Equity Research

Flash Comment



June 24, 2015 CALA-NASDAQ--Outperform (1) / V

Biotechnology

CALA: Arginase Inhibitor CB-1158 Puts CALA On Track To Enter Immunooncology By Q2 2016+

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Sector Rating: Biotechnology, Market Weight

**On 6/23/15 CALA announced it selected CB-1158 as its lead clinical candidate for its arginase inhibitor program. CALA has had a dedicated internal effort targeting arginase inhibitors and recently augmented the portfolio with its December 2014 Mars Corporation's Symbioscience Division license.

**Recall within the cancer cell microenvironment myeloid-derived suppressor cells (immune cells like T-regulatory cells) secrete the arginase enzyme which depletes the semi-essential amino acid arginine, important for activation of cytotoxic T-cell and natural killer-cells' activities against tumor cells. Proof-of-concept is being validated clinically by indoleamine 2,3-dioxygenase inhibitors (IDO) which look to block depletion of tryptophan. Arginase inhibitors have an identical mechanism to IDO inhibitors as they allow for the CD3 zeta chain for the T-cell co-receptor to reach the T-cell surface so it can begin signaling activities.

**Investigational new drug application-enabling studies are underway with a goal of filing an INDA by late Q1 2016.

**Similar to IDO inhibitors where synergies are expected with programmed death-1/programmed death ligand-1 (PD-1/PD-L1) inhibitors, CALA believes CB-1158 may also be synergistic with PD-1/PD-L1 antibodies. We note that based on a 2013 Immunity Review paper by Mellman and Chen, multiple other combination approaches (for CB-1158) may be feasible, including with cancer vaccines (which target increase in CD4/CD8 T-cells), Chimeric Antigen Receptor and T-cell receptor agents, and Yervoy to name a few. As a result, following Ph. I proof-of-concept for CB-1158 it would not be a surprise to see CALA enter non-exclusive/exclusive development agreements or initiate CB-1158+PD-1/PD-L1 combination studies independently.

**We believe a number of preclinical data could be presented at the American Association for Cancer Research-"Triple Meeting" in November 2015, along with updated Ph. I single-agent cohort expansion results for CB-839.

**BOTTOM LINE: While not the key near-term value driver, CB-1158 provides a potential additional long-term shot on goal in the important area of immuno-oncology. Following positive CB-839 updates in June and ahead of important data readouts later in 2015, we continue to see CALA shares as under-valued.

Calithera Biosciences, Inc. (CALA-NASDAQ)

Price as of 6/23/2015: \$8.84

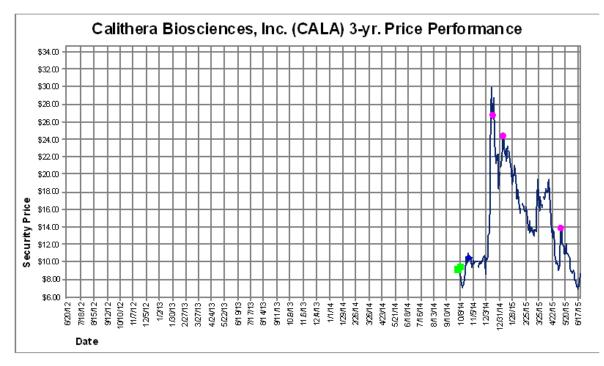
FY 15 EPS: \$-2.02 FY 16 EPS: \$-2.32 Shares Out.: 17.9 MM Market Cap.: \$158.24 MM

Please see Disclosure Appendix for rating definitions, important disclosures, and required analyst certifications.

All estimates/forecasts are as of 06/24/15 unless otherwise stated.

DISCLOSURE APPENDIX

Required Disclosures



	D ate	Publication Price (\$)	Rating Code	Val. Rng. Low	Val. Rng. High	Close Price (\$)
	10/3/2014	IPO at \$10.00				
	10/27 /2014		Andrews			
•	10/27 /2014	11.04	1	19.00	20.00	10.40
•	12/15/2014	29.85	1	50 0 0	55.00	26.75
•	1/7/2015	23.85	1	35 0 0	40.00	24.40
•	5/8/2015	9.52	1	27 0 0	30.00	13.87

Source: Wells Fargo Securities, LLC estimates and Reuters data

Symbol Key

- Ratin g Downgrad e
- 📤 Rating Upgrade
- Valuation Range Change

Initiation, Resumption, Drop or Suspend Analyst Change

🕽 Split Adjustment

Rating Code Key

1 Outperform/Buy SR Suspended 2 Market Perform/Hold NR Not Rated 3 Underperform/Sell NE No Estimate

Additional Information Available Upon Request

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CALA: Key risks include clinical trial failure, a safety signal for CB-839, and financing risk.

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SECTOR RATING

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- M = Market Weight: Industry expected to perform in-line with the relevant broad market benchmark over the next 12 months.
- **U = Underweight:** Industry expected to underperform the relevant broad market benchmark over the next 12 months.

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As of: June 24, 2015

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