Lee Kalowski Chief Financial Officer Bicycle Therapeutics Ltd. 4 Hartwell Place Lexington, Massachusetts 02421

Re: Bicycle Therapeutics Ltd.
Amendment No. 1 to Draft Registration Statement on Form S-1
Submitted March 22, 2019
CIK 0001761612

Dear Mr. Kalowski:

We have reviewed your amended draft registration statement and have the following comments. In some of our comments, we may ask you to provide us with information so we

may better understand your disclosure.

Please respond to this letter by providing the requested information and either submitting $% \left(1\right) =\left(1\right) \left(1\right) +\left(1\right) \left(1\right) \left(1\right) +\left(1\right) \left(1\right) \left(1\right) \left(1\right) +\left(1\right) \left(1\right) \left$

an amended draft registration statement or publicly filing your registration statement on $% \left(1\right) =\left(1\right) \left(1\right$

 ${\tt EDGAR.}$ If you do not believe our comments apply to your facts and circumstances or do not

believe an amendment is appropriate, please tell us why in your response.

After reviewing the information you provide in response to these comments and your amended draft registration statement or filed registration statement, we may have additional

Amendment No. 1 to Draft Registration Statement on Form S-1 submitted March 22, 2019

Prospectus Summary Beyond Oncology, page 5

1. We note your response to prior comment 4. Please also revise your prospectus summary

to describe the ongoing Phase I clinical trial of THR-149.

Lee Kalowski

Bicycle Therapeutics Ltd.

April 4, 2019

Page 2

comments.

Business Our Collaborations

Our Other Collaborators

AstraZeneca, page 144

2. We note your statement that, pursuant to your collaboration agreement with ${\color{black} }$

AstraZenaca, you could receive more than \$1 billion in milestone payments and royalties.

We also note your disclosure that, pursuant to the collaboration

agreement, AstraZeneca owes you milestone fee of \$8 million for the first drug candidate

selected from each research program, and you are eligible to receive up to an additional

\$162 million in development, regulatory and commercial milestones on a research

program by research program basis, as well as tiered royalty payments of mid-single digits $% \left(1\right) =\left(1\right) +\left(1\right)$

based on a percentage of net sales. Please provide further details regarding how these

amounts could total over \$1 billion.

Management

Executive Officers, page 168

3. We note your updated disclosure regarding your Chief Business Officer, Peter Leone.

Please clarify Mr. Leone's activities between April 2016 and 2012. Please see Item $\,$

401(e)(1) of Regulation S-K.

You may contact Jim Dunn at 202-551-3724 or Isaac Esquivel at 202-551-3395 if you

have questions regarding comments on the financial statements and related matters. Please $\,$

contact Liz Walsh at 202-551-3696 or Christopher Edwards at 202-551-6761 with any other questions.

FirstName LastNameLee Kalowski

Corporation Finance Comapany NameBicycle Therapeutics Ltd.

& Insurance
April 4, 2019 Page 2
cc: Jonathan Schur
FirstName LastName

Sincerely,

Division of

Office of Healthcare