



**RYVU THERAPEUTICS S.A.**  
**Q1 2023 Report**

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## 1. ECONOMIC AND FINANCIAL HIGHLIGHTS

### 1.1 Financial Results Obtained in the Reporting Period

Condensed Interim Financial Statements of Ryvu Therapeutics S.A. (“Company”, “Issuer”, “RyvU”) for the period from January 1, 2023 to March 31, 2023 are prepared in accordance with the requirements of the International Accounting Standard No. 34 “Interim Financial Reporting” endorsed by the EU (“IAS 34”).

Selected balance sheet data are as follows:

RyvU Therapeutics S.A. Item	Data in PLN thousand		Data in EUR thousand	
	31.03.2023	31.12.2022	31.03.2023	31.12.2022
Total assets	<b>449,183</b>	474,977	<b>96,072</b>	101,277
Short-term receivables	<b>25,556</b>	16,931	<b>5,466</b>	3,610
Cash from the issue on the account of the brokerage house	-	242,962	-	51,805
Cash and cash equivalents	<b>300,839</b>	101,917	<b>64,344</b>	21,731
Other current and non-current financial assets	<b>13,417</b>	604	<b>2,870</b>	129
Total liabilities	<b>120,399</b>	131,586	<b>25,751</b>	28,057
Long-term liabilities	<b>83,664</b>	86,772	<b>17,894</b>	18,502
Short-term liabilities	<b>36,734</b>	44,814	<b>7,857</b>	9,555
Total equity	<b>328,784</b>	343,390	<b>70,321</b>	73,219
Share capital	<b>9,248</b>	7,342	<b>1,978</b>	1,565

**Selected income statement data are as follows:**

Ryvü Therapeutics S.A.	Data in PLN thousand		Data in EUR thousand	
	From 01.01.2023 to 31.03.2023	From 01.01.2022 to 31.03.2022	From 01.01.2023 to 31.03.2023	From 01.01.2022 to 31.03.2022
Revenues from sales	5,012	32	1,066	7
Revenues from subsidiaries	5,455	6,754	1,161	1,453
Revenues from R&D projects	7,849	0	1,670	0
Other operating revenues	238	196	51	42
Revenues from operating activities	18,554	6,982	3,947	1,502
Operating expenses	-36,657	-33,738	-7,799	-7,260
Operating expenses without Incentive Scheme and valuation of Nodthera shares	-32,883	-25,587	-6,996	-5,506
Depreciation	-2,782	-3,375	-592	-726
Valuation of Incentive Scheme	-2,991	-8,149	-636	-1,754
Profit/loss from operating activities (EBIT)	-18,103	-26,756	-3,851	-5,757
Profit/loss from operating activities (EBIT) without Incentive Scheme and valuation of Nodthera shares	-14,329	-18,605	-3,048	-4,003
Profit/loss before income tax	-17,597	-26,583	-3,744	-5,720
Net profit/loss	-17,597	-26,536	-3,744	-5,710
Net profit/loss without Incentive Scheme	-14,606	-18,387	-3,107	-3,957
EBITDA	-15,321	-23,381	-3,259	-5,031
EBITDA without Incentive Scheme and valuation of Nodthera shares	-11,547	-15,230	-2,457	-3,277
Net cash flows from operating activities	-33,268	-21,807	-7,078	-4,693
Net cash flows from investing activities	-11,358	4,143	-2,416	892
Net cash flows from financing activities	241,549	-768	51,388	-165
Total net cash flow	196,923	-18,432	41,894	-3,966
Number of shares (weighted average)	22,220,154	18,355,474	22,220,154	18,355,474
Profit (loss) per share (in PLN)	-0.79	-1.45	-0.17	-0.31
Diluted profit (loss) per share (in PLN)	-0.79	-1.45	-0.17	-0.31
Book value per share (in PLN)	14.80	7.79	3.16	1.66
Diluted book value per share (in PLN)	14.80	7.79	3.16	1.66
Declared or paid dividend per share (in PLN)	-	-	-	-

**Selected financial data presented in the quarterly report were converted to Euro as follows:**

- Items relating to the profit and loss statement and the cash flow statement were converted using the exchange rate constituting the arithmetic average of the exchange rates, applicable as of the last day of every month in the given period, based on the information published by the National Bank of Poland (NBP):
  - for the period from 01/01/2023 – 31/03/2023: PLN 4.7005;
  - for the period from 01/01/2022 – 31/03/2022: PLN 4.6472;
- Balance sheet items were converted using the average exchange rate announced by the NBP applicable as of the balance sheet date; which were:
  - as of 31 March 2023: PLN 4.6755;
  - as of 31 December 2022: PLN 4.6899.

## 1.2 Management Board comments to the financial results

In the first quarter of 2023, Ryvu Therapeutics S.A. recognized total operating revenue of PLN 18,554 thousand, which constitutes an increase compared to the corresponding period in 2022, when total operating revenue amounted to PLN 6,982 thousand. This results from an increase in revenues from R&D projects (an increase of PLN 7,849 thousand) and increase in revenues from sales (an increase of PLN 4,980 thousand) partially offset by a decrease in revenues from subsidies (a decrease of PLN 1,299 thousand) compared to the corresponding period in 2022.

Revenues from R&D projects in the first quarter of 2023 resulted from the following transactions:

- achievement of a milestone in the amount of USD 1 million from the exclusive license agreement with Exelixis Inc. The agreement combines Ryvu's proprietary small molecule STING agonists and STING biology know-how with Exelixis' network of expertise and resources in antibody engineering, antibody-drug conjugate (ADC) technologies, and oncology therapeutics development and commercialization experience.
- recognition of a portion of the upfront payment in the amount of PLN 3,514 thousand from the exclusive research collaboration and license agreement with BioNTech SE. In accordance with the accounting policy of Ryvu and IFRS 15, in 2022 Ryvu recognized only a part of the upfront revenues. The remaining amount is recognized equally in each period for 5 years.

Revenues from sales resulted mostly from research collaboration with BioNTech SE. Under the License Agreement, Ryvu provides appropriately qualified employees and BioNTech funds all discovery, research and development activities under the multi-target research collaboration.

In the first quarter of 2023, Ryvu reported a net loss, as well as an operating loss. The net and operating losses are the result of the fact that the Company focuses on increasing the value of the ongoing projects that will be commercialized at a later stage of development.

The Company's net loss for the period ended March 31, 2023, amounted to PLN 17,597 thousand compared to the net loss of PLN 26,536 thousand in the corresponding period of 2022. The smaller loss in 2023 is related to the abovementioned transactions and lower non-cash cost of valuation of the incentive program for its employees of PLN 2,991 thousand (described below), partially compensated by a higher negative change in NodThera shares valuation of PLN 783 thousand (described below), as well as higher expenditure incurred on research and clinical projects.

### Valuation of shares in NodThera Inc.

#### Valuation of shares

As of March 31, 2023, four types of shares existed in NodThera Inc.: ordinary stock and preferred stock (Junior Preferred Stock, Series A1 and A2 Preferred Stock, Series B Preferred Stock and Series C Preferred Stock). Ryvu is a holder of the Junior Preferred Stock.

Associated with the Series A, B and C Preferred Stock is the right to receive dividends in the form of cash or the issuance of shares of the same class and the non-dilution right. The payment of dividends and execution of the anti-dilution right may be made only in cases specified in the investment agreement, in particular in the event of a sale of the company or the admission of its shares to trading on a stock exchange. The shares held by Ryvu, i.e. Junior Preferred Stock, do not have the aforementioned right to pay dividends or the non-dilution right.

Series C Preferred Stock was issued by NodThera Inc. on September 20, 2022. The issue comprised of 8,698,375 shares at a price of USD 2.8741 per share. As a result of this issue, NodThera received financing in the total amount of USD 25,000,002.47. The issue was addressed only to existing investors. Ryvu did not participate in the issue.

Thanks to the receipt of funds raised from the Series C share issue, according to information obtained from NodThera Inc., NodThera has the necessary financial resources to fully implement the projects currently underway. In addition, the proceeds will provide enough cash for the company to operate smoothly until the end of 2023 and to seek additional capital for development in 2024 and the following years.

The Management Board of Ryvu has decided to include in the valuation of the shares held by Ryvu in NodThera, a 16.42% discount (reflecting no right to dividend and non-dilution right) to the price at which they were subscribed under the last share capital increase, i.e. issuance of series C on September 20, 2022, and the above approach was applied as of March 31, 2023.

Therefore, a share valuation of GBP 2.4022/share (share price from the last financing round from September 20, 2022, including a discount corresponding to the class of shares held by the Issuer) should be used as a basis for the calculations. As of 31.03.2023, Ryvu held 3.15% shares in NodThera on a fully diluted basis and the total valuation of the Issuer's shares in NodThera Inc. amounts to PLN 19,692,100 (at the average NBP exchange rate of 4,2934 PLN/USD).

#### Valuation of shares in NodThera Inc. according to fair value:

new share issue price (in USD)	2.4022
average NBP exchange rate from March 31, 2023	4.2934
new share issue price (in PLN)	10.31
number of the Company's shares in NodThera Inc.	1,910,000
value of shares in the balance sheet as of March 31, 2023	19,692,100
value of shares in the balance sheet as of December 31, 2022	20,475,200
<b>change in valuation – gross impact on the valuation of shares</b>	<b>-783,100</b>

#### Incentive Scheme

On May 17, 2021, the General Shareholders Meeting adopted the non-dilutive Stock Grant Program for 2021-2024 for all employees in the form of the right to acquire shares of the Company. The Stock Grant Program is comprised of 1,247,720 ordinary shares of the Company that have been donated free of charge by Mr. Paweł Przewięźlikowski – founder, President of the Management Board, and Company's largest shareholder to the Company, constituting a total of 25% of the Company's shares held by Mr. Paweł Przewięźlikowski. The Stock Grant Program provides employees with the right to acquire shares at a preferential price of PLN 0.19 per share, covering the Company's administrative costs incurred to execute the Stock Grant Program. The fair value of the shares granted is determined as of the grant date and recognized over the vesting period in remuneration costs in correspondence with the capital increase at the time of vesting by employees during the program. For the period ending March 31, 2023, the Company recognized the non-cash cost of valuation of this

incentive program of PLN 2,991 thousand – more details are described in note 32 to the financial statements.

### Issue of Series ‘J’ Shares

In Q4 2022, the Company carried out a successful issue of Series “J” Shares, as a result of which the Company secured over PLN 242.5 million net. As of December 31, 2022, proceeds from the issue were presented in the item "Cash from the issue on the account of the brokerage house." Ryvu was eligible to receive the funds from the issue after the registration of the capital increase, which took place in January 2023.

### 1.3 The Company’s Assets and the Structure of Assets and Liabilities

As of March 31, 2023, the value of the Company’s assets was PLN 449,183 thousand and decreased by PLN 25,794 thousand compared to the end of 2022 (PLN 474,977 thousand), mainly due to the expenditures on R&D projects. At the end of March 2023, the highest value of assets was cash which amounted to PLN 300,839 thousand (at the end of 2022, it was PLN 101,917 thousand) and other financial assets of PLN 13,417 thousand (at the end of 2022, it was PLN 604 thousand). As of March 31, 2023, other financial assets were mostly bonds (PLN 12,870 thousand). The increase in cash resulted mainly from the transfer of funds from the brokerage house accounts to Ryvu’s accounts because of the successful issue of Series “J” Shares. Fixed assets were mainly the Research and Development Centre for Innovative Drugs (named ‘CBR’) and laboratory equipment, as well as the valuation of NodThera shares (PLN 19,692 thousand).

The main item in Ryvu’s equity and liabilities is equity, which amounted to PLN 328,784 thousand as of March 31, 2023, and decreased by PLN 14,606 thousand compared to December 31, 2022. The decrease in equity was mainly a result of the net loss recognized for the period. The other source of asset’ funding are long-term liabilities which amounted to PLN 83,664 thousand at the end of March 2023. The long-term liabilities are mainly related to the deferred income linked mainly to the deferred revenue from the BioNTech agreement and the infrastructure subsidy for CBR.

**The asset structure demonstrates the Company’s high financial liquidity, which is confirmed by the following ratios:**

	31.03.2023	31.12.2022
<b>Current ratio</b>		
<b>current assets/current liabilities, including short-term provisions and accruals (excl. deferred revenues)</b>	9.57	8.82
<b>Quick ratio</b>		
<b>(current assets-inventory)/current liabilities, including short-term provisions and accruals (excl. deferred revenues)</b>	9.52	8.77

Cash surpluses, not used in the operating activities, are deposited in the low-risk financial instruments like short term bank deposits and bonds.

#### **1.4 Current and Projected Financial Condition**

The Company's financial position as of the report date is very good considering the current cash position and the expected financing from the European Investment Bank. As of March 31, 2023, the value of the Company's cash amounted to PLN 313,709 thousand (PLN 300,839 thousand in cash at the banks and PLN 12,870 thousand in bonds), and as of May 11, 2023, it was PLN 297,474 thousand (PLN 284,604 thousand in cash at the banks and PLN 12,870 thousand in bonds). The decrease in cash has resulted from expenditure on R&D projects.

The Company meets its obligations in a timely manner and maintains sustainable cash levels ensuring its financial liquidity. Cash inflow from previous share issues, funds obtained from subsidies from the EU, funds supporting R&D projects and cash generated from the commercialization of projects allow the Company to execute its planned investments, in particular, the development of the ongoing and new innovative projects and expansion of laboratory infrastructure. The future Company's revenues will strongly depend on the ability to commercialize the research projects.

## 2 MANAGEMENT BOARD INFORMATION ON ACTIVITES

### 2.1 The pipeline

Ryvu Therapeutics is advancing a broad pipeline addressing emerging targets in oncology.

Ryvu's pipeline includes candidates with differentiated therapeutic mechanisms, including programs directed at kinase, synthetic lethality, immuno-oncology and immunometabolism pathways.

These research and development projects are represented below.

#### CLINICAL PROJECTS

PROGRAM / TARGET NAME	INDICATION	DISCOVERY	PRECLINICAL	PHASE I	PHASE II	PARTNER	NEXT ANTICIPATED MILESTONE
RVU120 CDK8/19	AML/MDS	[Progress bar: Discovery to Phase I]				LEUKEMIA & LYMPHOMA SOCIETY	Additional Phase I data in Q2 2023
	SOLID TUMORS	[Progress bar: Discovery to Phase I]					Additional Phase I data in Q2 2023
SEL24 (MEN1703) PIM/FLT3	AML	[Progress bar: Discovery to Phase I]				MENARINI	

#### DISCOVERY & PRECLINICAL PROJECTS

PROGRAM / TARGET NAME	INDICATION	DISCOVERY	PRECLINICAL	PHASE I	PHASE II	PARTNER	NEXT ANTICIPATED MILESTONE
<b>SYNTHETIC LETHALITY</b>							
PRMT5	SOLID TUMORS	[Progress bar: Discovery]					Development candidate in 2023
WRN	SOLID TUMORS	[Progress bar: Discovery]					In vivo POC in 2023
Novel Targets	ONCOLOGY	[Progress bar: Discovery]					
<b>IMMUNO-ONCOLOGY</b>							
STING Standalone	ONCOLOGY	[Progress bar: Discovery]				BIONTECH	
STING ADC	ONCOLOGY	[Progress bar: Discovery]				EXELIXIS	
HPK1	SOLID TUMORS	[Progress bar: Discovery]					
<b>IMMUNE MODULATION RESEARCH COLLABORATION (MULTI-TARGET) DISCOVERY COLLABORATION</b>							
		[Progress bar: Discovery]				BIONTECH MERCK	

Source: Company's own data.

### SEL24 (MEN1703)

SEL24 (also known as MEN1703) is a selective, small molecule, dual inhibitor of PIM and FLT3 kinases, two enzymes that are strongly implicated in malignant transformation of hematopoietic cells. The compound has been discovered by Ryvu and is currently in development in collaboration with Menarini Group as a therapeutic option for cancers including acute myeloid leukemia (AML). The licensing contract with Menarini was executed in March 2017, and currently, Menarini is the sole sponsor of the ongoing phase I/II clinical study. Details of this study can be found at ClinicalTrials.gov under the identifier . Ryvu has also been assisting in translational research on the project.

The data that have been generated in the SEL24 Cohort Expansion part of the study were presented in June 2021 during the American Society of Clinical Oncology (ASCO) and European Hematology Association (EHA) Virtual Congresses. Data reported in the posters confirmed the manageable safety

profile of the drug at the recommended dose and showed preliminary single agent efficacy in relapsed/refractory AML, particularly in patients with IDH mutant disease either naïve or previously exposed to IDH inhibitors.

In the above-mentioned posters, a total of four objective responses across the dose escalation (n=25) and cohort expansion (n=23) in patients with AML were reported, with 3 of those 4 responders harboring an IDH mutation. Notably, three out of five patients with IDH mutations treated at doses of 75-125 mg achieved a CR/CRI, including a patient that previously relapsed on the IDH-inhibitor enasidenib. Furthermore, one patient with an IDH1 mutation achieved a CRI and underwent allogeneic HSCT.

Menarini stated that these results warrant further investigation of SEL24 in AML, with the potential to focus on the IDH subset. A subsequent study in this patient population started in July 2021.

On November 4, 2021, Menarini announced that the U.S. Food and Drug Administration (FDA) has granted orphan drug designation (ODD) to SEL24 for the treatment of AML.

In June 2022 during the ASCO Annual Meeting and at the EHA Hybrid Congress 2022 Menarini presented a poster entitled: “Phase 1/2 study of SEL24/MEN1703, a first-in-class dual PIM/FLT3 kinase inhibitor, in patients with IDH1/2-mutated acute myeloid leukemia: The DIAMOND-01 trial”.

As of 21 April 2022 (cut-off date), 25 patients were enrolled in the IDHm cohort. Fourteen patients had IDH2, 1 had IDH1/2, and 9 had IDH1 mutations. Concomitant mutations in FLT3-ITD were detected in 4 patients. The median duration of treatment was 2 cycles. In total, 15 patients completed  $\geq 1$  treatment cycle and were efficacy evaluable. The ORR was 13%. One patient with IDH2 and NPM1 mutations had a partial remission at cycle 4 and achieved a CR at cycle 13. One patient with an IDH1 mutation achieved a CRh at cycle 3 and underwent hematopoietic stem cell transplant. These preliminary results in the IDHm cohort confirm that SEL24/MEN1703, has a manageable safety profile and single-agent activity in patients with R/R IDHm AML.

During the ASH Annual Meeting & Exposition in December 2022, Menarini and its collaborators presented translational data on SEL24 (MEN1703). There were four posters on combination therapy of SEL24 (MEN1703) with gilteritinib and SEL24 (MEN1703)-induced PIM inhibition and mechanism of action demonstrated *in vitro* in multiple myeloma (MM), classical Hodgkin lymphoma-tumor-associated macrophages (cHL-TAMs), and diffuse large B-cell lymphoma (DLBCL) models showing the potential of SEL24 (MEN1703) in these malignancies.

Ryvu receives information on the study progress from Menarini during periodic technical and joint steering committee meetings. Based on information received by Ryvu in March 2023, Menarini is considering additional clinical trials in order to explore better the potential of SEL24 (MEN1703) in various disease settings, however, as of the date of the report no further clinical development plans have yet been approved by the Menarini management.

## **RVU120 (SEL120)**

RVU120 (also known as SEL120) is a clinical stage, selective, first-in-class dual inhibitor of CDK8 and CDK19 kinases. RVU120 has demonstrated efficacy in a number of solid tumors and hematologic malignancies in *in vitro* and *in vivo* models. CDK8 and its paralog CDK19 are kinase submodules of the mediator complex, involved in both transcriptional activation and repression, having central roles in

the maintenance of cancer cell viability and undifferentiated state for a variety of tumor types (Dannappel et al. 2019; Rzymiski et al. 2015; Philip et al. 2018). CDK8/19-mediator complex integrates basal transcriptional machinery with the activity of oncogenic transcriptional and epigenetic factors. Inhibition of CDK8/19 can repress key oncogenic transcriptional programs and induce lineage commitment genes in AML. CDK8 and CDK19 are preclinically validated novel targets for the treatment of breast and prostate cancers. Targeting CDK8 and CDK19 using RVU120 may be an effective treatment for both hematologic malignancies and solid tumors with deregulated transcription.

RVU120 has been internally discovered by Rvuu and has received support from the Leukemia & Lymphoma Society Therapy Acceleration Program® (TAP), a strategic initiative to partner directly with innovative biotechnology companies and leading research institutions to accelerate the development of promising new therapies for blood cancers.

On March 25, 2020, the U.S. Food and Drug Administration (FDA) granted an orphan drug designation (ODD) to RVU120 for the treatment of patients with AML.

At present, Rvuu is conducting two clinical studies with RVU120: (i) Phase Ib in patients with AML/HR-MDS (NCT04021368) and (ii) Phase I/II in patients with relapsed/refractory metastatic or advanced solid tumors (NCT05052255). Additionally, multiple translational research activities are underway, aimed at further confirmation of the RVU120 mechanism of action, defining the target patient population and potential combination partners, as well as validating RVU120 in other hemato-oncology as well as solid tumor indications.

The primary aim of the ongoing first-in-human (FIH) Phase Ib study with RVU120 in relapsed or refractory AML or high-risk MDS (CLI120-001 [RIVER-51], NCT04021368) is to evaluate the safety and tolerability of RVU120 as well as to determine the recommended dose for Phase II (RP2D). The secondary endpoints include measurements of pharmacokinetic (PK) properties and an assessment of signs of clinical activity. Response to RVU120 will be evaluated by individual response criteria per each disease predefined in the study protocol. In addition, the exploratory objective of the study is the investigation of the relevant pharmacodynamic (PD) response by studying biomarkers of target engagement in patient samples, such as STAT5 phosphorylation, and identification of molecular markers that might point to a better response to treatment with RVU120.

The first patient in the RIVER-51 clinical trial was dosed in September 2019. The study is currently enrolling at seven investigational sites in the US and Poland. An abstract was accepted for poster presentation at the European Hematology Association Congress in Frankfurt in June 2023. At the data cut off for the abstract (February 28, 2023), 22 patients had been treated at doses up to 110 mg. RVU120's safety profile continues to be favorable. No DLTs were observed and no study drug interruptions due to adverse drug reactions occurred. Clinically significant signs of efficacy were observed in 10 out of 19 evaluable patients with either a complete remission, blast reductions, or evidence of hematologic improvement. The cohort at the dose of 135 mg was open for enrollment at the data cut-off.

The other ongoing clinical study with RVU120 (RVU120-SOL-021 [AMNYS-51], NCT05052255) is a Phase I/II study aiming to investigate the safety and efficacy of RVU120 in patients with relapsed/refractory metastatic or advanced solid tumors. The study is designed in two parts. Part 1 of the study (Phase I) is a dose escalation part according to a standard 3+3 design and is aimed at the enrollment of adult patients with solid malignancies who have failed available standard therapies. The primary objective of the Phase I part is to determine safety, tolerability and the RP2D. The secondary objectives include

the determination of the pharmacokinetic (PK), pharmacodynamic (PD), and preliminary anti-tumor activity of RVU120 as a single agent. Part 2 (Phase II) is aimed both at safety and efficacy expansion. Part 2 will enroll patients with specific tumor types, either as a single agent or combined with standard anticancer medicinal agents. Additional translational and biomarker studies are currently ongoing to confirm which target patient populations will be selected.

The study is currently enrolling at five investigational sites in Poland and Spain. Preliminary data of the dose escalation part were presented as a poster at the 34th EORTC-NCI-AACR Symposium in October 2022. After the data cut-off for that conference, a biomarker inhibition of >70% has been achieved in a patient dosed at the 135mg cohort. Based on preclinical assumptions, this threshold is sufficient to obtain high efficacy in selected patient groups with hematologic malignancies. As of March 3, 2023, enrollment was ongoing at the dose of 375 mg EOD.

**Recent achievements in RVU120 clinical development:**

- **Poster presentation at the 34th EORTC-NCI-AACR Symposium** in Barcelona in October 2022, Preliminary data from the ongoing dose escalation part of AMNYS-51 patients with relapsed/refractory metastatic or advanced solid tumors were presented. As of the cut-off date, 17 patients had been treated with RVU120 at doses between 75 mg and 175 mg. The adverse event profile was favorable with mild or moderate gastrointestinal events as the most frequent. There were no drug-related serious adverse events (SAEs), no dose-limiting toxicities (DLTs), and no adverse event was leading to drug discontinuation. A dose-dependent increase of RVU120 exposure was observed with expected variability. pSTAT5 inhibition as a marker of target engagement correlated with exposure and a more than 60% inhibition was observed at a dose of 135 mg. Disease stabilization was achieved in 4 out of 11 evaluable patients, of which 3 lasted for more than 4 months.
- **Poster presentation at the ASH Annual Meeting & Exposition** in December 2022, updated safety and efficacy data were presented of a total of 19 patients (16 patients with AML, 3 patients with HR-MDS). Nine of 16 evaluable patients showed clinical benefit: one patient with AML had a complete response, four patients had hematologic improvement and four patients had blast reductions. RVU120 demonstrated a favorable safety profile. A meaningful inhibition of pSTAT5 of >70% has been observed in patients treated with RVU120 in a dose- and exposure-dependent manner.

In the opinion of Ryvu's Management Board, the data warrant continuation of dose escalation and collection of additional clinical data.

Further translational research showed that patient-derived AML cells with DNMT3A and NPM1 mutations are more sensitive to RVU120 treatment both *in vitro* and *in vivo*. This observation is consistent with the clinical responses to RVU120 in CLI120-001 (RIVER-51) study in two patients that harbored DNMT3A and NPM1 mutations. Anti-cancer efficacy of RVU120 was associated with transcriptomic reprogramming and lineage commitment.

## PRECLINICAL AND DISCOVERY STAGE PROJECTS

### Synthetic lethality projects

Ryvu is carrying out several discovery stage projects in the area of synthetic lethality. Ryvu's most advanced project in the field of synthetic lethality focuses on cancers with a deletion of the metabolic gene MTAP, which occurs in 10 to 15% of all human tumors. MTAP deletion results in massive accumulation of methylthioadenosine (MTA) in cells. MTA at high concentrations is a very selective inhibitor of PRMT5 methyltransferase, competitive for the substrate: S-adenosylmethionine (SAM). Accumulation of MTA in cells with MTAP deletion causes a partial inhibition of the methylation activity of PRMT5, which in turn reduces the level of symmetric arginine dimethylation of the whole proteome, and thus an increased sensitivity of cells to modulation of methylosome activity. The Company's strategy is to develop MTA-cooperative PRMT5 inhibitors, which will selectively inhibit the growth of MTAP-deleted cancer cells.

The work carried out in Q1 2023 focused on the expansion of the main chemical series into a lead series with the key aim to demonstrate in vivo proof of concept, which would then allow for the nomination of a preclinical candidate in 2023. Experimental works on improving the properties of the chemical series were continued with respect to potency, selectivity measured by the inhibition of SDMA in MTAP-deleted versus MTAP WT cells, and particularly PK parameters in rodent species (necessary for pharmacological and toxicological characterization). Ryvu compounds selectively inhibit growth of MTAP-deleted cancer cells in prolonged 3D culture, which strongly correlates with inhibition of PRMT5-dependent protein symmetric arginine dimethylation (SDMA) in those cells. Selectivity between effects observed in MTAP-deleted and WT cells exceeds for multiple compounds in the series 100-fold both for SDMA and growth inhibition. Additionally, in vivo efficacy studies have demonstrated that tool lead compounds have achieved significant tumor growth inhibition in a colorectal cancer model with MTAP deletion.

Further optimization allowed for selection of new, improved derivatives for larger scale synthesis and efficacy studies in animal models, which are planned in Q2 and Q3 2023.

Results on the development of MTA-cooperative PRMT5 inhibitors including summary of optimization progress and an early lead compound profile, together with in vivo results in a mouse model showing tumor growth inhibition and pharmacodynamic biomarkers in MTAP-deleted tumors were presented at the annual AACR American Association for Cancer Research conference in Orlando, United States in April 2023.

The second disclosed project focuses on the identification and development of first-in-class, small molecule inhibitors of the WRN (Werner Syndrome helicase) helicase, which plays an important role in cell proliferation, the replication stress response, and DNA repair. Scientific reports reveal promising synthetic lethal interaction between inactivation/inhibition of the WRN protein and impaired viability of cancer cells with microsatellite instability (MSI). This strategy is an attractive therapeutic approach in the treatment of MSI cancers, responsible for 10-30% of colorectal, ovarian, endometrial and gastric tumors.

Ryvu's WRN project was initiated by several high-throughput screening (HTS) campaigns that provided a range of small-molecule WRN-inhibiting actives, characterized by different scaffolds. In order to enhance the essential properties, further expansion and profiling have been conducted on the selected chemotypes, distinguished by their varied physicochemical characteristics. During the initial quarter of

2023, ligands with confirmed binding to the target protein and low molar WRN helicase inhibitory activity were developed and confirmed in recombinant biophysical and biochemical assays. In the next quarters, expansion of further derivatives will be continued, which will allow the assessment of the structure-activity relationship (SAR) for the newly obtained chemical series. The applied strategy assumes further optimization of physicochemical properties, activity, and ADME parameters in order to select a development candidate in 2024.

### **New, undisclosed targets**

In addition to the two disclosed projects, Ryvu is currently running a number of internal initiatives focused on identifying and validating new targets in the field of synthetic lethality, with potential for first-in-class drug discovery. Work is currently underway to validate several therapeutic targets identified so far.

### **Target discovery**

Ryvu's continues also the efforts within an innovative target discovery platform based on genome-wide screening in cancer cells with defined genotype. The methodology enables the detection of new biological targets that meet the definition of synthetic lethality and other candidates for targeted therapies (e.g. disease specific, actionable oncogenic drivers). These therapies will target genetically stratified patient populations in which the tumor genotype significantly increases the chances of a clinical response. The Ryvu platform enables modeling the influence of the tumor microenvironment and the use of cells directly isolated from patients' tumors (primary cells) in high-throughput screening. Our platform is currently used for genomic alterations (mutations or deletions of genes) with the biggest unmet medical need.

### **Collaboration with BioNTech on Immunotherapy and STING**

On November 29, 2022, BioNTech and Ryvu entered into a multi-target research collaboration for several small molecule immunotherapy programs as well as an exclusive license agreement for Ryvu's STING agonist portfolio as standalone small molecules. BioNTech received a global, exclusive license to develop and commercialize Ryvu's STING agonist portfolio as standalone small molecules, including as monotherapy and in therapeutic combinations. Under the collaboration a selected candidate molecule will enter further preclinical development necessary to complete the IND package as well as to initiate first-in-human trials. Ryvu know-how on STING agonism combined with the vast expertise of BioNTech in immunology and immune-oncology will enable new clinical development strategies to fully utilize the potential of the candidate STING agonist. Current project progress remains confidential.

In addition, BioNTech and Ryvu will jointly undertake drug discovery and research projects to develop multiple small molecule programs directed at exclusive targets selected by BioNTech, primarily focused on immune modulation within oncology, with potential applications in other disease areas. BioNTech has the option to license global development and commercialization rights to these programs at the development candidate stage. Multiple research programs are underway jointly but remain confidential.

### **Collaboration with Exelixis on STING ADCs**

On July 7, 2022, Exelixis and Ryvu entered into an exclusive license agreement focused on the development of novel targeted therapies utilizing Ryvu's STING technology. As part of the optimization of the STING agonists developed by Ryvu, the company identified active compounds with a variety of chemical groups that allow easy combination with a reactive chemical group. This modification allows for further development of agonists in the form of antibody-drug conjugates (ADC), where the antibody enables targeted delivery of the active STING agonist. The high potential of compounds developed by Ryvu led to an exclusive licensing agreement with Exelixis. Under the terms of agreement, Exelixis develops antibody-drug conjugates leveraging Ryvu's expertise in STING protein biology and Exelixis' expertise in antibody engineering and ADC technology.

In January 2023, the first milestone within the collaboration was achieved, which in line with the agreement made Ryvu entitled to a payment of USD 1 million. Further project progress currently remains confidential.

### **Collaboration with Galapagos on Inflammation**

On April 16, 2020, Galapagos and Ryvu entered into a collaboration focused on the discovery and development of novel small molecule drugs in inflammation. The subject of the Agreement was the research and development of novel small molecule compounds with therapeutic potential in inflammatory diseases, based on the drug target identified by the Company and its scientific platform. The Company informed about the conclusion of the Agreement in the current report no. 7/2020 dated April 16th of 2020. On December 14, 2021, the companies announced that Galapagos exercised its exclusive option for the program. The joint research collaboration is focused on the discovery and development of novel small molecule drugs in inflammation. In November 2022, Galapagos announced a strategy to focus its R&D investment in the areas of immunology and oncology and subsequently, as a result of this strategic decision, on April 5, 2023, Ryvu received a notice of termination with reference to Option, License and Research Collaboration Agreement ("Agreement"). After analyzing the data package received from the Galapagos, the Management Board of Ryvu will consider the use of intellectual property opportunities in Ryvu projects or its re-commercialization.

### **OTHER PROJECTS**

Ryvu is also developing small molecule modulators of HPK1 (MAP4K1), a hematopoietic cell-restricted member of Ste 20 serine/threonine kinases. HPK1 is known as a negative regulator of TCR signaling. Inhibition of HPK1 leads to TCR-induced phosphorylation of SLP-76, which undergoes phosphorylation-dependent ubiquitination and results in its degradation, thereby blocking signal transduction - required for immune system activation and elimination of cancer cells. The results of the project so far are compounds with high selectivity for HPK1 kinase, metabolic stability and in vivo activity in a mouse model for selective pharmacodynamic biomarkers. At the same time, the main chemical series and the lead compound require further improvement of the safety parameters related to the potential risk of cardiotoxicity and an increase of the therapeutic window.

## 2.2 Significant events in Q1 2023

### A) DURING THE REPORTING PERIOD

#### Registration of amendment of the Company's Articles of Association concerning share capital

On January 17, 2023 the District Court for Kraków-Śródmieście in Kraków, XI Commercial Division of the National Court Register, registered an amendment to the Company's Articles of Association concerning increasing the Company's share capital from the amount of PLN 7,342,189.60 PLN (seven million three hundred forty-two thousand one hundred eighty-nine zlotys and sixty groszy) to the amount of 9,248,059.20 PLN (nine million two hundred forty-eight thousand fifty-nine zlotys and twenty groszy), by way of issue of 4,764,674 (four million seven hundred sixty-four thousand six hundred seventy-four) new series J ordinary bearer shares with a nominal value of PLN 0.40 (forty groszy) each ("Series J Shares") within the authorized capital, made pursuant to Resolution No 1 of the Company's Management Board of 5 October 2022 on increasing the Company's share capital within the limits of the authorized capital through the issue of series J shares, excluding the pre-emptive rights of the existing shareholders in full and amending the Company's Articles of Association (the "Issue Resolution"), of which the Issuer informed in a current report No 22/2022 of 5 October 2022 (the "Registration of Amendments").

After the Registration of Amendments, the share capital of the Company equals PLN 9,248,059.20 and is divided into 23,120,148 shares with a nominal value of PLN 0.40 (forty groszy) each.

#### Admission and introduction of the series J shares of the Company to trading on the regulated market of the WSE

On January 20th, 2023 the Management Board of the Warsaw Stock Exchange S.A. adopted Resolution No. 51/2023 on the admission and the introduction to exchange trading on the main market of the WSE of series J ordinary bearer shares of the Company, pursuant to which the Management Board of the Warsaw Stock Exchange S.A. stated that 4,764,674 series J ordinary bearer shares of the Issuer with a nominal value of PLN 0.40 each with ISIN code PLSELV00013 ("Series J Shares") are admitted to exchange trading on the main market. The WSE Management Board decided to introduce on January 25th, 2023 the Series J Shares to exchange trading on the main market, subject to the registration of the Series J Shares by the National Depository for Securities S.A. with the ISIN code PLSELV00013 on January 25th, 2023.

#### Achievement of the first milestone under the license agreement with Exelixis Inc.

On January 25th, 2023 the Company has received notice that the first milestone has been achieved in the research collaboration with Exelixis Inc. with its registered office in Alameda, California ("Exelixis") under the license agreement, which was described by the Company in current report 11/2022 of July 7th, 2022. The purpose of the Agreement is to develop novel targeted therapies using the STING (STimulator of INterferon Genes) technology developed by Ryvu (the "Agreement"). Based on the achievement of the milestone, Ryvu is entitled to receive a payment of USD 1 million (PLN 4,326,500 converted at the average exchange rate of the National Bank of Poland on January 25th, 2023 1 USD = 4.3265 PLN).

### **Conclusion of an agreement concerning the operational execution of Phase II of Phase I/II clinical trial of RVU120 in Patients with Relapsed/Refractory Solid Tumors**

On March 2nd, 2023 the Company entered into an agreement with Labcorp Drug Development Inc. ("LabCorp Drug Development"), based in New Jersey, USA to conduct Phase II of Phase I/II of a clinical study to determine the safety and efficacy profile of RVU120 in patients with relapsed/refractory metastatic or advanced solid tumors (the "Agreement").

The Phase I clinical study of RVU120 began on August 13th, 2021. Labcorp Drug Development (then known as Covance Inc.) has cooperated with Rvuv in the operational execution of Phase I clinical trial of RVU120 in solid tumors under the contract about which the Company informed in the current report 5/2021 of March 8th, 2021.

The primary objective of Phase I/II study is to evaluate the anti-tumor activity of RVU120 as a single agent in patients with selected tumor types and to further evaluate the safety and tolerability profile of RVU120. Phase II will be conducted at selected clinical investigational sites in Europe and will start after the selection of the recommended Phase II dose based on Phase I results.

Labcorp Drug Development will be responsible for the operational execution of the Phase II clinical study. The estimated cost of the Agreement is EUR 3,872,088.22 (PLN 18,102,012.43 converted at the average exchange rate of the National Bank of Poland of March 2nd, 2023, EUR 1 = PLN 4,6750) and will be co-financed by the European Regional Development Fund and the Government of Poland as part of the project titled "Clinical development of an innovative drug candidate in solid tumors" within the Smart Growth Operational Programme 2014-2020, measure 1.1.1. "Fast Track". The value of the Agreement may change in the event of extending the scope of the order.

### **B) EVENTS OCCURRED BETWEEN THE END OF REPORTING PERIOD UNTIL THE APPROVAL OF FINANCIAL STATEMENT**

#### **Conclusion of the agreements concerning Phase II start-up services for RVU120 clinical studies in AML/HR-MDS**

On April 20th, 2023 the Company entered into two agreements with Labcorp Drug Development Inc. (Labcorp), based in New Jersey, USA to conduct Phase II study start-up services for the clinical development of RVU120 in hematologic malignancies: (i) RIVER-52 (in patients with AML/HR-MDS; RVU120 monotherapy) and (ii) RIVER-81 (in patients with AML; RVU120 combination), (the "Agreements").

The commencement of both aforementioned Phase II studies in AML/HR-MDS is expected in H2 2023. The start-up activities covered by the Agreements outlined above constitute the second and third RVU120 Phase II studies planned to be initiated by the end of 2023 and are parts of the planned broad RVU120 clinical development across multiple indications (hematology and solid tumors) and in diverse treatment settings (monotherapy and combination therapy), aimed at maximizing the potential of RVU120 and diversifying development risks. Additional clinical trials investigating RVU120 in patients with low-risk MDS or with an MDS/MPN overlap syndrome are in planning.

The estimated total cost of services under Agreements is EUR 1,221,627.57 (PLN 5,632,802.56 converted at the average exchange rate of the National Bank of Poland of April 20th, 2023, EUR 1 = PLN 4.6109). The value of Agreements may change in the event of extending the scope of the order.

## Conclusion of an agreement for the issuance of subscription warrants to the European Investment Bank

On May 4th, 2023, the Company entered into an agreement with the European Investment Bank ("EIB") for the issuance of subscription warrants to the EIB (the "Warrant Agreement"). The execution of the Warrant Agreement is one of the conditions for the disbursement of the first tranche of financing by the EIB under the financing agreement with the Company dated August 16th, 2022 (the "Financing Agreement"), which the Company announced in the current report No. 14/2022 dated August 17th, 2022.

Pursuant to the Warrant Agreement, the Company agreed to issue 592,825 subscription warrants (the "Warrants") to the EIB, entitling it to subscribe for a total of 592,825 shares of the Company with a total par value of PLN 237,130 (the "Shares"). The essential provisions of the Warrant Agreement are as follows: (i) the Warrants will be acquired by the EIB free of charge and will entitle the holder to subscribe for Shares of the Company at an issue price equal to the par value of each Share; (ii) the rights under the Warrants to subscribe for Shares may be exercised over a period of 10 years. The Warrant Agreement regulates the terms and conditions for the exercise of the rights under the Warrants to subscribe for Shares, making this right contingent, in particular, on the disbursement of further tranches of financing under the Financing Agreement and the occurrence of other events specified in the Warrant Agreement; (iii) the Warrants will be transferable. The Warrant Agreement sets forth the rules for the transfer and purchase of the Warrants, including providing for the Company's obligation to purchase the Warrants from the holder of the Warrants for redemption against payment in the cases specified in the Warrant Agreement; (iv) in the occurrence of events causing dilution of the Company's share capital, EIB will be entitled to acquire additional Subscription Warrants, in a number ensuring that EIB maintains a level of 2.5% of the Company's fully diluted share capital, subject to the exceptions provided for in the Warrant Agreement.

The Agreement regulates the Company's obligations to obtain the EIB's approval for certain activities and its disclosure obligations to the EIB. The issuance of the Warrants to the EIB is part of the remuneration to the EIB for providing financing under the Financing Agreement.

## New clinical and preclinical data on RVU120 to be presented at the upcoming 2023 European Hematology Association Congress

The updated safety and efficacy data from the Phase 1b dose-escalation study of RVU120 in patients with Acute Myeloid Leukemia (AML) and High-Risk Myelodysplastic Syndromes (HR-MDS) and nonclinical data of RVU120 in combination with JAK1/2 inhibitor Ruxolitinib (RUX) in myeloproliferative neoplasms will be presented at the Annual European Hematology Association (EHA) 2023 Hybrid Congress, taking place June 8-11, 2023 in Frankfurt, Germany.

As of February 2023, treatment with single-agent RVU120 has led to complete remission in one patient, an increase of hemoglobin and platelets in four patients, and bone marrow blast reduction in five patients, including in a patient with TP53 double-hit. Based on these encouraging clinical benefits, favorable safety, and no dose-limiting toxicities, the Company's Management Board plans to continue the dose escalation phase, expecting further increases in RVU120's anti-leukemic and erythroid-stimulating activity at increased doses.

The data suggest synergistic effects between RVU120 and RUX in myelofibrosis by demonstrating a significant reduction of disease manifestation in vivo. In the opinion of the Company's Management

Board, these data reinforce the potential emerging role of targeting both CDK8/19 and JAK1/2 in myeloproliferative neoplasms.

## **2.3 Unusual events occurring in the reporting period**

### **Conflict in Ukraine**

Due to the outbreak of the conflict in Ukraine, the Issuer's Management Board has analyzed the potential impact of the ongoing war on the Issuer's operations. In the opinion of the Management Board, apart from the currency risk, the Management Board did not identify any other significant risks that could affect the Issuer's operations.

In particular, it should be noted that the Issuer does not have any assets in Ukraine and does not conduct business and operations in Ukraine and Russia. The share of entities from Ukraine or Russia as suppliers in the Issuer's structure remains insignificant and is mostly limited to the provision of compound libraries for discovery stage projects at their early stage.

Nevertheless, the Management Board of the Company analyzes the Issuer's situation on an ongoing basis. Any new circumstances having a significant impact on the financial results and business situation of the Issuer will be communicated to investors.

## 3. THE ISSUER'S CORPORATE BODIES

### Issuer's Management Board:

- 1) Paweł Przewięźlikowski – President of the Management Board
- 2) Krzysztof Brzózka – Vice President of the Management Board
- 3) Kamil Sitarz – Member of the Management Board
- 4) Vatnak Vat-Ho – Member of the Management Board
- 5) Hendrik Nogai – Member of the Management Board

### Issuer's Supervisory Board:

- 1) Piotr Romanowski – Chairman of the Supervisory Board
- 2) Tadeusz Wesołowski – Vice Chairman of the Supervisory Board
- 3) Rafał Chwast – Supervisory Board Member
- 4) Axel Glasmacher – Supervisory Board Member
- 5) Jarl Ulf Jungnelius – Supervisory Board Member
- 6) Thomas Turalski – Supervisory Board Member

### Issuer's Audit Committee:

- 1) Rafał Chwast – Chairman of the Audit Committee
- 2) Piotr Romanowski – Member of the Audit Committee
- 3) Tadeusz Wesołowski – Member of the Audit Committee
- 4) Jarl Ulf Jungnelius – Member of the Audit Committee

### Issuer's Remuneration Committee:

- 1) Piotr Romanowski – Chairman of the Remuneration Committee
- 2) Axel Glasmacher – Member of the Remuneration Committee
- 3) Thomas Turalski – Member of the Remuneration Committee

## 4. INFORMATION ON THE SHAREHOLDERS HOLDING (DIRECTLY OR INDIRECTLY) AT LEAST 5% OF THE TOTAL NUMBER OF VOTES AT THE GENERAL SHAREHOLDERS' MEETING OF THE COMPANY AND ON SHARES HELD BY MEMBERS OF THE ISSUER'S MANAGEMENT BOARD AND SUPERVISORY BOARD

### Shares held by members of the Management and Supervisory Board of the Company as of 31.03.2023

Shareholder	Preferred shares*	Ordinary shares	Number of shares	% of Share Capital	Number of Votes	% of Votes at SM
<b>The Management Board</b>						
<b>Paweł Przewięźlikowski</b>	3 500 000	639 544	<b>4 139 544</b>	17,90%	<b>7 639 544</b>	28,12%
<b>Krzysztof Brzózka</b>		267 321	<b>267 321</b>	1,16%	<b>267 321</b>	0,98%
<b>Kamil Sitarz</b>		21 365	<b>21 365</b>	0,09%	<b>21 365</b>	0,08%
<b>Vatnak Vat-Ho</b>		18 500	<b>18 500</b>	0,08%	<b>18 500</b>	0,07%
<b>Hendrik Nogai</b>		9 000	<b>9 000</b>	0,04%	<b>9 000</b>	0,03%
<b>The Supervisory Board</b>						
<b>Tadeusz Wesołowski (directly)</b>		92 975	<b>92 975</b>	0,40%	<b>92 975</b>	0,34%
<b>Tadeusz Wesołowski (indirectly through Augebit FIZ**)</b>		1 279 738	<b>1 279 738</b>	5,54%	<b>1 279 738</b>	4,71%
<b>Piotr Romanowski</b>		174 000	<b>174 000</b>	0,75%	<b>174 000</b>	0,64%
<b>Rafał Chwast</b>		121 115	<b>121 115</b>	0,52%	<b>121 115</b>	0,45%
<b>Thomas Turalski</b>		20 100	<b>20 100</b>	0,09%	<b>20 100</b>	0,07%

\*A single preferred share entitles to two votes at the Shareholder Meeting.

\*\*The beneficiary of Augebit FIZ is Tadeusz Wesołowski - Vice-Chairman of the Issuer's Supervisory Board.

## Shares held by members of the Management and Supervisory Board of the Company as of the date of Report publication

Shareholder	Preferred shares*	Ordinary shares	Number of shares	% of Share Capital	Number of Votes	% of Votes at SM
<b>The Management Board</b>						
Paweł Przewięźlikowski	3 500 000	639 544	<b>4 139 544</b>	17,90%	<b>7 639 544</b>	28,12%
Krzysztof Brzózka		267 321	<b>267 321</b>	1,16%	<b>267 321</b>	0,98%
Kamil Sitarz		21 365	<b>21 365</b>	0,09%	<b>21 365</b>	0,08%
Vatnak Vat-Ho		18 500	<b>18 500</b>	0,08%	<b>18 500</b>	0,07%
Hendrik Nogai		9 000	<b>9 000</b>	0,04%	<b>9 000</b>	0,03%
<b>The Supervisory Board</b>						
Tadeusz Wesołowski (directly)		92 975	<b>92 975</b>	0,40%	<b>92 975</b>	0,34%
Tadeusz Wesołowski (indirectly through Augebit FIZ**)		1 279 738	<b>1 279 738</b>	5,54%	<b>1 279 738</b>	4,71%
Piotr Romanowski		100 000	<b>100 000</b>	0,43%	<b>100 000</b>	0,37%
Rafał Chwast		121 115	<b>121 115</b>	0,52%	<b>121 115</b>	0,45%
Thomas Turalski		20 100	<b>20 100</b>	0,09%	<b>20 100</b>	0,07%

\*A single preferred share entitles to two votes at the Shareholder Meeting.

\*\*The beneficiary of Augebit FIZ is Tadeusz Wesołowski - Vice-Chairman of the Issuer's Supervisory Board.

## Shares held by significant shareholders of the Company as of 31.03.2023 and as of the date of Report publication

Shareholder	Shares	% [Shares]	Votes	% [Votes]
Paweł Przewięźlikowski	4 139 544	17,90%	7 639 544	28,12%
Bogusław Sieczkowski	825 348	3,57%	1 375 348	5,06%
Tadeusz Wesołowski (with Augebit FIZ)	1 372 713	5,94%	1 372 713	5,05%
Nationale Nederlanden OFE	1 900 980	8,22%	1 900 980	7,00%
PTE Allianz Polska S.A.	2 132 540	9,22%	2 132 540	7,85%
TFI Allianz Polska S.A.	1 689 419	7,31%	1 689 419	6,22%
BioNTech SE	1 917 437	8,29%	1 917 437	7,06%

The above information on the state of possession of the Issuer's shares by shareholders (including those being members of the Company's bodies) holding directly and indirectly at least 5% of the total number of votes at the Company's General Meeting has been prepared on the basis of information obtained from shareholders through their performance of duties imposed on shareholders of public companies by virtue of relevant legal regulations, including on the basis of the provisions of the Act of 29. July 2005 on public offering and the conditions for introducing financial instruments to the organized trading system and on public companies (art. 69 and art. 69a) and on the basis of the provisions of the Regulation (EU) No 596/2014 of the European Parliament and of the Council of 16 April 2014 on market abuse and repealing Directive 2003/6/EC of the European Parliament and of the Council and Commission Directive 2003/124/EC, 2003/125/EC and 2004/72/EC (MAR Regulation, art. 19). In addition, information on the ownership of the Company's shares is provided on the basis of publicly available data on the portfolio exposure and asset structure of investment funds or pension funds, including information on the number of shares registered at the Company's General Meeting (data available periodically, inter alia, based on information from the financial statements of investment funds and pension funds - data may be subject to change since the date of publication of the last information).

## 5. STATEMENT OF THE MANAGEMENT BOARD REGARDING APPLICABLE ACCOUNTING PRINCIPLES

The Management Board of Ryvu Therapeutics S.A. confirms that, to the best of its knowledge, the financial statements have been prepared in accordance with the applicable accounting principles and reflect in a true, reliable and clear manner the financial situation of Ryvu Therapeutics S.A. and its financial results. Report of the Management Board on the activities of Ryvu Therapeutics S.A. contains a true picture of the development and achievements, including a description of the basic threats and risks.

## 6. ADDITIONAL INFORMATION

### **Proceedings pending at court, before an arbitration institution or a public administration authority**

Company has filed a lawsuit against Mota-Engil Central Europe S.A. in connection with construction of the Research and Development Center for the payment of PLN 13,756,717.07. With this lawsuit, the Company seeks claims related to the agreement for "Construction of the Research and Development Center of Innovative Drugs Selvita S.A.", the conclusion of which was announced by the Company in the current report No. 27/2018 of August 13, 2018. The total value of the Contract was PLN 68.783.585,34 including VAT. Proceedings are on the stage of a pre-trial hearing.

Mota-Engil has filed a lawsuit for payment against to the Regional Court in Kraków in connection with the performance of the general contractor agreement for the project entitled: "Construction of the Research and Development Center for Innovative Drugs Selvita S.A.". In the lawsuit the Contractor is claiming damages for the costs incurred in connection with prolonged performance of the Contract, the unpaid portion of the lumpsum fee as well as supplementary remuneration for additional, replacement and omitted works (PLN 5,391,425.63) as well as damages resulting from the Company's unauthorized - in the Contractor's opinion - application of the performance bond and removal of the defects and faults (PLN 2,063,507.56). With the statutory interests, the Contractor demands from the Company a total amount of PLN 7,671,285. At a court hearing held on March 31, 2023, a schedule for the testimony of all witnesses and questioning of the parties was set. The dates of the hearings were planned between 30.06.2023 and 22.11.2023.

### **Significant non-arm's length transactions with related entities**

Not applicable.

### **Information on organizational or capital relations of the Issuer with other entities**

As at the publication date of the report, the Issuer does not form a Capital Group. As at the date of this Report, the Issuer holds 3.15% of shares in NodThera Inc.

### **Warranties for loans and borrowings and guarantees granted**

Not applicable.

### **Other information significant for the assessment of the Issuer's position in the area of human resources, assets, cash flows, financial results and changes thereof and information significant for the assessment of the Issuer's ability to settle its liabilities**

Not applicable.

### **Factors which, in the Issuer's opinion, will affect the results over at least the following quarter**

The results of the subsequent quarters will depend primarily on the execution of the Company's strategy, which assumes in particular that the following business objectives will be met:

- Completing the ongoing Phase I clinical studies of RVU120 in AML/HR-MDS and solid tumors;

- Expanding therapeutic potential of RVU120 by initiating broad Phase II clinical development across multiple indications (hematology and solid tumors) and in diverse treatment settings (monotherapy and combination therapy);
- Supporting clinical development of our partnered candidate, SEL24 (MEN1703) by Menarini Group;
- Conducting preclinical development and advancing into Phase I clinical trials of one new program;
- Strengthening of our Synthetic Lethality Platform and accelerating progress in the early pipeline;
- Achieving financial milestones in the existing R&D collaborations (i.e. BioNTech, Exelixis, Menarini);
- Signing at least one new partnering agreement per year.

**Description of factors and events, in particular of an unusual nature, having a significant effect on the financial performance**

In the reported period, the Covid-19 pandemic occurred. The Issuer described its effect on the Company's operations under Significant events that occurred in the reporting period.

**Explanations regarding the seasonal or cyclical nature of the Issuer's operations in the reported period**

Not applicable.

**Information on inventory write-downs to the net realizable amount and reversal of such write-downs**

Not applicable.

**Information on impairment write-downs in respect of financial assets, tangible fixed assets, intangible assets or other assets and the reversal of such write-downs**

Not applicable.

**Information on the set-up, increase, utilization and reversal of provisions**

Information on the changes in provisions for holidays and bonuses is provided in note 27 to the financial statements.

**Information on deferred income tax provisions and assets**

Information on deferred income tax provisions and assets is provided in note 10 to the financial statements.

**Information on significant purchases or disposals of tangible fixed assets**

Information on tangible fixed assets is provided in note 12 to the consolidated financial statements.

**Information on significant liabilities in respect of purchases of tangible fixed assets**

Information on the liabilities in respect of purchases of tangible fixed assets is provided in note 33 to the consolidated financial statements.

**Information on significant settlements resulting from court cases**

Not applicable.

**Error corrections relating to previous periods**

Not applicable.

**Information on changes in the economic situation and business conditions, which have a significant effect on the fair value of the entity's financial assets and financial liabilities**

Not applicable.

**Information on the failure to repay a loan or borrowing or a breach of significant terms and conditions of a loan agreement, with respect to which no corrective action had been taken by the end of the reporting period**

Not applicable.

**Information on changes in the method of valuation of financial instruments measured at the fair value**

Not applicable.

**Information on changes in the classification of financial assets due to a change in their purpose**

Not applicable.

**Information on the issue, redemption and repayment of non-equity and equity securities**

Not applicable.

**Information on dividends paid (or declared) in the total amount and per share, divided into ordinary and preference shares**

Not applicable.

**Events that occurred after the date for which the quarterly financial statements were prepared, not disclosed in these financial statements although they may have a significant effect on the Issuer's future financial results**

Information on events that occurred after the date for which the financial statements were prepared is provided in note 36 to the financial statements.

**Information on changes in contingent liabilities or contingent assets that occurred after the end of the last financial year**

Information on changes in contingent liabilities or contingent assets is provided in note 34 to the financial statements.

**Other disclosures which may have a material impact on the assessment of the Issuer's financial position and results of operations**

Not applicable.

**Amounts and types of items affecting the assets, liabilities, equity, net profit/ (loss) or cash flows, which are unusual in terms of type, amount or frequency**

Not applicable.

Krakow, May 15, 2023

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Paweł Przewięźlikowski  
President of the Management Board

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Krzysztof Brzózka  
Vice-President of the Management Board

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Kamil Sitarz  
Management Board Member

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Vatnak Vat-Ho  
Management Board Member

---

Hendrik Nogai  
Management Board Member

# CONTACT



## RYVU THERAPEUTICS

2 Sternbacha Street

30-394 Krakow, Poland

P: +48 12 314 02 00



## GENERAL INQUIRIES

[ryvu@ryvu.com](mailto:ryvu@ryvu.com)