This review article covers biological effects of soluble PD-L1 (sPD-L1) in tumor pathogenesis, immune response, and immunotherapy and its role in cancer biology.
This specific strategy presented in this review is designed to describe other specific effects as potential prognostic and predictive biomarker for immunotherapy. It is evident that elevated levels of PD-L1 is related to disease severity, clinicopathological characteristics and recommendation of possible cancer therapy. The compiled data are supported with 2 figures and one table. The article concludes with 59 very recent literature references. This consolidated study constitutes important developments, which were never ever reported in such systematic and specific order and sequences.

The following suggested changes and recommendations should be introduced before the publication of the manuscript:

- Page 2. Line 68. "The PD-L1 gene" It is biomarker (as stated in line 15) or gene? This must be clarified with supporting literature reference!
- Page 2, Line 82. "Nonetheless, other possible sources cannot be excluded" Which are the other possible sources referred? Please, insert the literature reference.
- Page 3, Line 119. Insert literature reference after "homodimeric structure"
- Page 4, Figure 1 should be moved to line 104, (page 3) where is first mentioned.
- Page 7. Table 1 legend, second column headline should be changed to "Finding/prognosis"
- Page 9, Line 318. Conclusions. This section must be expanded. In its present format, the authors do not fully describe the desired/anticipated effects without citing the literature references. Authors should also include comparative data available in the literature by inserting the references numbers specifically in the first paragraph in lines 319-324.

  Additionally, the last sentence in line 329is not specific and authors should name the "future studies", which are needed to understand the behavior …

The manuscript is of good quality, well-written, and meets the standard for articles published in *Life.* I recommend it for publication after the correction of these suggested changes.