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**OUTPERFORM** 

Reason for report: FLASH NOTE

Michael Schmidt, Ph.D.

(617) 918-4588

Michael.Schmidt@Leerink.com

**Jonathan Chang, Ph.D.** (617) 918-4015

Jonathan.Chang@Leerink.com



#### HEALTHCARE EQUITY RESEARCH

## Kev Stats: (NASDAQ:KPTI)

S&P 600 Health Care Index:	1,279.01
Price:	\$16.91
52 Week High:	\$19.09
52 Week Low:	\$15.50
Shares Outstanding (mil):	28.7
Market Capitalization (mil):	\$485.3

## KARYOPHARM THERAPEUTICS, INC.

#### Selinexor Phase I Update at ASH Encouraging

- Bottom Line: KPTI provided an incremental update from its ongoing single-agent Phase I dose-escalation trial in hematological cancers at ASH this weekend. Recall, the last update on this trial included data through September 20. Presentations at ASH included data from additional patients across the Phase I study which we believe confirms the efficacy and tolerability profile of Selinexor seen previously in patients with Multiple Myeloma, AML, and NHL. KPTI also enrolled three additional patients with Richter's Syndrome, all of which had disease stabilization at the time of data analysis. We continue to believe Selinexor has clear single agent activity across a broad range of indications and a manageable tolerability profile. Reiterate Outperform.
- Multiple Myeloma (MM) data now available on 25 heavily pretreated patients. Disease control rate improved to 80% (PR+MR+SD) from 71% at the last update (see attached table). Duration of disease control was also prolonged. Five patients are remaining on the therapy for over five months and two patients for over one year without clinically significant cumulative toxicity.
- AML data further improved. In 33 patients evaluable for response, complete remissions with complete (CR, n=4) or incomplete (CRi, n=1) hematologic recovery were observed (CR+CRi 15%). This compares favorably to the prior update, which had 2 CRs and one CRi (see attached data table). Overall, the disease control rate improved by 14 percentage points to 61% from 47%. Five patients remained on study for at least three months, with three of them on study for at least five months with no cumulative toxicity.
- Non-Hodkin's Lymphoma (NHL) update includes now data on 11 DLBCL patients. Selinexor's disease control rate improved from 60% to 73% in DLBCL. KPTI now has 2 additional PRs, and 3 new patient with disease stabilization.
- **Next up**: KPTI plans to report more data from the Phase I trial and investigator-sponsored single agent and combination studies in the first half of 2014. KPTI also reiterate its plans to initiate two randomized pivotal studies in the first half of next year.
- Glossary: AML=Acute Myelogenous Leukemia, CR=Complete response, CRi=Complete Response with Incomplete Hematological Recovery, DLBCL=Diffuse Large B-Cell Lymphoma, MR=Minimal Response, NHL=Non-Hodgkins Lymphoma, PR=Partial Response, SD=Stable Disease

# Phase I Dose-Escalation (Hematological Ca

MM (Arm 1)	ASH 2013 Update				Septem 2013 U	•
	Pts	%	Pts	%		
Enrolled	25		20			
Evaluated	25		17			
PR	1	4%	1	6%		
MR	4	16%	4	24%		
SD	15	60%	7	41%		
PD	4	16%	4	24%		
WC	1	4%	1	6%		
NE	0	0%	0	0%		
PR+MR+SD	20	80%	12	71%		

WM (Arm 1)	ASH 2 Upda		Septem 2013 U	-
	Pts	%	Pts	%
Enrolled	3		2	
Evaluated	3		2	
PR	0	0	0	0%
MR	3	100%	2	100%
SD	0	0%	0	0%
PD	0	0%	0	0%
WC	0	0%	0	0%
NE	0	0%	0	0%

AML (Arm 2)	ASH 2		September 20 2013 Update		
	Pts	%	Pts	%	
Enrolled	38		34		
Evaluated	33		32		
CR	4	12%	2	6%	
CRi	1	3%	1	3%	
PR	3	9%	0	0%	
SD	12	36%	12	38%	
PD	11	33%	11	34%	
WC	4	12%	4	13%	
NE	2	6%	2	6%	
CR+CRi+PR+SD	20	61%	15	47%	

Source: Karyopharm data

**Arm1 B-Cell Cancers** 

September 20 Update	Evaluated	PR		SD		PD		wc	NE		PR+SE	)
CLL	4	2	50%	2	50%						4	100%
Richter's Syndrome	1	1	100%								1	100%
NHL	15	2	13%	8	53%						10	67%
DLBCL	5	1	20%	2	40%	2	40%				3	60%
MCL	3	1	33%	1	33%				1	33%	2	67%
FL	5			5	100%						5	100%
Transformed	2					2	100%				0	0%

ASH Update	Evaluated	PR		SD		PD		wc		NE		PR+SI	)
CLL	4	2	50%	2	50%							4	100%
Richter's Syndrome	4	1	25%	3	75%							4	100%
NHL	22	5	23%	10	45%							15	68%
DLBCL	11	3	27%	5	45%	2	18%			1	9%	8	73%
MCL	3	1	33%	1	33%					1	33%	2	67%
FL	6	1	17%	4	67%			1	17%			5	83%
Transformed	2					2	100%					0	0%

Source: Karyopharm Data

#### **Event**

#### Phase I program

Phase I data update

End of Phase I FDA/EMA meeting

Phase I dose expansion data

Phase I dose expansion data

Phase I dose expansion data

Phase I data

Phase I data

### **Hematological cancers**

Initiation of pivotal Phase II/III (single agent)

Initiation of pivotal Phase II/III (single agent)

Pivotal Phase II/III data

Pivotal Phase II/III data

Launch

Launch

## **Solid tumors**

Initiate Phase II

Initiate Phase II

Phase II data

Phase II data

### **Investigator-Initiated Studies**

Initiation Phase I/II

Source: Company filings and Leerink Swann estimates

#### Indication

Hematological cancers

Pivotal trial protocol review

Heme Arm 1 (MM, WM, DLBCL)

Heme Arm 2 (AML)

Solid tumors

Heme Arm 3 (TCL)

Food effect study in soft tissue/bone sacromas

elderly r/r AML

DLBCL or MM

elderly r/r AML

DLBCL or MM

r/r AML

DLBCL or MM

single agent solid tumor (gynecological)

single agent solid tumor (squamous cell cancers [lung, head and neck, esophageal)

single agent solid tumor (gynecological)

single agent solid tumor (squamous cell cancers [lung, head and neck, esophageal)

combination

<b>-11</b>
Timing
ASH Dec. 12, 2013
4Q13
mid-14
mid-14
mid-14
2014
2014
2Q14
2Q14
1H16
1H16
1H17
1H17
1Q14
1Q14
2H15
2H15
2013/14



# Disclosures Appendix Analyst Certification

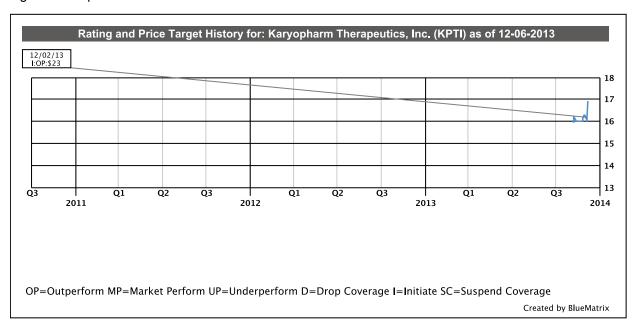
I, Michael Schmidt, Ph.D., certify that the views expressed in this report accurately reflect my views and that no part of my compensation was, is, or will be directly related to the specific recommendation or views contained in this report.

## **Valuation**

We estimate a \$23 fair value for KPTI shares in 12 months, based on a discounted cash flow (DCF) analysis. We apply a 12% discount rate to 35% probability of success-weighted Selinexor cash flows in three relapsed/refractory hematological cancer indications (AML, DLBCL, and MM). Potential Selinexor revenues derived from solid tumor indications as well as the preclinical and pet pipeline are upside to our valuation. Our valuation uses a terminal value derived by applying a 5x multiple to 2032 Selinexor revenue, discounted back by 18.25 periods. We believe use of a 5x revenue multiple is conservative compared to that of biotech industry average (6x). Based on our DCF analysis, we attribute \$20/share to Selinexor and the rest to the preclinical pipeline and platform and the remainder to expected cash in one year.

### **Risks to Valuation**

Early stage biotech companies such as KPTI face significant clinical and regulatory development risk, as well as commercial risks. KPTI also faces execution risk and financial risk. We estimate that KPTI's current cash will be sufficient to fund into early 2017, and the company may have additional financing needs before turning cash flow positive. The vast majority of our KPTI valuation is based on selinexor, the company's only clinical stage product candidate, so potential setbacks due to possible safety and/or efficacy related issues of selinexor could have a significant impact to our valuation.





	Distribution of Ratings/Investment Bank	ring Services (IE		erv./Past 12 Mos.
Rating	Count	Percent	Count	Percent
BUY [OP]	111	64.90	27	24.00
HOLD [MP]	60	35.10	0	0.00
SELL [UP]	0	0.00	0	0.00

# **Explanation of Ratings**

Outperform (Buy): We expect this stock to outperform its benchmark over the next 12 months.

<u>Market Perform (Hold/Neutral)</u>: We expect this stock to perform in line with its benchmark over the next 12 months.

<u>Underperform (Sell):</u> We expect this stock to underperform its benchmark over the next 12 months. The degree of outperformance or underperformance required to warrant an Outperform or an Underperform rating should be commensurate with the risk profile of the company.

For the purposes of these definitions the relevant benchmark will be the S&P 600® Health Care Index for issuers with a market capitalization of less than \$2 billion and the S&P 500® Health Care Index for issuers with a market capitalization over \$2 billion.



## **Important Disclosures**

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In the past 12 months, the Firm has received compensation for providing investment banking services to Karyopharm Therapeutics, Inc.

Leerink Swann LLC makes a market in Karyopharm Therapeutics, Inc.

Leerink Swann LLC has acted as the manager for a public offering of Karyopharm Therapeutics, Inc. in the past 12 months.

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Leerink Swann LLC Equity Research							
Director of Equity Research	John L. Sullivan, CFA	(617) 918-4875	john.sullivan@leerink.com				
Associate Director of Research	Alice C. Avanian, CFA	(617) 918-4544	alice.avanian@leerink.com				
Healthcare Strategy	John L. Sullivan, CFA	(617) 918-4875	john.sullivan@leerink.com				
	Alice C. Avanian, CFA	(617) 918-4544	alice.avanian@leerink.com				
Biotechnology	Howard Liang, Ph.D.	(617) 918-4857	howard.liang@leerink.com				
Biotechnology	Joseph P. Schwartz	(617) 918-4575	joseph.schwartz@leerink.com				
	Marko Kozul, M.D.	(415) 905-7221	marko.kozul@leerink.com				
	Michael Schmidt, Ph.D.	(617) 918-4588	michael.schmidt@leerink.com				
	Irene Lau	` '	irene.lau@leerink.com				
	Gena Wang, Ph.D., CFA	(415) 905-7256 (212) 277-6073	gena.wang@leerink.com				
	Paul Matteis	(617) 918-4585	paul.matteis@leerink.com				
	. da matoro	(0.1.) 0.10 1000	paammanoid Sissimmosiin				
Life Science Tools	Dan Leonard	(212) 277-6116	dan.leonard@leerink.com				
and Diagnostics	Justin Bowers, CFA	(212) 277-6066	justin.bowers@leerink.com				
Pharmaceuticals/Major	Seamus Fernandez	(617) 918-4011	seamus.fernandez@leerink.com				
	Ario Arabi	(617) 918-4568	ario.arabi@leerink.com				
Specialty Pharmaceuticals,	Jason M. Gerberry, JD	(617) 918-4549	jason.gerberry@leerink.com				
Generics	Christopher W. Kuehnle, JD	(617) 918-4851	chris.kuehnle@leerink.com				
Medical Devices, Cardiology &	Danielle Antalffy	(212) 277-6044	danielle.antalffy@leerink.com				
Orthopedics	Richard Newitter	(212) 277-6088	richard.newitter@leerink.com				
	Robert Marcus, CFA	(212) 277-6084	robert.marcus@leerink.com				
	Ravi Misra	(212) 277-6049	ravi.misra@leerink.com				
Healthcare Services	Ana Gupte, Ph.D.	(212) 277-6040	ana.gupte@leerink.com				
	7a • ap.o., 12	(212) 277 0010	and gapto chommicom				
Healthcare Technology	David Larsen, CFA	(617) 918-4502	david.larsen@leerink.com				
& Distribution	Christopher Abbott	(617) 918-4010	chris.abbott@leerink.com				
Sr. Editor/Supervisory Analyst	Mary Ellen Eagan, CFA	(617) 918-4837	maryellen.eagan@leerink.com				
Supervisory Analysts	Robert Egan		bob.egan@leerink.com				
	Amy N. Sonne		amy.sonne@leerink.com				

**New York** 299 Park Avenue, 21<sup>st</sup> floor New York, NY 10171 (888) 778-1653 Boston One Federal Street, 37<sup>th</sup> Floor Boston, MA 02110 (800) 808-7525

San Francisco 201 Spear Street, 16<sup>th</sup> Floor San Francisco, CA 94105 (800) 778-1164