrials.gov: NCT05280314.

ADDRESSABLE PATIENTS | Cylembio with potential to redefine treatment of hard-to-treat solid tumors and expanding beyond 1L advanced melanoma



1. 2025 estimated incidence of resectable Melanoma and SCCHN patients in the US. SEER; For melanoma refer to references in footnote #3; Ko C, Citrin D. Radiotherapy for the management of locally advanced squamous cell carcinoma of the head and neck. Oral Dis. 2009 Mar;15(2):121-32.) 2. 2023 estimated 1L mSCCHN CPS>20 patients in the US. SEER; For melanoma refer to references in footnote #3; Ko C, Citrin D. Radiotherapy for the management of locally advanced squamous cell carcinoma of the head and neck. Oral Dis. 2009 Mar;15(2):121-32.) 2. 2023 estimated 1L mSCCHN CPS>20 patients in the US. SEER; For melanoma refer to references, 2023; Seallo Cancer; 2023; Seallo Cancer; 2023; Seallo Cancer Teatment and Diagoois, 2021; Broak Med Sci. 2023; Gramer. Oral Oncol. 2019; Siegel. CA Cancer J Clin. 2023; Wusiman. Pathol Res Prac. 2022; estimated 1L addressable mNSCLCPD-L1250%: SEER; American Cancer Society; Mack Cancer; 2020; Evans Pathology and Oncology Research, 2018; Ganti. JAMA. 2021; Rodak. Cancers. 2023; Hotst. Lancet. 2019; Shah Cancer Treatment and Research Communications. 2023; Hotst. Lancet. 2019; Chancer Teatment and Research Communications. 2023; Hotst. Lancet. 2019; Chancer Cancer Society; Mack Cancer and C

3. 2025 estimated addressable 1L advanced melanoma patients in the US. SEER Melanoma of the Skin; Bajaj. J Natl Cancer Inst. 2020; Bensimon. J Med Econ. 2019; Leeneman. EJC. 2021; Luke. ASCO Presentation. 2023; Zhang. Adv Ther. 2023; Eggermont. J of Clin Onc. 2019; Egger NEJM. 2022

PRE-CLINICAL PIPELINE | Expanding the clinical portfolio with Arginase 1 program; IND filing expected in 2025





PLATFORM | T-Win[®] therapeutic cancer vaccines activate the immune system, targeting both tumor cells and immune-suppressive cells in the TME

The T-win[®] platform provides a **new therapeutic strategy** that may improve patient outcomes with a **novel mechanism of action** that **addresses multiple TME suppressive elements in solid tumors**.



Kjeldsen JW, et al. Nat Med 2021;27:2212–23. Erratum in: Nat Med 2022;28:871; Munir S, et al. PLoS One 2012;7:e34568; Munir S, et al. Oncoimmunology 2013;2:e23991; Ahmad SM, et al. Blood Cancer J 2014;4:e230; Andersen MH. Semin Immunopathol 2019;41:87–95; IO Biotech and Lankenau Institute (unpublished data); Andersen MH, et al. Semin Immunopathol 2019;41:87–95; IO Biotech and Lankenau Institute (unpublished data); Andersen MH, et al. Semin Immunopathol 2019;41:87–95; IO Biotech and Lankenau Institute (unpublished data); Andersen MH, et al. Semin Immunopathol 2023;45:253–64.

OFF-THE-SHELF TREATMENT | IOBT's immune-modulatory therapeutic cancer vaccine candidates designed to ensure all eligible patients can receive treatment without delay*

A 4 steps process from Cylembio® (imsapepimut and etimupepimut, adjuvanted) production to the patient vaccination...



... enhancing the overall patient experience.

Time to treatmentSite of careConvenienceIOBT's immune-modulatory therapeutic
cancer vaccine candidates provides fast
access to the medicine ensuring the
patients don't have to wait*The patients have access to therapy
where they get their cancer treatments in
both community and academic settingsRapid subcutaneous administration along
with SOC with no additional visits
required**



MILESTONES | Important clinical milestones expected through 2025

Cash into 2Q2026 including first three tranches of EIB loan facility and 1Q cash balance of \$37M*

	H1 2025	H2 2025	2026
Cylembio 1L Advanced Melanoma		 Readout of primary endpoint expected 3Q25 Potential BLA submission 	 Potential US approval Potential US launch
Cylembio Neoadjuvant / adjuvant Melanoma	 ✓ Complete enrollment in neoadj/adj Ph₂ cohorts 	Initial data	
Cylembio 1L Lung (NSCLC)		 Final data from Ph2 Decision on next steps 	
Cylembio Head & Neck (SCCHN)	 ✓ Complete enrollment in neoadj/adj Ph₂ cohort 	 Final data from first-line Ph2 Initial data from neoadjuvant/adjuvant Ph2 Decision on next steps 	
IO112 Solid Tumors (pre-clinical)	 IND filing Initiate enrollment in Ph1 study 		
IO170 Solid Tumors (pre-clinical)	IND enabling studies		

* On May 6, 2025, the Company drew on the EIB tranche A loan facility and obtained funding in the principal amount of *€*10.0 million before payment of certain fees and transaction related expenses.

BLA, biologics license application; IA, interim analysis; IDO, indoleamine 2,3-dioxygenase; IND, investigational new drug; IST, investigator-sponsored trial; NSCLC, nonsmall cell lung cancer; PD-L1, programmed cell death ligand 1; SCCHN, squamous cell carcinoma of the head and neck; TGF-β1, transforming growth factor-beta 1.

DRIVING THE NEXT FRONTIER OF IMMUNOTHERAPY | Cylembio[®], a potential first-in-class, immune-modulatory, off-the-shelf therapeutic cancer vaccine candidate

POISED FOR SUCCESS WITH POTENTIAL CYLEMBIO® LAUNCH Significant Off-the-shelf Cylembio® Large growing therapeutic potentially unmet medical transforming markets of cancer IO treatment need in melanoma, vaccine immunecandidate NSCLC, paradigm **SCCHN** oncology

BUILDING UPON SEVERAL SUCCESSFUL MILESTONES



Compelling Phase 1/2 data in metastatic melanoma patients



Promising Phase 2 results in NSCLC and SCCHN indications



Fully enrolled Phase 3 in advanced melanoma -PFS primary endpoint readout expected 3Q2025



IOBT preparing to submit BLA by year end 2025 and to potentially launch Cylembio in 1L advanced Melanoma in 2026

HIGHLIGHTS | BREAK BOUNDARIES. IGNITE CHANGE.

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THE TEAM | We have a strong management team with large biopharma and biotech experience





THE TEAM | Our management team is supported by the Board of Directors and the Scientific Advisory Board

Scientific Advisory Board

Board of Directors



IOBT HISTORY | Since its foundation in 2014, IO Biotech has built a strong platform and has the potential for US market launch in 2026

