## **EDITORIAL**



# Standardisation of protocols can be crucial in long non-coding RNA research

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In this issue, Traversa et al. [1] reviewed our current knowledge about the role of circular and linear forms of PVT1 non-coding RNA in cancer and human diseases. They highlighted the technical challenges of these studies and raised a potential bias in the publications, which require more attention from researchers.

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#### MAIN

Along with the increasing number of discovered circular RNAs (circRNAs) the need of understanding their functional roles became an important topic. Plasmacytoma variant translocation 1 (*PVT1*) lncRNA gene produces both circular (circPVT1) and linear (lncPVT1) transcripts, and their expression has been found to correlate with tumour progression in different types of cancers.

Traversa et al. reviewed the literature about the lncPVT1 and circPVT1 transcripts, which have been reported to have similar functions in the pathophysiology of diseases such as solid tumours and haematological malignancies. They highlighted in Table 2 the frequent association between lncPVT1 and circPVT1 upregulation and poor prognosis of the patients with different types of cancers, including but not limited to colorectal, ovarian and gastric cancer [1]. Therefore, both the linear and the circular transcripts have been implicated to be potential prognostic biomarkers.

The authors suggest, that since the two molecules share the IncPVT1 exon 2 sequence, it is difficult to distinguish the expression of the two transcripts, which can result in a biased interpretation of the data. They pointed out that some of the observations regarding the shared functions of these transcripts can be a result of this technical issue, and therefore lead to incorrect conclusions.

Even though their review focus on the expression of circular and linear PVT1, their point goes beyond that. Their case raises attention to the potentially misleading interpretation of data during the investigation of circRNAs and can be valid for other long non-coding RNAs as well.

This review highlights the importance of solid understanding and the appropriate choice of the technical approaches to be used for the dissection of complex genomic loci. Besides, using several independent techniques to underneath experimental findings can be also critical.

The issue raised in the review is not entirely new. Researchers have been discussing about a "replication crisis" in science, including cancer research, for several years now, referring to the uncovered phenomena that a lot of published data cannot be reproduced due to different reasons. This reminds us that despite the constant publication pressure and the difficulties of publish

negative results, a thoroughly understanding of the used methods, careful planning of the experiments, preliminary measurements and application of different independent methods are extremely important. Besides, this can lead to expanding the discussion in the scientific community about standardise the experimental protocols and the interpretation of the data.

Generating and following guidelines similar to the MIQE (Minimum Information for Publication of Quantitative Real-Time PCR Experiments) guideline can highly increase the reproducibility and reliability of the results.

#### **OUTLOOK**

During these days, when generating a large amount of data is easily accessible, it is especially important to generate clear criteria on how we can improve our daily routine to not only generate a huge amount of data but to be sure that those data are going to be reliable, durable, and be a valuable part of our common scientific knowledge. Moreover, it is important to give space for debate and be open for discussion about how we can improve our standards.

## REFERENCE

 Traversa D, Simonetti G, Tolomeo D, Visci G, Macchia G, Ghetti M, et al. Unraveling similarities and differences in the role of circular and linear PVT1 in cancer and human disease. Br J Cancer. 2021 (this issue).

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KN and GAC equally contributed to the manuscript writing.

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#### ETHICS APPROVAL AND CONSENT TO PARTICIPATE

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# **CONSENT TO PUBLISH**

Not applicable.

## **COMPETING INTERESTS**

GAC is the scientific founder of Ithax Pharmaceuticals. KN declares no competing interests.

# **ADDITIONAL INFORMATION**

Correspondence and requests for materials should be addressed to George A. Calin.

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