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Reason for report:

**COMPANY UPDATE** 

### ZAFGEN, INC.

Mtg With Management Provides Insight Into Ph. III PWS Changes

- Bottom Line: After meeting with ZFGN on our bus tour we have a greater appreciation for the factors that led to changes in the design and stat. analysis for the Ph. III trial of beloranib in Prader-Willi Syndrome (PWS). We are lowering our prob.of success to 70% from 80% because we believe that the new conditions make it slightly harder for Ph.III to succeed although we believe even in less bullish scenarios that the totality of the Ph. III PWS data in early 1Q16 as well as Ph. II severe obesity data in late Q4/early 1Q16 could still support approval in this rare disease with sig. unmet need. We believe that ZFGN's business model and suite of metAP2 inhibitors are very differentiated from other obesity stories and that recent stock weakness presents an attractive buying opportunity. Reiterate OP, lowering PT to \$56 from \$61.
- As shown in our slides within, we believe the company has managed to evolve their strategy according to personnel changes at the FDA in a way that still supports a good likelihood of success in Ph III. Although the new reviewer at the FDA prefers body weight to body mass (DEXA), and views both body weight and hyperphagia behavior as equally important to hit, our analysis of Ph. II data suggests that this outcome is also achievable. Multiple studies of beloranib have demonstrated weight loss across several types of obesity which builds over time. Interestingly, the Street appears to be more comfortable with the ability to succeed on the weight loss co-primary endpoint, however we believe that the hyperphagia effects are actually more impressive and easier to hit, particularly in PWS.
- Ph. II data suggests that co-primary endpoints are still within reach in the Ph. III bestPWS study. For the first of the two primary endpoints, 1.2mg and 1.8mg doses of beloranib showed ~1.5% weight loss vs. placebo in Ph2, which was limited by short duration (4wks), s mall numbers (~5 pts/arm), one placebo patient losing a lot of weight and pts. not complying with specific weighing procedures (fasting, voiding and gowning). We are encouraged that Ph. III will employ higher dosing (1.8mg and 2.4mg), longer duration (6 mos), larger numbers (~35 pts/arm), and procedural improvements. On the second of the two primary endpoints (hyperphagia), beloranib actually showed over a 50% improvement vs. 40% worsening for placebo (delta of ~36 pts).
- Mathematically, powering in Ph.III is unchanged at 90% to show at least a 1.5% improvement in weight and a 4.5 point improvement in hyperphagia; although ZFGN must now hit both endpoints, the hurdle has been lowered to p<0.05 on both from p<.025 on either one previously. Even if beloranib were to miss one of the co-primary endpoints, we believe that the risk/benefit could be positive for the FDA to approve the drug considering the safety of the drug (fewer dropouts than expected and only using the higher dose in the EU Ph. III bodes well in our view), strong presence of the patient pop. at the FDA, and high unmet medical need in this debilitating orphan disease.

Key Stats: (OTC Un:ZFGN)

 S&P 600 Health Care Index:
 1,625.40

 Price:
 \$30.28

 Price Target:
 \$56.00 from \$61.00

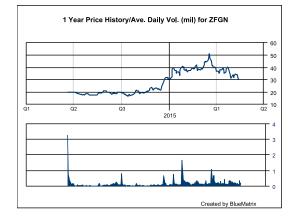
Methodology:

Probability-weighted DCF analysis, 11% discount rate

52 Week High: \$55.36 52 Week Low: \$16.01 Shares Outstanding (mil): 29.6 Market Capitalization (mil): \$896.3 Book Value/Share: \$0.00 Cash Per Share: \$7.50 Dividend (ann): \$0.00 Dividend Yield: 0.0%

Shares Outstanding (mil): Diluted; includes stock

Cash Per Share: On a net basis as of 1Q15E



| Dec Yr | 1Q   | 2Q  | 3Q  | 4Q  | FY Rev | 1Q        | 2Q       | 3Q       | 4Q       | FY EPS   | P/E |
|--------|------|-----|-----|-----|--------|-----------|----------|----------|----------|----------|-----|
| 2014A  | 0.0  | 0.0 | 0.0 | 0.0 | 0.0    | (\$6.18)  | (\$2.96) | (\$0.65) | (\$0.48) | (\$3.00) | NM  |
| 2015E  | 0.0A | 0.0 | 0.0 | 0.0 | 0.0    | (\$0.53)A | (\$0.81) | (\$0.89) | (\$0.99) | (\$3.23) | NM  |
| 2016E  |      |     |     |     | 0.0    | j         |          |          |          | (\$3.13) | NM  |

Source: Company Information and Leerink Partners LLC Research

GAAP EPS presented



### **INVESTMENT THESIS**

We rate ZFGN Shares Outperform. Zafgen (NASDAQ: ZFGN) is a biopharmaceutical company dedicated to the development of medicines to address the unmet need in obesity, with an initial focus on two orphan diseases that offer a potentially streamlined development path and high margin business model. We believe that ZFGN has an experienced management team with an outstanding track record executing in the field of drug discovery and commercialization. ZFGN's lead asset, beloranib, is a MetAP2 inhibitor that has generated compelling Phase II data in Prader-Willi syndrome (PWS) and severe obesity on a number of clinically relevant endpoints, in our view, and is now being studied in a registration-enabling Phase III PWS study expected to read out data by 1Q16. We project a 70% probability of PWS approval in 2017, and peak gross PWS sales of ~\$700MM worldwide in 2029. Zafgen is also developing beloranib in hypothalamicinjury associated obesity (HIAO), where it also recently released positive proof-of-concept Phase Ila data and expects to initiate a registration-enabling study in HIAO in the near future. We project 60% probability of HIAO approval in 2018, and peak gross HIAO sales of ~\$445MM worldwide in 2029. The commercial opportunity presented by severe obesity holds the potential to be orders of magnitude larger than PWS and HIAO, though ZFGN will likely need support from a larger partner to unlock its full potential. Thus, while severe obesity afflicts ~16MM Americans in the US, we only model ~\$200MM in peak beloranib sales in non-PWS/HIAO patients, though in a partnership/acquisition (P&A) scenario, ZFGN is likely to receive considerably more value for beloranib in high prevalence indications. In the meantime, we believe that establishing broader proof-of-concept in orphan sub-populations offers a less risky and more rapid development path, and over the long term could position ZFGN as a very attractive partnership or takeout target. Likewise, a second generation MetAP2 inhibitor in preclinical development for general obesity, and a novel chemical class MetAP2 inhibitor in preclinical development for NASH/diabetes could provide significant upside to our price target as their development advances further.

### **VALUATION**

We derive a ~\$56 per share value for ZFGN using an 11% discount rate and a 2% terminal growth rate. Our price target assumes 70% and a 60% probabilities of beloranib approval in PWS and HIAO, respectively, which leads to our risk adjusted peak sales estimates of ~\$485MM and ~\$270MM in each indication. We only model ~\$200MM in peak sales in severe obesity, which we believe holds the potential to be very conservative if/when ZFGN generates pivotal beloranib data in orphan indications.

### **RISKS TO VALUATION**

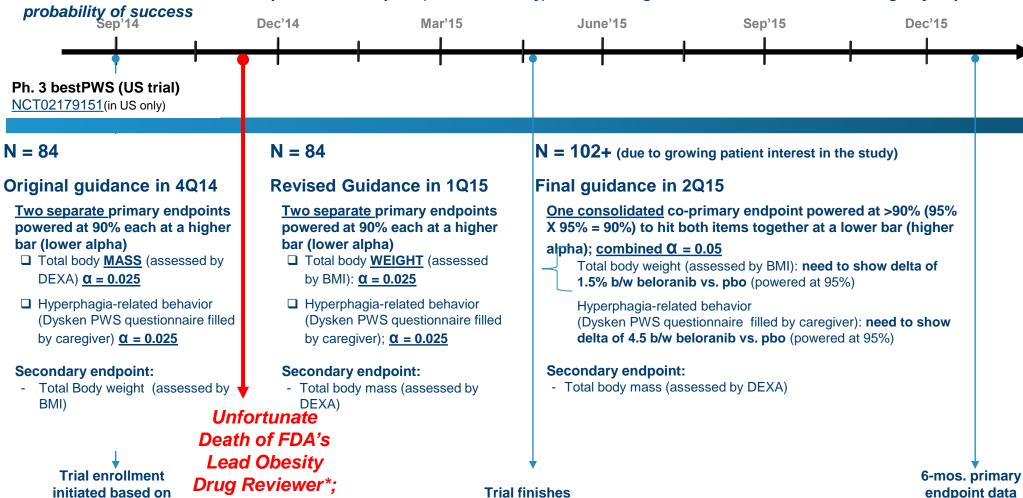
Risks to our valuation include disappointing clinical data, regulatory setbacks, dilution risk from an additional equity offering, and commercial shortfalls. Because ZFGN has only one late stage product, the occurrence of any of these could impact the stock significantly.



# Beloranib Prader-Willi Ph. 3 Clinical Development

Bull Case of Launch in 2H16 Based on US Trial; Base Case of Launch in 2H17 After EU Trial Readout

The recent revision to statistical plan has no impact (mathematically) on trial design and hence, shouldn't meaningfully impact



enrollment

Eric Coleman, Ex-Deputy Director, Division of Metabolism and Endocrinology, FDA

**Review Process** 

Restarted

Source: Leerink Partners Research, Company Reports, SEC Filings

original guidance

Ph. 3 bestPWS|EU (EU trial)

(in EU sites only)

N = 150

Same primary endpoint as bestPWS US study

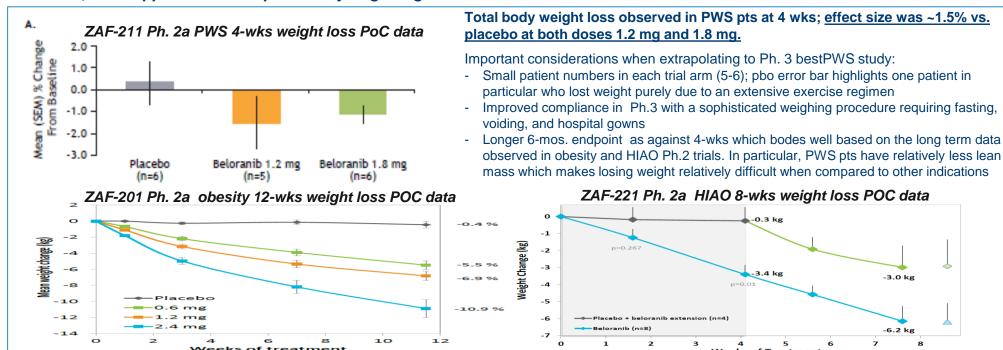
expected in 1Q16

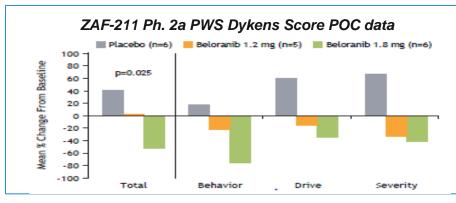


# Beloranib Prader-Willi Ph. III Clinical Development

How Much Should We Extrapolate From Ph. I/II Trials?

4-wks PoC data in PWS, 8-wks PoC in HIAO and 12-wks data in severe obesity renders new 6-mos. co-primary endpoint bar achievable, and supports our 80% probability of getting to market





Improvement in Hyperphagia-related behavior in PWS pts at 4 wks; <u>effect size</u> SIGNIFICANTLY higher than 4.5 units at both doses 1.2 mg and 1.8 mg.

Important considerations when extrapolating to Ph. 3 bestPWS study:

- 40-points score of 10 questions changed to a 36-points score with exclusion of 1 question that was indicator of the caregiver burden (as against patient burden), based on FDA's feedback
- An impressive ~20 points differential for 1.8mg beloranib in Ph.2 has more than
  enough room we believe to work around investors' concerns over this endpoint. We
  believe this endpoint should be relatively easier to hit the ~4.5 points delta, as
  requested by FDA

In worst case of beloranib missing the primary endpoint, pre-specified stratification may facilitate looks at the two endpoints in isolation, which if stat. sig. with a clinically meaningful benefit and when coupled with a benign safety profile may still justify approval in this debilitating orphan disease with a high unmet need for a therapeutic.

| ZFGN P&L (\$MM) GAAP         | 2013       | 1Q14         | 2Q14      | 3Q14      | 4Q14   | 2014      | 1Q15      | 2Q15E     | 3Q15E     | 4Q15E     | 2015E     | 2016E     | 2017E     | 2018E     | 2019 | ЭE   |
|------------------------------|------------|--------------|-----------|-----------|--------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|------|------|
|                              |            |              |           |           |        |           |           |           |           |           |           |           |           |           |      |      |
| Beloranib PWS                | -          | -            | -         | -         | -      | -         | -         | -         | -         | -         | -         | -         | 4.0       | 46.5      |      | 26.6 |
| Beloranib HIAO               | -          | -            | -         | -         | -      | -         | -         | -         | -         | -         | -         | -         | -         | 10.0      | 3    | 34.4 |
| Beloranib Severe Obesity     | -          | -            | -         | -         | -      | -         | -         | -         | -         | -         | -         | -         | -         | -         |      | -    |
| Total Revenue (p/w)          | -          | -            | -         | -         | -      | -         | -         | -         | -         | -         | -         | -         | 4.0       | 56.5      | 16   | 51.0 |
| cogs                         | -          | -            | -         | -         | -      | _         | -         | _         | -         | _         | _         | -         | 0.4       | 5.7       | 1    | 16.1 |
| R&D                          | 9.6        | 3.3          | 4.7       | 12.1      | 7.3    | 27.4      | 10.2      | 18.0      | 20.0      | 22.0      | 70.2      | 70.2      | 63.2      | 56.9      | 5    | 6.4  |
| SG&A                         | 4.2        | 1.2          | 1.3       | 2.3       | 3.3    | 8.1       | 3.0       | 4.0       | 4.3       | 5.0       | 16.3      | 22.9      | 30.0      | 33.9      | 4    | 18.3 |
| Operating Expenses           | 13.8       | 4.5          | 6.0       | 14.4      | 10.7   | 35.5      | 13.2      | 22.0      | 24.3      | 27.0      | 86.5      | 93.1      | 93.6      | 96.4      | 12   | 20.8 |
| Operating Income             | (13.8)     | (4.5)        | (6.0)     | (14.4)    | (10.7) | (35.5)    | (13.2)    | (22.0)    | (24.3)    | (27.0)    | (86.5)    | (93.1)    | (89.6)    | (39.9)    | 4    | 10.3 |
| Interest Income (Expense)    | -          | (0.0)        | (0.4)     | (0.2)     | (0.2)  | (0.8)     | (0.2)     | (0.2)     | (0.2)     | (0.2)     | (0.8)     | (0.4)     | (0.3)     | -         |      | -    |
| FX Gains/Losses              | (0.2)      | 0.1          | 0.0       | (0.1)     | (0.1)  | (0.1)     | (0.1)     | -         | -         | -         | (0.1)     | -         | -         | -         |      | -    |
| Total Other Income (expense) | (0.2)      | 0.1          | (0.4)     | (0.3)     | (0.3)  | (0.9)     | (0.2)     | -         | -         | -         | (0.2)     | (0.4)     | (0.3)     | -         |      | -    |
| ЕВТ                          | (14.0)     | (4.5)        | (6.4)     | (14.7)    | (10.9) | (36.5)    | (13.5)    | (22.0)    | (24.3)    | (27.0)    | (86.8)    | (93.4)    | (89.8)    | (39.9)    | 4    | 10.3 |
| Тах                          | -          | -            | -         | -         | -      | -         | -         | -         | -         | -         | -         | -         | -         | -         |      |      |
| Net Income (Loss)            | (14.2)     | (4.5)        | (6.4)     | (14.7)    | (10.9) | (36.6)    | (13.5)    | (22.0)    | (24.3)    | (27.0)    | (86.8)    | (93.4)    | (89.8)    | (39.9)    | 4    | 10.3 |
| Diluted EPS                  | \$ (19.53) | \$ (6.18) \$ | (2.96) \$ | (0.65) \$ | (0.48) | \$ (3.00) | \$ (0.53) | \$ (0.81) | \$ (0.89) | \$ (0.99) | \$ (3.23) | \$ (3.13) | \$ (2.78) | \$ (1.20) | \$ 1 | l.17 |
| Basic Shares Outstanding     | 0.7        | 0.7          | 2.2       | 22.7      | 22.9   | 12.2      | 25.6      | 27.1      | 27.2      | 27.3      | 26.8      | 29.8      | 32.3      | 33.3      | 3    | 34.3 |
| Diluted Shares Outstanding   | 0.7        | 0.7          | 2.2       | 22.7      | 22.9   | 12.2      | 29.6      | 27.1      | 27.2      | 27.3      | 27.8      | 30.8      | 33.3      | 34.3      |      | 35.3 |

Source: SEC Filings and Leerink Partners Research

| ZFGN BS & CFS (\$MM) GAAP  | 2013   | 1Q14  | 2Q14  | 3Q14   | 4Q14   | 2014E  | 1Q15   | 2Q15E  | 3Q15E  | 4Q15E  | 2015E  | 2016E  | 2017E  | 2018E  | 2019E  |
|----------------------------|--------|-------|-------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|--------|
|                            |        |       |       |        |        |        |        |        |        |        |        |        |        |        |        |
| Net Cash                   | 35.5   | 31.1  | 126.9 | 119.6  | 102.4  | 102.4  | 222.1  | 201.9  | 178.5  | 151.7  | 151.7  | 65.6   | (19.0) | (46.2) | 6.6    |
| Cash & Equivalents         | 35.5   | 38.5  | 134.2 | 127.0  | 115.5  | 115.5  | 234.2  | 212.9  | 188.5  | 160.6  | 160.6  | 70.4   | (15.3) | (46.2) | 6.6    |
| Debt                       | -      | 7.4   | 7.4   | 7.4    | 13.1   | 13.1   | 12.0   | 11.0   | 10.0   | 8.9    | 8.9    | 4.7    | 3.7    | -      | =      |
| Change in Cash             | 25.6   | 3.0   | 95.8  | (7.2)  | (11.5) | 80.0   | 118.6  | (21.3) | (24.4) | (27.9) | 45.1   | (90.2) | (85.6) | (30.9) | 52.8   |
| Operating Cash Flow        | (15.0) | (4.0) | (6.9) | (6.7)  | (17.2) | (34.9) | (9.9)  | (20.2) | (22.4) | (24.8) | (77.3) | (81.1) | (74.9) | (20.9) | 62.8   |
| Net Income (Loss)          | (14.0) | (4.5) | (6.4) | (14.7) | (10.9) | (36.5) | (13.5) | (22.0) | (24.3) | (27.0) | (86.8) | (93.4) | (89.8) | (39.9) | 40.3   |
| SOE                        | 0.4    | 0.2   | 0.4   | 0.9    | 0.6    | 2.0    | 1.1    | 1.8    | 1.9    | 2.2    | 6.9    | 8.4    | 9.3    | 10.0   | 12.6   |
| D&A                        | 0.0    | 0.0   | -     | -      | -      | 0.0    | -      | -      | -      | -      | -      | 4.0    | 5.6    | 9.0    | 10.0   |
| Other                      | (1.4)  | 0.3   | (0.9) | 7.1    | (6.9)  | (0.4)  | 2.5    | -      | -      | -      | 2.5    | -      | -      | -      | -      |
| Investing Cash Flow        | (0.0)  | (0.0) | -     | -      | -      | (0.0)  | -      | -      | (1.0)  | (2.0)  | (3.0)  | (5.0)  | (7.0)  | (10.0) | (10.0) |
| CapEx                      | (0.0)  | (0.0) | -     | -      | -      | (0.0)  | -      | -      | (1.0)  | (2.0)  | (3.0)  | (5.0)  | (7.0)  | (10.0) | (10.0) |
| Other                      | -      | -     | -     | -      | -      | -      | -      | -      | -      | -      | -      | -      |        |        |        |
| Financing Cash flow        | 40.6   | 7.0   | 102.7 | (0.5)  | 5.7    | 114.9  | 128.6  | (1.0)  | (1.0)  | (1.0)  | 125.4  | (4.2)  | (3.7)  | -      | -      |
| Equity Issuance (Buyback)  | 40.8   | 0.4   | 102.7 | -      | -      | 103.1  | 129.6  | -      | -      | -      | 129.6  | -      |        | -      | -      |
| Debt Issuance (Retirement) | -      | 7.4   | -     | (0.5)  | 5.7    | 12.6   | (1.0)  | (1.0)  | (1.0)  | (1.0)  | (4.2)  | (4.2)  | (3.7)  | -      | =-     |
| Other                      | (0.2)  | (0.8) | -     | -      | -      | (0.8)  | -      | -      | -      | -      | -      | -      | -      | -      | -      |

Source: SEC Filings and Leerink Partners Research

| ZFGN DCF Analysis                | 2014 | 2015E | 2016E | 2017E | 2018E | 2019E | 2020E | 2021E | 2022E | 2023E | 2024E | 2025E | 2026E | 2027E | 2028E | 2029E | 2030E | 2031E | TV   |
|----------------------------------|------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|-------|------|
| Cash Flow From Operations (\$MM) | (35) | (77)  | (81)  | (75)  | (21)  | 63    | 128   | 229   | 266   | 291   | 332   | 364   | 395   | 425   | 448   | 461   | 346   | 177   |      |
| Cash Flow From Investing (\$MM)  | (0)  | (3)   | (5)   | (7)   | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  | (10)  |      |
| Net Borrowing (Repayment) (\$MM) | 5    | (4)   | (4)   | (4)   | -     | -     | -     | -     | -     | -     | -     | -     | -     | -     | -     | -     | -     | -     |      |
| Free Cash Flow (\$MM)            | (30) | (85)  | (90)  | (86)  | (31)  | 53    | 118   | 219   | 256   | 281   | 322   | 354   | 385   | 415   | 438   | 451   | 336   | 167   | 1889 |
| Discount Periods                 | -    | -     | 1     | 2     | 3     | 4     | 5     | 6     | 7     | 8     | 9     | 10    | 11    | 12    | 13    | 14    | 15    | 16    |      |
| NPV FCF (\$MM)                   | -    | (63)  | (83)  | (71)  | (23)  | 36    | 72    | 120   | 126   | 125   | 129   | 128   | 125   | 122   | 116   | 107   | 72    | 32    | 365  |

| Sum NPV FCF (\$MM)          | 1434        |
|-----------------------------|-------------|
| Net Cash 1Q15E              | 222         |
| Implied ZFGN Mkt Cap (\$MM) | \$<br>1,657 |
| ZFGN Per Share Value        | \$<br>55.96 |

| Cost of Equity            | 11%  |
|---------------------------|------|
| TG Rate                   | 2%   |
| Diluted Shares Oustanding | 29.6 |

Source: Leerink Partners Research

| Prader Willi Syndrome Revenue Model   | 2014                   | 2015E                  | 2016E                  | 2017E                  | 2018E                   | 2019E                     | 2020E                     | 2021E                     | 2022E                       | 2023E                       | 2024E                       | 2025E                       | 2026E                       | 2027E                       | 2028E                       | 2029E                       | 2030E                       | 2031E                     |
|---|------------------------|------------------------|------------------------|------------------------|-------------------------|---------------------------|---------------------------|---------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|-----------------------------|---------------------------|
| PWS Patients in the US  | 7,500                  | 7,568                  | 7,636                  | 7,704                  | 7,774                   | 7,844                     | 7,914                     | 7,985                     | 8,057                       | 8,130                       | 8,203                       | 8,277                       | 8,351                       | 8,426                       | 8,502                       | 8,579                       | 8,656                       | 8,734                     |
| % >12 years old   | 50%                    | 50%                    | 50%                    | 50%                    | 50%                     | 50%                       | 50%                       | 50%                       | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                       |
| US PWS Patients >12 yr/old  | 3,750                  | 3,784                  | 3,818                  | 3,852                  | 3,887                   | 3,922                     | 3,957                     | 3,993                     | 4,029                       | 4,065                       | 4,102                       | 4,138                       | 4,176                       | 4,213                       | 4,251                       | 4,289                       | 4,328                       | 4,367                     |
| % treated with Beloranib<br>PWS Patients on Beloranib                           | 0.0%                   | 0.0%                   | 0.0%                   | 1.0%<br>39             | 8.0%<br>311             | 16.0%<br>627              | 21.0%<br>831              | 25.0%<br>998              | 28.0%<br>1,128              | 30.0%<br>1,219              | 32.0%<br>1,312              | 33.0%<br>1,366              | 34.0%<br>1,420              | 35.0%<br>1,475              | 35.0%<br>1,488              | 35.0%<br>1,501              | 28.0%<br>1,212              | 14.0%<br>611              |
| Annual Cost of Therapy<br>Gross Revenues (\$MM)                                 | \$150,000<br>\$0.0     | \$150,000<br>\$0.0     | \$150,000<br>\$0.0     | \$150,000<br>\$5.8     | \$150,000<br>\$46.6     | \$150,000<br>\$94.1       | \$150,000<br>\$124.6      | \$150,000<br>\$149.7      | \$150,000<br>\$169.2        | \$150,000<br>\$182.9        | \$150,000<br>\$196.9        | \$150,000<br>\$204.9        | \$150,000<br>\$213.0        | \$150,000<br>\$221.2        | \$150,000<br>\$223.2        | \$150,000<br>\$225.2        | \$150,000<br>\$181.8        | \$150,000<br>\$91.7       |
| %<12 years old  | 50%                    | 50%                    | 50%                    | 50%                    | 50%                     | 50%                       | 50%                       | 50%                       | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                       |
| US PWS Patients <12 yr/old  | 3,750                  | 3,784                  | 3,818                  | 3,852                  | 3,887                   | 3,922                     | 3,957                     | 3,993                     | 4,029                       | 4,065                       | 4,102                       | 4,138                       | 4,176                       | 4,213                       | 4,251                       | 4,289                       | 4,328                       | 4,367                     |
| % treated with Beloranib<br>PWS Patients on Beloranib<br>Annual Cost of Therapy | 0.0%<br>-<br>\$150,000 | 0.0%<br>-<br>\$150,000 | 0.0%<br>-<br>\$150,000 | 0.0%<br>-<br>\$150,000 | 1.0%<br>39<br>\$150,000 | 8.5%<br>333<br>\$150,000  | 16.0%<br>633<br>\$150,000 | 21.0%<br>838<br>\$150,000 | 25.0%<br>1,007<br>\$150,000 | 28.0%<br>1,138<br>\$150,000 | 30.0%<br>1,230<br>\$150,000 | 32.0%<br>1,324<br>\$150,000 | 33.0%<br>1,378<br>\$150,000 | 34.0%<br>1,433<br>\$150,000 | 35.0%<br>1,488<br>\$150,000 | 35.0%<br>1,501<br>\$150,000 | 24.5%<br>1,060<br>\$150,000 | 12.3%<br>535<br>\$150,000 |
| Gross Revenues (\$MM)  Approval Probability                                     | \$0.0<br>70%           | \$0.0<br>70%           | \$0.0<br>70%           | \$0.0<br>70%           | \$5.8<br>70%            | \$50.0<br>70%             | \$95.0<br>70%             | \$125.8<br>70%            | \$151.1                     | \$170.7<br>70%              | \$184.6<br>70%              | \$198.6<br>70%              | \$206.7                     | \$214.9<br>70%              | \$223.2<br>70%              | \$225.2<br>70%              | \$159.1<br>70%              | \$80.2<br>70%             |
| US P(w) Beloranib PWS Revenues  | \$0.0                  | \$0.0                  | \$0.0                  | \$4.0                  | \$36.7                  | \$100.9                   | \$153.7                   | \$192.8                   | \$224.2                     | \$247.6                     | \$267.0                     | \$282.4                     | \$293.8                     | \$305.2                     | \$312.5                     | \$315.3                     | \$238.6                     | \$120.4                   |
|   | , , ,                  |                        | - '                    |                        |                         |                           |                           |                           |                             |                             |                             |                             |                             |                             |                             |                             |                             |                           |
| PWS Patients in the EU  | 12,000                 | 12,108                 | 12,217                 | 12,327                 | 12,438                  | 12,550                    | 12,663                    | 12,777                    | 12,892                      | 13,008                      | 13,125                      | 13,243                      | 13,362                      | 13,482                      | 13,604                      | 13,726                      | 13,850                      | 13,974                    |
| % >12 years old   | 50%                    | 50%                    | 50%                    | 50%                    | 50%                     | 50%                       | 50%                       | 50%                       | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                       |
| EU PWS Patients >12 yr/old  | 6,000                  | 6,054                  | 6,108                  | 6,163                  | 6,219                   | 6,275                     | 6,331                     | 6,388                     | 6,446                       | 6,504                       | 6,562                       | 6,621                       | 6,681                       | 6,741                       | 6,802                       | 6,863                       | 6,925                       | 6,987                     |
| % treated with Beloranib<br>PWS Patients on Beloranib                           | 0.0%                   | 0.0%                   | 0.0%                   | 0.0%                   | 2.5%<br>155             | 4.0%<br>251               | 6.0%<br>380               | 8.0%<br>511               | 11.0%<br>709                | 13.0%<br>846                | 15.0%<br>984                | 16.0%<br>1,059              | 17.0%<br>1,136              | 18.0%<br>1,213              | 19.0%<br>1,292              | 20.0%<br>1,373              | 16.0%<br>1,108              | 8.0%<br>559               |
| Annual Cost of Therapy<br>Gross Revenues  | \$90,000<br>\$0.0      | \$90,000<br>\$0.0      | \$90,000<br>\$0.0      | \$90,000<br>\$0.0      | \$90,000<br>\$14.0      | \$90,000<br>\$22.6        | \$90,000<br>\$34.2        | \$90,000<br>\$46.0        | \$90,000<br>\$63.8          | \$90,000<br>\$76.1          | \$90,000<br>\$88.6          | \$90,000<br>\$95.3          | \$90,000<br>\$102.2         | \$90,000<br>\$109.2         | \$90,000<br>\$116.3         | \$90,000<br>\$123.5         | \$90,000<br>\$99.7          | \$90,000<br>\$50.3        |
| %<12 years old  | 50%                    | 50%                    | 50%                    | 50%                    | 50%                     | 50%                       | 50%                       | 50%                       | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                         | 50%                       |
| US PWS Patients <12 yr/old  | 6,000                  | 6,054                  | 6,108                  | 6,163                  | 6,219                   | 6,275                     | 6,331                     | 6,388                     | 6,446                       | 6,504                       | 6,562                       | 6,621                       | 6,681                       | 6,741                       | 6,802                       | 6,863                       | 6,925                       | 6,987                     |
| % treated with Beloranib<br>PWS Patients on Beloranib                           | 0.0%                   | 0.0%                   | 0.0%                   | 0.0%                   | 0.0%                    | 2.5%<br>157               | 4.0%<br>253               | 6.0%<br>383               | 8.0%<br>516                 | 11.0%<br>715                | 13.0%<br>853                | 15.0%<br>993                | 17.0%<br>1,136              | 18.0%<br>1,213              | 19.0%<br>1,292              | 20.0%<br>1,373              | 10.0%<br>692                | 4.0%<br>279               |
| Annual Cost of Therapy  | \$90,000               | \$90,000               | \$90,000               | \$90,000               | \$90,000                | \$90,000                  | \$90,000                  | \$90,000                  | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                    | \$90,000                  |
| Gross Revenues (\$MM)   | \$0.0                  | \$0.0                  | \$0.0                  | \$0.0                  | \$0.0                   | \$14.1                    | \$22.8                    | \$34.5                    | \$46.4                      | \$64.4                      | \$76.8                      | \$89.4                      | \$102.2                     | \$109.2                     | \$116.3                     | \$123.5                     | \$62.3                      | \$25.2                    |
| Approval Probability  | 70%                    | 70%                    | 70%                    | 70%                    | 70%                     | 70%                       | 70%                       | 70%                       | 70%                         | 70%                         | 70%                         | 70%                         | 70%                         | 70%                         | 70%                         | 70%                         | 70%                         | 70%                       |
| EU P(w) Beloranib PWS Revenues  | \$0.0                  | \$0.0                  | \$0.0                  | \$0.0                  | \$9.8                   | \$25.7                    | \$39.9                    | \$56.3                    | \$77.2                      | \$98.3                      | \$115.8                     | \$129.3                     | \$143.1                     | \$152.9                     | \$162.8                     | \$172.9                     | \$113.4                     | \$52.8                    |
| WW Beloranib Gross Sales<br>WW Beloranib P(w) Sales                             | \$0.0<br><b>\$0.0</b>  | \$0.0<br><b>\$0.0</b>  | \$0.0<br><b>\$0.0</b>  | \$5.8<br><b>\$4.0</b>  | \$66.5<br><b>\$46.5</b> | \$180.8<br><b>\$126.6</b> | \$276.6<br><b>\$193.6</b> | \$356.0<br><b>\$249.2</b> | \$430.5<br>\$301.4          | \$494.1<br><b>\$345.9</b>   | \$546.8<br>\$382.8          | \$588.2<br>\$411.8          | \$624.1<br>\$436.9          | \$654.5<br>\$458.1          | \$679.0<br><b>\$475.3</b>   | \$697.5<br>\$488.2          | \$502.9<br><b>\$352.0</b>   | \$247.4<br><b>\$173.2</b> |

| Assumptions             |           |
|-------------------------|-----------|
| Beloranib US Cost       | \$150,000 |
| Beloranib EU Cost       | \$90,000  |
| Probability of Approval | 70%       |

Source: Company Presentations and Leerink Partners Research

| HIAO Revenue Model                             | 2014      | 2015E     | 2016E     | 2017E     | 2018E     | 2019E     | 2020E     | 2021E     | 2022E     | 2023E     | 2024E     | 2025E     | 2026E     | 2027E     | 2028E     | 2029E     | 2030E     | 2031E     |
|--|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|-----------|
|  |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |           |
| HIAO Patients in the US                        | 6,260     | 6,316     | 6,373     | 6,431     | 6,488     | 6,547     | 6,606     | 6,665     | 6,725     | 6,786     | 6,847     | 6,908     | 6,971     | 7,033     | 7,097     | 7,160     | 7,225     | 7,290     |
| % with post-treatment hypothalamic dysfunction | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       |
| beloranib HIAO candidates                      | 3,130     | 3,158     | 3,187     | 3,215     | 3,244     | 3,273     | 3,303     | 3,333     | 3,363     | 3,393     | 3,423     | 3,454     | 3,485     | 3,517     | 3,548     | 3,580     | 3,612     | 3,645     |
| % treated with Beloranib                       | 0.0%      | 0.0%      | 0.0%      | 0.0%      | 2.0%      | 6.0%      | 13.0%     | 18.0%     | 24.0%     | 29.0%     | 32.0%     | 34.0%     | 36.0%     | 38.0%     | 40.0%     | 40.0%     | 32.0%     | 16.0%     |
| Patients on Beloranib                          | -         | -         | -         | -         | 65        | 196       | 429       | 600       | 807       | 984       | 1,095     | 1,174     | 1,255     | 1,336     | 1,419     | 1,432     | 1,156     | 583       |
| Annual Cost of Therapy                         | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 | \$150,000 |
| Gross Revenues (\$MM)                          | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$9.7     | \$29.5    | \$64.4    | \$90.0    | \$121.1   | \$147.6   | \$164.3   | \$176.2   | \$188.2   | \$200.4   | \$212.9   | \$214.8   | \$173.4   | \$87.5    |
| Approval Probability                           | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       |
| US P(w) Beloranib HIAO Revenues                | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$5.8     | \$17.7    | \$38.6    | \$54.0    | \$72.6    | \$88.6    | \$98.6    | \$105.7   | \$112.9   | \$120.3   | \$127.7   | \$128.9   | \$104.0   | \$52.5    |
| HIAO Patients in the EU                        | 14,850    | 14,984    | 15,119    | 15,255    | 15,392    | 15,530    | 15,670    | 15,811    | 15,953    | 16,097    | 16,242    | 16,388    | 16,536    | 16,684    | 16,835    | 16,986    | 17,139    | 17,293    |
| % with post-treatment hypothalamic dysfunction | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       | 50%       |
| beloranib HIAO candidates                      | 7,425     | 7,492     | 7,559     | 7,627     | 7,696     | 7,765     | 7,835     | 7,906     | 7,977     | 8,049     | 8,121     | 8,194     | 8,268     | 8,342     | 8,417     | 8,493     | 8,569     | 8,647     |
| % treated with Beloranib                       | 0.0%      | 0.0%      | 0.0%      | 0.0%      | 1.0%      | 4.0%      | 8.0%      | 12.0%     | 16.0%     | 20.0%     | 22.0%     | 24.0%     | 26.0%     | 28.0%     | 30.0%     | 30.0%     | 24.0%     | 12.0%     |
| Patients on Beloranib                          | -         | -         | -         | -         | 77        | 311       | 627       | 949       | 1,276     | 1,610     | 1,787     | 1,967     | 2,150     | 2,336     | 2,525     | 2,548     | 2,057     | 1,038     |
| Annual Cost of Therapy                         | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    | 90,000    |
| Gross Revenues (\$MM)                          | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$6.9     | \$28.0    | \$56.4    | \$85.4    | \$114.9   | \$144.9   | \$160.8   | \$177.0   | \$193.5   | \$210.2   | \$227.3   | \$229.3   | \$185.1   | \$93.4    |
| Approval Probability                           | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       | 60%       |
| EU P(w) Beloranib HIAO Revenues                | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$4.2     | \$16.8    | \$33.8    | \$51.2    | \$68.9    | \$86.9    | \$96.5    | \$106.2   | \$116.1   | \$126.1   | \$136.4   | \$137.6   | \$111.1   | \$56.0    |
| WW Gross Beloranib HIAO Revenues               | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$16.7    | \$57.4    | \$120.8   | \$175.4   | \$235.9   | \$292.5   | \$325.1   | \$353.2   | \$381.7   | \$410.7   | \$440.2   | \$444.1   | \$358.5   | \$180.9   |
| WW P(w) Beloranib HIAO Revenues                | \$0.0     | \$0.0     | \$0.0     | \$0.0     | \$10.0    | \$34.4    | \$72.5    | \$105.2   | \$141.6   | \$175.5   | \$195.1   | \$211.9   | \$229.0   | \$246.4   | \$264.1   | \$266.5   | \$215.1   | \$108.5   |

| Assumptions             |           |
|-------------------------|-----------|
| Beloranib US Cost       | \$150,000 |
| Beloranib EU Cost       | \$90,000  |
| Probability of Approval | 60%       |

Source: Company Presentations and Leerink Partners Research

| Product   | Event                                   | Timing    |
|-----------|---|-----------|
| Beloranib | Initiate US Phase III PWS Trial         | 3Q14      |
| Beloranib | Initiate Phase IIb Severe Obesity Trial | 4Q14      |
| Beloranib | Phase IIa HIAO Data                     | 1Q15      |
| ZGN-839   | File NASH/Type II Diabetes IND          | mid-2015  |
| Beloranib | Initate EU Phase III PWS Trial          | mid-2015  |
| Beloranib | 6 Month Interim Severe Obesity Data     | 4Q15/1Q16 |
| Beloranib | Initiate Phase III HIAO Study           | 2016      |
| Beloranib | 6 Mo. Phase III PWS Data                | 1Q16      |

Source: Company Presentations and Leerink Partners Research



# **Disclosures Appendix Analyst Certification**

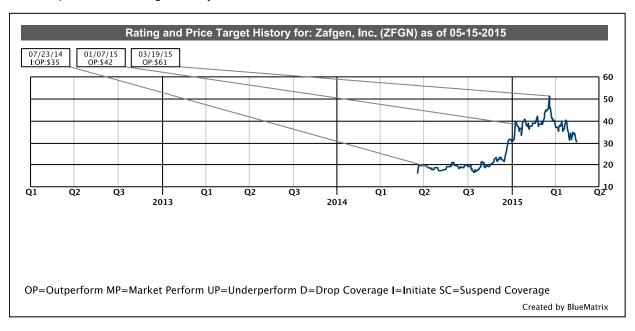
I, Joseph P. Schwartz, certify that the views expressed in this report accurately reflect my views and that no part of my compensation was, is, or will be directly related to the specific recommendation or views contained in this report.

### **Valuation**

We derive a  $\sim$ \$56 per share value for ZFGN using an 11% discount rate and a 2% terminal growth rate. Our price target assumes 70% and a 60% probabilities of beloranib approval in PWS and HIAO, respectively, which leads to our risk adjusted peak sales estimates of  $\sim$ \$485MM and  $\sim$ \$270MM in each indication. We only model  $\sim$ \$200MM in peak sales in severe obesity, which we believe holds the potential to be very conservative if/when ZFGN generates pivotal beloranib data in orphan indications.

#### **Risks to Valuation**

Risks to our valuation include disappointing clinical data, regulatory setbacks, dilution risk from an additional equity offering, and commercial shortfalls. Because ZFGN has only one late stage product, the occurrence of any of these could impact the stock significantly.





| Dis       | stribution of Ratings/Investment Ban | king Services (IB) |       | erv./Past 12<br>Mos. |
|-----------|--------------------------------------|--------------------|-------|----------------------|
| Rating    | Count                                | Percent            | Count | Percent              |
| BUY [OP]  | 151                                  | 70.20              | 55    | 36.00                |
| HOLD [MP] | 64                                   | 29.80              | 2     | 3.00                 |
| 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 Zafgen, Inc. .

Leerink Partners LLC makes a market in Zafgen, Inc.

Leerink Partners LLC has acted as the manager for a public offering of Zafgen, Inc. in the past 12 months.

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