


May 2025 Update

About Ichnos Glenmark Innovation (IGI)

IGI, a global fully integrated clinical-stage biotech company developing multispecifics™ in oncology, with the aim to accelerate new drug discovery in cancer treatment. IGI combines research and development proficiencies in novel biologics with those in new small molecules to continue developing cutting-edge therapy solutions that treat hematological malignancies and solid tumors. Harnessing the combined proficiency of over 100 scientists and a robust pipeline of novel molecules, IGI looks to leverage the capabilities of its centers of innovation spread across the USA, Switzerland and India to propel Innovation. For more information, visit www.iginnovate.com.

Headquartered in New York City, IGI has research and manufacturing operations at two sites in Switzerland. As a fully integrated biotechnology company with approximately 125 employees, IGI has strong capabilities in research, antibody engineering, small molecule, CMC, and clinical development of biotechnologies.

IGI is guided by an accomplished management team with experience developing immune cell engagers and small molecules within the biopharmaceuticals industry, and is led by Cyril Konto, M.D., President, Executive Director and Chief Executive Officer.

LEADERSHIP TEAM			PREVIOUS EXPERIENCE	BY THE NUMBERS
 Cyril Konto, M.D. President and Chief Executive Officer	 Lida Pacaud, M.D. Chief Medical Officer	 Mario Perro, Ph.D. Head of Biologics Research	 	100+ Years combined experience in biotech and pharmaceuticals
 Roberto Giovannini, Ph.D. Chief Process & Manufacturing Officer	 Dean Thomas, LL.M. General Counsel	 Sebastien Chenuet, Ph.D. Head of Business Development	 	30+ Products developed or launched
 Karishma Sipahimalani, Ph.D. Head of Human Resources			 	
			 	
			 	40+ Mergers, acquisitions, IPOs and other transactions

The proprietary BEAT® technology platform¹ is one of the bases for IGI's clinical-stage oncology pipeline. Using this technology, coupled with the proprietary common light chain library, the company is developing novel multispecific immune cell engagers and modulators, with the goal of realizing its mission to provide breakthrough, potentially curative therapies that may extend and improve lives, writing a new chapter in healthcare.

¹Bispecific Engagement by Antibodies based on the TCR

Oncology Pipeline

IGI's multispecific antibody and small molecule immune modulator pipeline, consists of three assets. This includes ISB 2001 (BCMAxCD38xCD3), which received orphan drug and fast track designations by the U.S. Food and Drug Administration (FDA) and is currently in dose-expansion Phase 1 clinical study for relapsed/refractory multiple myeloma (TRIgnite-1). GRC 65327 (Cbl-b inhibitor small molecule) is awaiting regulatory approval for initiating clinical development in India for solid tumors. ISB 2301 (NK-Cell Engager) is in the discovery stage for application in solid tumors. ISB 1442 (CD38 biparatopic x CD47) development has been discontinued, and the asset is available for licensing.

Updates of note in the last quarter are outlined below:

- + ISB 2001 completed enrollment of the phase 1 escalation (part-1) in March and initiated/dosed the first patient in the expansion (part-2) in April 2025
- + ISB 2001 EU CTA has passed validation on 27 March 2025
- + ISB 2001 FDA Fast Track designation has been granted on 23 April 2025
- + Three posters were presented at AACR in April 2025:
 - o ISB 2001: Clinical validation of a quantitative systems pharmacology (QSP) model of ISB 2001 used for deriving first in human (FIH) dose and efficient phase 1 dose escalation design in relapsed refractory multiple myeloma (RRMM) patients ([Link](#))
 - o ISB 2001: Pharmacokinetics (PK) and pharmacodynamics (PD) of ISB 2001, a novel BCMAxCD38xCD3 trispecific antibody from the First-in-Human (FIH) Phase 1 study in relapsed/refractory multiple myeloma patients ([Link](#))
 - o GRC 65327: Discovery of GRC 65327: A Best-in-Class, Selective and potent Cbl-b E3 ligase inhibitor for the treatment of advanced solid cancers ([Link](#))
- + ISB 2001 clinical data (encore) was presented at COMy in May 2025
- + ISB 2001 clinical abstract has been accepted at multiple conferences in H1 2025 and new clinical data will be presented in June 2025 at [ASCO2025](#) in the Rapid Oral Abstract Session, followed by Poster Presentation at EHA2025 (encore) also in June 2025

Oncology-Focused Pipeline to Drive Long-Term Value Growth

ASSET	DESCRIPTION	INDICATION	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	STATUS
CLINICAL ASSETS							
ISB 2001	BCMA x CD38 x CD3 TREAT™ trispecific T-Cell Engager	Multiple Myeloma					PHASE 1 ORPHAN DRUG
GRC 65327	Cbl-b Inhibitor Small Molecule	Solid Tumors					PRE-CLINICAL
CANDIDATES							
ISB 2301	IMMUNITE™ NK-Cell Engager	Solid Tumors					DISCOVERY

Partnering-Ready Asset to Accelerate Short-Term Value Creation

ASSET	DESCRIPTION	INDICATION	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	STATUS
CLINICAL ASSETS							
ISB 1442	CD38 biparatopic x CD47 BEAT® Myeloid-Cell Engager	Multiple Myeloma					PHASE 1 ORPHAN DRUG

IGI is looking for asset-level and platform-level collaboration partners in development and research. For more information, visit <https://IGInnovate.com/contact/>.

Overview of Oncology Candidates in Development

ISB 2001 TREAT™ TRISPECIFIC ANTIBODY

- ISB 2001 is a first-in-class T cell-engaging antibody that targets BCMA and CD38 on multiple myeloma cells. It is a trispecific antibody based on IGI's proprietary BEAT® platform, allowing maximal flexibility and excellent manufacturability of full-length multispecific antibodies.
- ISB 2001 combines three proprietary Fab antigen-binding arms, each targeting a different antigen, with one arm binding to the epsilon chain of CD3 on T cells, and the other two binding BCMA and CD38 on multiple myeloma cells. Its Fc domain was fully silenced to suppress Fc effector functions.
- ISB 2001 redirects CD3+ T lymphocytes to kill tumor cells expressing low to high levels of both BCMA and CD38. With two different tumor-associated antigens instead of one, ISB 2001 is expected to be more resistant to antigen escape associated with treatment of multiple myeloma patients.
- At the AACR Annual Meeting in 2024, an oral presentation showcased the results of ISB 2001 anti-myeloma activity ex-vivo in bone marrow aspirates from patients who have relapsed after CD38 and BCMA targeted therapies. ISB 2001 demonstrated superior cytotoxicity relative to teclistamab in the samples of patient relapsing from CD38 and BCMA targeted immunotherapies.
- The preclinical data package for ISB 2001 was in 2024 published in [Nature Cancer](#) and shows that:
 - + ISB 2001 can overcome resistance mechanisms by dual tumor targeting via binding and cytotoxicity of tumor cells with low expression of CD38 and/or BCMA.
 - + ISB 2001's architecture is optimized to support robust killing of tumor cells while limiting CD38 on-target, off-tumor activity.
 - + ISB 2001 demonstrated increased killing of tumor cells compared to BCMA-targeted T cell engagers in vitro, in vivo and ex vivo; induced complete tumor regression in humanized mouse models; and demonstrated superior potency compared to standard combination of therapies.
- The advantages of the trispecific ISB 2001 antibody was highlighted in the accompanying [News and Views article](#) written by S.R. Ruuls and P.W.H.I. Parren and was further emphasized in a [Fierce Biotech article](#) in which the mode of action of ISB 2001 and promise of IGI's BEAT® platform were described by IGI's CEO, Cyril Konto.
- In April 2023, IGI received approvals from HREC in Australia and the FDA to initiate a Phase 1 first-in-human study of ISB 2001 for the treatment of r/r MM. In April 2024, IGI received approval from DCGI to expand the clinical Phase 1 study into India. The phase 1 study is divided into a dose escalation part and a dose expansion part, with the latter being designed to meet the goals of FDA Project Optimus. First patient was dosed in November 2023 and the trial is now active in US, Australia and India, with dose expansion initiated in April 2025. Enrollment in Europe is targeted for end of Q2 2025.
- In July 2023, ISB 2001 received Orphan Drug Designation from the FDA for the treatment of MM and in April 2025, FDA also granted Fast Track designation to ISB 2001 ([press release](#)).
- IGI declared clinical Proof-of-Concept for ISB 2001 in r/r MM in July 2024, based on the data generated in the ongoing dose escalation phase, and decided to accelerate the development of this asset.
- The first clinical data of the ongoing ISB 2001 trial was presented in an oral presentation at [ASH 2024](#) on December 9th, 2024 ([press release](#)) and showed:

- + ISB 2001 is well tolerated with no dose limiting toxicities up to 1200 µg/kg, low grade cytokine release syndrome, no neurological Adverse Events or ICANs, low infection and hematological toxicity rates, no Adverse Events leading to discontinuation¹
- + Early, deep and sustained responses were observed across effective dose levels (DL3 to DL7) with antimyeloma activity from 50 µg/kg (MRD negative sCR) and higher¹
- + Overall Response rate (ORR) was 83% (22% Complete response (CR) or better, 50% Very Good Partial Response (VGPR) and 11% Partial Response (PR). The ORR was 75 % in patients pretreated with CAR-T or bispecific T cell engagers and 90 % in patients who had not been treated with T-cell directed therapies¹
- + Dose proportional PK with long half-life supports less frequent dosing and T cell activation observed at effective doses¹
- Next presentation of clinical data will be in June 2025 at [ASCO2025](#)
- ISB 2001 has been raising the interest from several global players, which has prompted the initiation of partnering discussions

¹Quach H. et al., ASH2024, Oral Presentation

CASITAS B-LINEAGE LYMPHOMA B (CBL/B) PROGRAM

- Casitas B-lineage lymphoma b (Cbl/b) is an E3 ubiquitin ligase that has been identified as a key inhibitor of T and NK cells activation in the absence of CD28 co-stimulation, regulate immune cells activity in PD-1, CTLA4, TIGIT etc positive cells. As an intracellular master regulator, Cbl/b inhibition may lead to robust immune cells activation in suppressed tumor microenvironment and induce strong single agent activity.
- The IND for the clinical candidate GRC 65327 was submitted to the Drugs Controller General of India (DCGI) on October 30, 2024. The meeting with the oncology subject matter expert committee (SEC) happened on December 13, 2024. The committee recommended the approval of the Phase 1 protocol with the condition of initiating the study with a 10 mg dose cohort and submitting data of the first subject of the same cohort before initiation into the second subject to the Central Drugs Standard Control Organization (CDSCO) for further deliberation by the committee. A second set of queries from DCGI SEC received on March 21, 2025, were addressed on April 23, 2025. Final reports of toxicology studies were submitted to DCGI along with the response. A formal approval of NOC is awaited.
- A poster entitled 'Discovery of GRC 65327: A Best-in-Class, Selective and potent Cbl-b E3 ligase inhibitor for the treatment of advanced solid cancers' was presented at [AACR2025](#).

Autoimmune Diseases

IGI has two monoclonal antibody drug product candidates addressing autoimmune diseases in the pipeline. To enhance the company's focus on oncology, future development of both assets is overseen by out-licensing partners.

The first asset, ISB 880, an anti-IL-1RAP antagonist, was licensed to Almirall, S.A. in December 2021. The initiation of dosing in a Phase 1 study of ISB 880/ALM27134 was announced by Almirall in September 2022.

The second antibody, ISB 830 (telazorlimab) and its follow-on molecule ISB 830-X8, was licensed to Astria Therapeutics in October 2023. Telazorlimab is an OX40 antagonist that successfully completed a Phase 2b study in moderate to severe atopic dermatitis in 2021. Both compounds have potential across a range of autoimmune diseases.

Assets in Autoimmune Diseases

MOLECULE MECHANISM/CLASS	POTENTIAL INDICATIONS	PHASE	STATUS
ISB 880 (ALM 27134) IL-1RAP Antagonist Monoclonal Antibody	Autoimmune Diseases	Phase 1	Licensed to Almirall S.A. in December 2021. <u>Dosing of participants in the Phase 1 study was announced by Almirall in September 2022.</u>
ISB 830 Telazorlimab OX40 Antagonist Antibody	Atopic Dermatitis	Phase 2b	Licensed to Astria Therapeutics in October 2023. Successfully completed a Phase 2b study in Atopic Dermatitis.
	Other autoimmune diseases, including Rheumatoid Arthritis	U.S. IND for Rheumatoid Arthritis and other autoimmune indications is active.	
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ISB 880 / ALM27134 (IL-1RAP ANTAGONIST)



IGI entered an exclusive global licensing agreement for ISB 880 in autoimmune diseases with Almirall in December 2021. Within the terms of the agreement, Almirall assumed full cost and responsibility for the global development and commercialization of the compound. IGI received an upfront payment of €20.8 million. The deal includes development and commercial milestone payments, and tiered royalties based upon future global sales. Almirall initiated a Phase I study in 2022, to evaluate the safety, pharmacokinetics, pharmacodynamics and clinical activity of the licensed asset. IGI received a milestone payment in March 2025 based on the successful phase I study.

For more information on this asset, please visit almirall.com

ISB 830 (TELAZORLIMAB, OX40 ANTAGONIST)



IGI entered an exclusive global licensing agreement for ISB 830 and its follow-on ISB 830-X8 with Astria Therapeutics in October 2023.

On January 23, Astria announced initiation of a phase 1a trial of STAR0310, a potential best-in-class monoclonal antibody OX40 antagonist for the treatment of atopic dermatitis. The dosing of the first human subject triggered the payment of a development milestone to IGI.

For more information, visit <https://IGInnovate.com/contact/>