Filed pursuant to Rule 433 of the Securities Act of 1933 Issuer Free Writing Prospectus dated June 25, 2021 Relating to the Preliminary Prospectus dated June 25, 2021 Registration Statement File No. 333-257427



### Disclaimer

This presentation is provided by Kiromic BioPharma, Inc. (the "Company") for the express purpose of giving prospective investors, bankers, employees, consultants, and corporate affiliations information regarding the Company to assist the recipient in evaluating a potential formal association with the Company. While the information contained herein or in any other materials that may be provided by the Company is believed to be true, accurate and reasonable, the Company makes no such representation or warranty, express or implied, as to the veracity, accuracy, reasonableness or completeness of such information. The Company expressly disclaims any and all liability which may be based on such information, any errors therein or omissions

NASDAQ: KRBP

therefrom. This presentation does not imply an offering of Securities. This presentation may contain forward-looking statements within the meaning of applicable securities regulations. All statements other than statements of historical facts are forward-looking statements. In some cases, forward-looking statements may be identified by the use of words such as "anticipate," "believe," "plan," "estimate," "expect," "intend," "may," "will," "would," "could," "should," "might," "potential," or "continue" and variations or similar expressions. Readers should not unduly rely on these forward-looking statements, which are not a guarantee of future performance. There can be no assurance that forward-looking statements will prove to be accurate, as all such forward-looking state-



ments involve known and unknown risks, uncertainties and other factors which may cause actual results or future events to differ materially from the forward-looking statements. Such risks include, but may not be limited to: general economic and business conditions; technology changes; competition; changes in strategy or development plans; governmental regulations and the ability or failure to comply with governmental regulations; the timing of anticipated results; and other factors referenced in the Company's business materials and prospectuses.





This presentation highlights basic information about us and the proposed offering. Because it is a summary, it does not contain all of the information that you should consider before investing. We have filed a registration statement (including a prospectus) with the SEC for the offering to which this presentation relates. The registration statement has not yet become effective. Before you invest, you should read the prospectus in the registration statement (including the risk factors described therein) and other documents we have filed with the SEC for more complete information about us and the offering.

You may access these documents for free by visiting EDGAR on the SEC Web site at http://www.sec.gov. Alternatively, we or any underwriter participating in the offering will arrange to send you the prospectus if you contact ThinkEquity, a division of Fordham Financial Management, Inc., Prospectus Department, 17 State Street, 22nd Floor, New York, New York 10004, telephone: (877) 436-3673 or e-mail: prospectus@think-equity.com.

This presentation shall not constitute an offer to sell, or the solicitation of an offer to buy, nor will there be any sale of these securities in any state or other jurisdiction in which such offer, solicitation or sale would be unlawful prior to the registration or qualification under the securities laws of such state or jurisdiction. The offering will only be made by means of a prospectus pursuant to a registration statement that is filed with the SEC after such registration statement becomes effective.



## **Offering Summary**

lssuer	Kiromic Biopharma, Inc.
Listing Symbol	NASDAQ / KRBP
Offering Size	~\$50,000,000 of Common Stock
<b>Over-Allotment Option</b>	15% (100% primary)
Use of Proceeds	ALEXIS-ISO-1 / ALEXIS-PRO-1 GMP Facility Expansion IP Protection and Reinforcement IND Applications / IND Enabling Trials General Working Capital
Sole Book-Running Manager	ThinkEquity, a division of Fordham Financial Management, Inc.
NASDAQ: KRBP	





We are connecting the dots in cancer research by using Al and machine learning to connect silos of informations and arrive at cancer targets which will be more effective vs. classic development, saving man-years and billions in development dollars.



### Non-Viral Genome edit and delivery

Our single-cut gene edits carry a lower mutagenesis risk vs. classic double-cut gene edits.

Our CAR receptors will also have higher safety with an on-demand cut-off switch vs. classic CAR therapies with no off-switch.

# **Kiromic at a Glance**

Revolutionizing Next-Gen Allogenic CAR Therapy for Solid Tumors



#### Gamma Delta T-cell Immune Cell Type

Our CAR Therapy will be using off-the-shelf Gamma-Delta T-cells and will have a higher yield and significantly lower yield variability vs. classic CAR-T therapies.



#### **Micro Tumor Environment**

Our CAR Therapies will be able to access the micro tumor environment due to our chPD-1 check-point activator vs. classic CAR-T therapies.

Classic CAR-T are limited to hematologic indications.

Solid Tumor



Diamond is a computational platform and a **neural network** that can identify new cancer immunological targets for T cells and B cells. Diamond is an **artificial intelligence** and **machine learning** approach that can identify novel surface tumor targets. It uses public and proprietary samples and can expand into the tumor target space.

## **ADVANCING CAR through A.I.**







**Big Data Science** 

Manual Target Identification

dramatically compressing

## Man-Years and Billions of Drug Development Dollars

meets

**Target Identification** 

to develop a live drug





# **ADVANCING CAR through A.I.**

**Our Therapeutic Products** 

# Allogenic CAR Immuno CAR-GD-T Therapy

in solid tumors

NASDAQ: KRBP

CAR = chimeric antigen receptors

## **Allogenic Engineered Immune Therapy**

## **Step 01: Fractionation**





Whole Blood



NASDAQ: KRBP



# **Our Pipeline**



	In vitro validation	Pre clinical	IND	Phase 1	Phase 2	Phase 3
Alexis - Pro 1						
Allogenic, Off-the-Shelf γδ chimeric T cells						
Multiple solid tumors						
Alexis - ISO 1						
Allogenic, Off-the-Shelf γδ CAR-T cells						
Multiple solid tumors						
C. L I Francisco	Kiromic plans to perform a dose-escalation schema			Once we reach the Of	BD, the trial will have a c	ohort expansion

followed by a Phase 2 trial for a specific tumor.

in multiple eligible solid tumors to determine the

optimal biologic dose (OBD).

NASDAQ: KRBP

**Cohort Expansion** 

## **Clinical Programs**



NASDAQ: KRBP



# **Recent Developments**



## chPD1 Licensing

In-licensed chPD1 from Longwood University

CAR-T chPD1 addresses the Micro Tumor Environment in Solid Tumors



## AACR

6 posters presented at AACR 2021

Showcased Kiromic's -- Al Engine -- Al Target validation -- Gamma Delta T-cell Manufacturing



## **IND Submission**

Submitted two expanded INDs for Off-the-Shelf, Allogenic CAR-T for Solid Tumors

Targets: chPD1, lso-msln

Mode of Admin: IV

# **Recent Developments (Cont')**



### Manufacturing

GMP Manufacturing facility certification, expansion

Appointed manufacturing veterant from Novartis who had manufactured for Kymriah (CAR-T)



IQVIA engaged as CRO to:

- -- ID clinical sites
- -- Select patient pool
- -- Select PI at sites



### Presentations

- -- Gamma Delta T Summit to showcase latest off-the-shelf allogenic CAR-T manufacturing
- -- Next-Gen Immuno Oncology Annual Congress to showcase latest AI Targets and Validation
- -- Cell Immuno Therapies for Solid Tumors Summit



# **Milestones**

3Q-2021	cGMP, multiple-room facility certified, staffed, and stocked
3Q-2021	First-in-Human dosing with Off-the-Shelf, Alexis-Pro1 (chPD1) CAR-T for Solid Tumors
3Q-2021	First-in-Human dosing with Off-the-Shelf, Alexis-ISO1 (Iso-Mesothelin) CAR-T for Solid Tumors
4Q-2021	Early safety and efficacy data from Alexis-Pro1 (chPD1)
4Q-2021	Early safety and efficacy data from Alexis-Iso1 (Iso-Mesothelin)
2H-2022	Termination of Phase 1 (Data: Safety, Efficacy)
2Q-2023	Beginning of Phase 2 / 3



# **The Road Ahead**

## **GMP**

Redundancy

## **AI Algorithm**

Reinforced InSilico Al Yale Al Bioinformatics Clinical Taskforce

### CMC Taskforce

## Regulatory

Taskforce Phase 2 planning

## **Vertical** Integration

Key vendors are brought in-house to improve response time



## **Comparables**

A.I. Targets + Small Molec	cules	CAR-T and CAR	-NK
SCHRÖDINGER	<b>\$5.37 B</b> IPO Feb 2020	Frete	<b>\$7.61 B</b> Public (Phase 1, solids)
BLACK DIAMOND THERAPEUTICS	<b>\$440 M</b> IPO Jan 2020	CODIAK	<b>\$493 M</b> Public (IND)
therapeutics	<b>\$920 M</b> IPO Mar 2018	<b>Kite</b> Pharma	hematologic <b>\$11.9 B</b> GILD acqui. post approval
<b>Lantern</b> Pharma	<b>\$162 M</b> IPO Jun 2020	* JUCO THERAPEUTICS	hematologic <b>\$9.0 B</b> BMS acqui. end Phase 2

NASDAQ: KRBP Market data as of 06/22/2021, Yahoo Finance intraday



# Capitalization

	As of June 25, 2021	
Common Stock	7,387,500	
Options (\$8.90 WAEP)	558,435	
Warrants (\$15.00 per share)	62,500	
<b>Restricted Stock Units</b>	1,100,281	
Fully Diluted	9,108,716	

NASDAQ: KRBP

## Management



### CEO Director

#### Maurizio Chiriva-Internati, PhD

Mr. Chiriva-Internati is an associate professor at MD Anderson Cancer Center. He has spent the past 28 years studying cancer targets and is the founder of Kiromic Artificial Intelligence Neural Network. He has published +160 articles (+peer reviews) on cancer targeting and on the use of AI to expedite the search for these targets. He holds PhD in immunology (U of Nottingham), PhD in morphological science (Milan), and a Certificate in Artificial Intelligence - M.I.T.

### CFO Director

#### Tony Tontat

Mr. Tontat brings to Kiromic over 2 decades of business experience from public (NASDAQ: SRNE, NK) and privately held biotechs. He had been healthcare analysts at specialist healthcare investment funds in New York, and Connecticut. He was also an investment banker at HSBC Securities in their New York, London, and Paris offices. Bachelor of Arts in Economics - Harvard University.

### CSIO Director

#### Gianluca Rotino

Mr. Rotino held CEO and Chairman roles in several Italian companies specializing in high-tech, and corporate consulting. He also worked at law firms in Milan where he specialized in M&A, intellectual property prosecution and corporate law. He holds a business development degree and bachelor of science (electronics). Mr. Rotino also has earned a certificate in Essential Epidemiologic Tools for Public Heath Practice from Johns Hopkins University, a certificate in the Artificial Intelligence Programme from Said Business School at the University of Oxford, and a certificate in Leadership and Innovation from M.I.T.

#### CMO Scott Dalhbeck, MD, PharmD

Dr. Dalhbeck was a radiation oncologist and was an adjunct professor of internal medicine, pathology, and urology at Texas Tech. He has also patented, manufactured, and commercialized IP and has more than a decade of experience in medical and oncology commerce. He holds an MD - Texas Health Science Center, and a PharmD - U of Nebraska. His residency was at Kaiser Permanente of Los Angeles.

NASDAQ: KRBP

### **COMO** Ignacio Núñez MSCHE, MBB

Mr. Núñez has over 20 years of global experience in corporate functions including manufacturing, research, operational excellence and strategy. He has held senior leadership positions in companies including General Electric, Johnson & Johnson and Novartis. Most recently, he was the Executive Director of Manufacturing at the Gene Therapy Program of the University of Pennsylvania. Before that, he was the Head of Manufacturing Strategy and Operations Excellence at Novartis, where he was charged with transforming manufacturing operations in support of the ramp up of Kymriah, the first FDA-approved CAR-T cell therapy, which was developed at the University of Pennsylvania. Mr. Núñez holds an MSC in Chemical Engineering from the University of Granada.



Genome edit and delivery

Our single-cut gene edits carry a

lower mutagenesis risk vs. classic double-cut gene edits.

Our CAR receptors will also have

higher safety with an on-demand

cut-off switch vs. classic CAR

therapies with no off-switch.

Non-Viral



We are connecting the dots in cancer research by using AI and machine learning to connect silos of informations and arrive at cancer targets which will be more effective vs. classic development, saving man-years and billions in development dollars.

### dCA53 Grue Megrase ABBIE

# **Value Drivers**

Revolutionizing Next-Gen Allogenic CAR Therapy for Solid Tumors

Our CAR Therapy will be using off-the-shelf Gamma-Delta T-cells and will have a higher yield and significantly lower yield variability vs. classic CAR-T therapies.



Gamma Delta T-cell Immune Cell Type



Micro Tumor Environment Our CAR Therapies will be able to access the micro tumor environment due to our chPD-1 check-point activator vs. classic CAR-T therapies.

Classic CAR-T are limited to hematologic indications.





THANK YOU

7707 Fannin Street, Suite 140 Houston, Texas 77054 +1 (806) 368 - 6731

ttontat@kiromic.com



Step 2



24

# **Artificial Intelligence Engine**

+histo-aminochemistry filters, +machine learning





Step 4



**Target Validation** 

*We rigorously validate all targets from our A.I. Prediction Engine Internal validations and then external validation* 



### **Algorithm Validation**

InSilico Solutions



## Wet Lab Validation

### MD ANDERSON CANCER CENTER

Baylor University University of Rome Humanitas Research Hospital (Milan)





non-exhaustive list of functions being applied by A.I. Engine

### Prioritizing T and B Cell Targets

Diamond generates a prioritized list of cancer immunological targets for T cells and B cells.

These targets can be used to create therapies such as antibody therapies, T cell therapies, T cell receptor therapies, CAR T cell therapies and vaccine therapies.

### Identify Highly Expressed Genes

Diamond's cognitive and deep learning capabilities extract information from our extensive digital library consisting of clinical studies, genomic and proteomic datasets.

Diamond harmonizes all the raw data and creates datasets which allows us to screen for cancer targets.

Diamond will identify and prioritize lists of genes (biomarkers, wild type, mutant, isoform, neoepitope, etc.) that are highly and specifically expressed in the disease of interest while providing its distribution and methylation status across the entire patient population.

It also maps out the exact portion of the gene that will elicit an immune response.

### Perform Meta Analysis

Diamond performs meta-analysis and convolution studies while standardizing and normalizing data across multiple and variable experimental platforms, then allows for the visualization of consistent and accurate results in a user-friendly fashion.

### Predict Isoform Targets

Cancer cells will down regulate or shed targets in order to avoid detection and destruction by T cells (the immune system).

These variations are known as isoforms.

CancerSplice also shows a box plot by tissue of expression of the isoform in normal cancer genome atlas tissues and a box plot of the matching isoform in genotype-tissue expression program normal data.

The sequence of amino acids that are specific for the selected cancer isoforms are then directly fed to Diamond's artificial neural capsule network for peptide design and prioritization.





A key A.I. Engine

Target isoforms are protein variants of the same targets that occur during the normal processing of immature gene transcripts to the mature form.

Target isoforms include variations in their primary amino acid sequence that can change both the final folded form of the target plus their ability to be recognized by modified T cells (autologous/allogeneic) and other cells, such as NK or invariant NKT cells (often used in the allogeneic setting). If they are the predominate form on the cell surface, these isoforms can make it impossible for T cells to outright bind the targets on cancer cells.

No binding or insufficient binding to the isoform results in no killing of cancer cells.

Our CancerSplice accurately predicts the most appropriate isoforms for T cells to bind and destroy cancer cells.



## Allogenic Engineered Immune Therapy Step 03: Gamma-Delta T Cells expanded invitro



K



## How We Know: GD-T cell Expansion Works





A large fold of expansion of highly pure  $\gamma\delta$  T cells during in vitro stimulation, culture, isolation and expansion process.





### CONCLUSION

Our method of  $\gamma\delta$  T expansion yields up to1,000-fold expansion of  $\gamma\delta$  T cells, which is over 95% purity for positive for CD3, V $\gamma$ 9, and V $\delta$ 2.

This has potential to produce enough number  $\gamma\delta\,T$  for clinical use.



## **Up-Armoring**

Accessory proteins can "up-armor" cellular therapies

## Strategic choice of proteins to improve cellular function and neutralize anti-immune responses



- \* Activated signaling molecules chosen to enhance cell persistence by stimulating cytokine pathways
- \* Targeting the immunosuppressive "reactive" stroma can enable tumor targeting by therapeutic cells while increasing anti-tumor efficacy









### **ACTIVATION Switch**

A rapidly deployed activation switch can provide a survival and proliferation signal to the therapeutic cells to enhance their efficacy and persistence in vivo.

### **ATTENUATION Switch**

A rapidly deployed attenuation switch can intercept activation signals transiently to minimize toxicity following successful anti-tumor interactions. Choice of two non-mutually exclusive Attenuation Switch approaches:

- (a) a protein-based switch that rapidly triggers attenuation of target cells in a dose-dependent fashion.
- (b) a small molecule-based approach to rapidly and reversibly attenuate cell signaling.



### **SAFETY Switch**

A rapidly deployed, protein-based safety switch can eliminate therapeutic cells in case of acute toxicity. The safety switch is designed to eliminate either:

- (a) essentially all active therapeutic cells.
- (b) only the most active cells, preserving a cohort of backup therapeutic cells for long-term control of residual relapsing tumor cells.

The Safety Switch will be co-expressed along with the bioactive chimeric activation receptor (CAR), the Activation Switch, and the Attenuation Switch.