August 29, 2015



Tags: Biotech, Investment, Drug development

RA Capital: Flexibility builds innovation



Peter Kolchinsky, Managing Director, RA Capital Management

Flexibility is the watchword for investment at RA Capital Management. As Peter Kolchinsky, managing director, explained, "Investors should be as flexible as possible. Therefore, we're a crossover fund manager. We invest in biotechnology companies whether they are public or private companies and, increasingly, are putting our efforts into company formation."

RA Capital Management chooses not to limit itself to early stage, late stage or other categories. Instead, he said, "We want the flexibility to invest in any way that is appropriate for a given concept. If, for example, a company already is public and is the best at what it does, we want the flexibility to invest." Conventional fund structures could restrict that option.

By eliminating such restrictions, the company retains many ways to invest. They may include direct investments, private placements, stock purchases on the open market or follow-on financings, for example. Additionally, it also has the flexibility to develop a company around a compelling idea.

"This flexibility creates a continuum, so we can approach a company at any point of its lifecycle. If we focused only on venture capital, for example, we would cut ourselves off from some great public companies," Kolchinsky said.

We originate conviction

That flexibility comes with a clear investment philosophy. Chief among its rather pithy values: "We originate conviction. We do not borrow it from others."

This means, Kolchinsky said, "that when presented with data we find compelling, we are willing to be the first to invest. If everyone waits for someone else to be first, who will capitalize the idea? Successful entrepreneurs by definition originate conviction in an idea and in themselves. They typically need at least one investor to appreciate the merits of their idea. We try to be that investor."

Formed by managers who all began their careers as scientists, RA Capital Management has the courage of its convictions. "You have to speak the language of science and parse data to invest well in life sciences," Kolchinsky said.

"We try to be as proficient in healthcare and in the life sciences as possible, to originate conviction based on our analysis of the data and assessment of professional competency of the management team." Once convinced, RA Capital can catalyze the formation of an investor syndicate.

Not all investment firms have deep scientific expertise and such confidence. As he said, "Some investors are confident they know the science and can distinguish between good and bad ideas and approaches. Others are clear that they will only invest once there is a lead investor. Many vary depending on the deal. An investor who leads an investment in one area he or she knows well, maybe cancer, might be less confident in another, like gene therapy, and only join after someone more specialized has validated the idea."

Stock fluctuations

To highlight the implications of adhering strictly to trend reports, he cited stock sell-offs. "When a stock drops, some people become desperate to sell it. They are losing confidence in that stock because other people are selling it. Prices dip lower, yet often the fundamentals of the company haven't changed."

When that happens, he said, ask why the stock price dropped and whether the fundamentals are changing or just perception. "Science moves slowly relative to the ups and downs of financial markets," Kolchinsky pointed out.

"Among investors who don't understand the data well enough to say the crowd is wrong, a declining stock price is a powerful motivation to sell," he said. "Investors who, instead, anchor themselves to the fundamental value of the company will buy that stock and halt the fall. At the nadir is an investor who originates conviction," and by that conviction catalyzes renewed enthusiasm among investors. Kolchinsky said RA Capital is always striving to be that that key investor.

Kolchinsky advises companies to take a long view of the stock market. "If you look at the past five or ten years, the recent fluctuations are merely a blip. If a company is well-capitalized, it can still move full speed ahead without changing its plans. If, however, a company has only one year of cash left, the near-term stock market conditions may affect its ability to raise funds." The implications of stock fluctuations, therefore, are relative.

Data-driven decisions

To build conviction, RA Capital focuses on evidence-based investing. Kolchinsky likens this to physicians who make decisions based on data rather than relying solely on their own personal experiences. In medicine, he says, evidence-based decisions ensure that treatments are chosen because they have been proven to work in trials, not because of anecdotal reports that a few patients improved. Like savvy physicians, "We're driven by the data."

RA Capital Management's employees, in identifying the company's pervading values, said, "We address the elephant in the room," which is to say that they do not hesitate to address their concerns so that they can be discussed and hopefully resolved. "Some people play their cards close. We believe in letting companies and each other see our thought process," Kolchinsky added. He expects the management team at companies of interest to do the same.

For example, when assessing the management teams of potential investments, "We look for earnestness and competence. We don't want embellishment. If there's a problem with a program – and they all have some problem – lay it out for us and show us several strategies to resolve it. Don't make us dig for it." If investors have to dig to find something, he says, it makes them wonder what else may be hidden.

Individual merits

In evaluating companies in which to invest, RA Capital looks at each option as a special case. Within the life sciences, "we have no hard and fast rules regarding what we consider as potential investments," Kolchinsky said.

"We believe in maximal flexibility and being prepared. We generate good returns on investment by finding underappreciated, innovative companies in our space. You never know where or when the next breakthrough will emerge. We look at public and private companies in medical devices, drugs, diagnostics and other areas, though typically drugs offer the most upside."

As part of its research, RA Capital maps drugs and technologies within therapeutic areas to understand the competitive landscape for treating each disease, including the drugs and the companies developing them. For example, he said, "dozens of companies are targeting lung cancer with EGFR, ALK, and other kinase inhibitors, PD-1, various kinds of vaccines, and chemotherapy. How do these technologies relate to one another and combine to give a patient the most benefit? Which companies have all the best elements so as to both beat the disease and beat other companies?

"Think of our maps as chessboard diagrams, showing individual drugs as the pieces. In this analogy, the maps facilitate collaborations between the bishop and knight, for example," he said. In practice, these maps help identify

promising companies for investment, as well as potential synergies among compounds or companies that may lead to innovative collaborations or acquisitions.

"We also hope to employ this approach in a collaboration with a leading contract research organization (CRO) to design some of the most sophisticated oncology trials ever conducted. It's in the vein of the earlier I-SPY trials and involves pooling drugs into one clinical trial, enabling patients to benefit from the therapies that are best for them," Kolchinsky said.

To further catalyze innovation, RA Capital also champions its Executive-in-Residence (EIR) program in which "people who have a background of forming or managing companies join our company, see the industry and hundreds of companies from our perspective, evaluate available technologies, and use these insights to inform their next career moves," Kolchinsky said.

"We want to see how someone we respect with real industry experience, when empowered by our platform, chooses from among all the options to invest their precious time and reputations, because maybe our capital should follow." For the EIR, it's an opportunity to see how investing is done from the inside as they consider forming new companies or joining existing organizations.

Focus on solutions

"Our philosophy focuses on solution development. A company should define itself by the problem it aims to solve, not the drug it hopes to develop," he said.

Hepatitis C therapeutics are one example. In 2014, three years after commercialization, Vertex took its first-in-class protease inhibitor Incivek™ (teleprevir) off the market, ceding that space to Gilead and AbbVie's better combination therapeutics. "Patients didn't need a protease inhibitor. They needed a cure."

Vertex, in the meantime, shifted its focus to cystic fibrosis, developing the first potentiator and corrector, Kolchinsky said. "It combined them into one regimen that did more than either the potentiator or corrector could do alone. Many of its competitors have only one or the other."

RA Capital Management's flexible approach to investing highlights the best in public and private companies, large and small. It is designed to bring bright people together to innovate in ways that would be unlikely or even impossible alone. There is no limitation on innovation, so why should there be limitations on investing?

#

Reprinted from partneringNEWS™, an online journal for executives in the life sciences focused on the people behind the deals.

RACAPITA L RA CAPITAL MANAGEMENT, L.P.

General Disclaimers

The information contained herein (the "Materials") is provided for informational and discussion purposes only and contains statements of opinion and belief. The Materials are not, and may not be relied on in any manner as, legal, tax, or investment advice. The Materials do not constitute an offer to sell, a solicitation to buy, or a recommendation for any security, nor do they constitute an offer to provide investment advisory or other services by RA Capital Management, L.P. and its affiliates and/or any investment products it advises (collectively, "RA Capital" or the "Firm"). Each recipient should make its own investigations and evaluations of RA Capital, and any investment products it advises, and should consult its own attorney, business adviser, and tax adviser as to legal, business, tax, and related matters thereto. The information contained in the Materials is not intended to be, and should not be viewed as, "investment advice" within the meaning of 29 C.F.R. §2510.3-21 or otherwise.

Any views expressed herein, unless otherwise indicated, are those of RA Capital as of the date indicated, are based on information available to RA Capital as of such date, and are subject to change, without notice, based on market and other conditions. No representation is made or assurance given that such views are correct and such views may have become unreliable for various reasons, including changes in market conditions or economic circumstances. Such views may have been formed based upon information, believed to be reliable, that was available at the time the Materials were published. Certain information contained herein concerning economic trends and/or data may be based on or derived from information provided by independent third-party sources. RA Capital believes that the sources from which such information has been obtained are reliable; however, it cannot guarantee the accuracy of such information and has not independently verified the accuracy or completeness of such information or the assumptions on which such information is based. RA Capital has no duty or obligation to update the information contained herein.

The content of the Materials neither constitutes investment advice nor offers any opinion with respect to the suitability of any security. Any references, either general or specific, to securities and/or issuers are for illustrative purposes only and are not intended to be, and should not be interpreted as, advice or recommendations to purchase, continue to hold, or sell such securities, or as an endorsement of any security or company. Certain current and prior investments may be highlighted in order to provide additional information regarding RA Capital's investment strategy, the types of investments it pursues, and current or anticipated exit strategies. In addition, due to confidentiality restrictions, the information contained herein might not reference investments in certain companies. Accounts managed by RA Capital may invest in certain companies referenced in the Materials; however, RA Capital makes no guarantees as to accuracy or completeness of views expressed in the Materials. Any strategies and companies referenced in the Materials may not be suitable for all investors

As stated above, the Materials are not an offer or solicitation for the purchase or sale of any security, including any interest in RA Capital Healthcare Fund, L.P. (the "Master Fund") or RA Capital Healthcare International Fund Ltd. (the "Offshore Fund," and, collectively with the Master Fund, the "Fund"), and should not be construed as such. Such an offer will only be made by means of a confidential Private Placement Memorandum (the "PPM") to be furnished to qualified investors upon request. The information contained herein is qualified in its entirety by reference to the PPM, which contains additional information about the investment objective, terms, and conditions of an investment in the Fund, and also contains certain disclosures that are important to consider when making an investment decision regarding the Fund. In the case of any inconsistency between any information contained herein or in the Materials and the PPM, the terms of the PPM shall control.

The Materials are proprietary and confidential and may include commercially sensitive information. As such, the Materials must be kept strictly confidential and may not be copied or used for an improper purpose, reproduced, republished, or posted in whole or in part, in any form, without the prior written consent of RA Capital. The recipient of the Materials must not make any communication regarding the information

contained herein, including disclosing that the Materials have been provided to such recipient, to any person other than its authorized representatives assisting in considering the information contained herein. Each recipient agrees to the foregoing and to return (or destroy upon RA Capital's instructions) the Materials promptly upon request.

Any investment strategies discussed herein are speculative and involve a high degree of risk, including loss of capital. Investments in any products described herein and the Fund's performance can be volatile, and investors should have the financial ability and be willing to accept such risks. An investor could lose all or a substantial amount of his or her investment. The Fund may be leveraged. Interests in the Fund are illiquid, as there is no secondary market for the Fund interests, and none is expected to develop. The Fund interests are subject to restrictions on transfer. Prior to investing in the Fund, investors should read the PPM and pay particular attention to the risk factors contained therein. Fees and expenses charged in connection with an investment in the Fund may be higher than the fees and expenses of other investment alternatives and may offset investment profits of the Fund. RA Capital has total trading authority over the Fund. The use of a single advisor applying generally similar trading programs could mean lack of diversification and, consequentially, higher risk. A portion of the trades executed for the Fund may take place on foreign exchanges. It should not be assumed, and no representation is made, that past investment performance is reflective of future results. Nothing herein should be deemed to be a prediction or projection of future performance. To the extent any prior or existing investments are described, RA Capital makes no representations, and it should not be assumed, that past investment selection is necessarily reflective of future investment selection, that any performance discussed herein will be achieved or that similar investment opportunities will be available in the future or, if made, will achieve similar results.

In particular, to the extent valuation information is provided for any unrealized investments, such valuations are RA Capital's estimates as of the date set forth in the Materials, and there can be no assurance that unrealized investments will be realized at such valuations. While RA Capital believes any valuations presented herein are reasonable, such valuations may be highly subjective, particularly for private investments, and are based on information provided by third parties and/or RA Capital's assumptions, any or all of which might be mistaken or incomplete. Actual realized returns will depend on, among other factors, future operating results, the value of the assets and market conditions at the time of disposition, any related transaction costs, and the timing and manner or sale, all of which may differ from the assumptions on which the valuations contained herein are based. As a result of the foregoing, actual realized returns may differ materially from the valuations contained herein.

Certain information contained in this document constitutes "forward-looking statements," which can be identified by the use of forward-looking terminology such as "may," "will," "should," "expect," "anticipate," "target," "project," "estimate," "intend," "continue," or "believe," or the negatives thereof or other variations thereon or comparable terminology. Due to various risks and uncertainties, actual events or results or the actual performance of any investment may differ from those reflected or contemplated in such forward-looking statements. Prospective investors should not rely on these forward-looking statements when making an investment decision.

None of the information contained herein has been filed with the U.S. Securities and Exchange Commission, any securities administrator under any securities laws of any U.S. or non-U.S. jurisdiction, or any other U.S. or non-U.S. governmental or self-regulatory authority. No such governmental or self-regulatory authority will pass on the merits of any offering of interests by RA Capital or the adequacy of the information contained herein. Any representation to the contrary is unlawful. The interests in the Fund have not been, and will not be, registered under the U.S. Securities Act of 1933, as amended, or qualified or registered under any applicable state, local, provincial, or other statutes, rules, or regulations. The Fund has not been, and will not be, registered as an investment company under the U.S. Investment Company Act of 1940, as amended.