Letter to the Editor



# Sequence analysis

# Comment on: 'ERGC: an efficient referential genome compression algorithm'

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Associate Editor: John Hancock

Received on 19 October 2015; revised on 24 November 2015; accepted on 25 November 2015

#### **Abstract**

**Motivation**: Data compression is crucial in effective handling of genomic data. Among several recently published algorithms, ERGC seems to be surprisingly good, easily beating all of the competitors.

**Results:** We evaluated ERGC and the previously proposed algorithms GDC and iDoComp, which are the ones used in the original paper for comparison, on a wide data set including 12 assemblies of human genome (instead of only four of them in the original paper). ERGC wins only when one of the genomes (referential or target) contains mixed-cased letters (which is the case for only the two Korean genomes). In all other cases ERGC is on average an order of magnitude worse than GDC and iDoComp.

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Supplementary information: Supplementary data are available at Bioinformatics online.

#### 1 Introduction

The rapid growth of genomic data in the last few years demands for efficient compression methods to facilitate their storage and transfer. One of the standard problems of this kind is (referential) compression of multiple individual genomes of the same species (Deorowicz *et al.*, 2015; Deorowicz and Grabowski, 2011; Kuruppu *et al.*, 2011; Ochoa *et al.*, 2014; Wandelt and Leser, 2013). As a pair of individual genomes differ only in variant loci, we can expect radical savings in storing a referentially encoded genome.

Recently, (Saha and Rajasekaran 2015) published a new referential genome compression algorithm, called ERGC. As the experimental results presented in this article seemed to suggest that ERGC is surprisingly good, winning easily in compression ratio over the previously proposed algorithms GDC (Deorowicz and Grabowski, 2011) and iDoComp (Ochoa *et al.*, 2014), we decided to take a closer look at ERGC and repeat the experiments on a wider set of data.

#### 2 Discussion

In the first experiment, we repeated the referential compression evaluation from the ERGC paper, where the  $D_1,\ldots,D_5$  experiments are specified in Table 1 in the cited paper and the actual results are shown there in Table 2. In each  $D_i$  experiment, all 24 chromosomes of one human genome are referentially compressed, given some other human genome as a reference. We repeated these experiments in our Table 1.

As it can be observed, the results differ significantly for cases  $D_1$  and  $D_5$ . First, the cited paper incorrectly reported 'NA' for the GDC algorithm in these two cases. Moreover, iDoComp achieves 6288 KB on  $D_1$  and 33 158 KB on  $D_5$ , rather than 65 708.47 KB and 209 380.79 KB as reported in the cited paper (note the order of magnitude difference). Finally, applying ERGC on  $D_5$  resulted in a compressed file of 9373 KB, instead of the reported 19 396.40 KB.

With the corrected results, we concluded that ERGC achieves the best compression ratio in all cases except in  $D_1$ , where it is

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Table 1. Rerun experiments from Table 2 in the ERGC paper

| Dataset          | Target size | GDC size | iDoComp size | ERGC size |  |
|------------------|-------------|----------|--------------|-----------|--|
| $\overline{D_1}$ | 3132        | 6439     | 6288         | 7890      |  |
| $D_2$            | 3132        | 29 831   | 30 437       | 9217      |  |
| $D_3$            | 3132        | 35 227   | 33 926       | 9404      |  |
| $D_4$            | 2938        | 12 293   | 7213         | 4914      |  |
| $D_5$            | 3124        | 34 806   | 33 158       | 9373      |  |

Table 2. Rerun experiments from Table 1 above, with the difference that the KOREAN genomes are converted to upper case

| Dataset          | Target size | GDC size | iDoComp size | ERGC size |  |
|------------------|-------------|----------|--------------|-----------|--|
| $\overline{D_2}$ | 3132        | 7660     | 7473         | 9273      |  |
| $D_3$            | 3132        | 7849     | 7691         | 9449      |  |
| $D_4$            | 2938        | 1073     | 1020         | 1246      |  |
| $D_5$            | 3124        | 7820     | 7754         | 9425      |  |

The  $D_1$  experiment is skipped run since both the reference and the target in it consist of upper case DNA symbols only.

Table 3. Summary of the results for all the considered pairs both for chromosomes 10 and 20

| Reference | Chromosome 10 |          |                     | Chromosome 20 |             |          |                   |           |
|-----------|---------------|----------|---------------------|---------------|-------------|----------|-------------------|-----------|
|           | Target size   | GDC size | iDoComp size        | ERGC size     | Target size | GDC size | iDoComp size      | ERGC size |
| CHM1_1.0  | 1 495 816     | 10 229   | 11 805              | 289 929       | 694 646     | 4242     | 5132              | 135 412ª  |
| CHM1_1.1  | 1 496 019     | 9505     | 11 054              | 290 014       | 694 641     | 4252     | 5088              | 100 306   |
| CSA       | 1 497 653     | 51 057   | 59 500 <sup>b</sup> | 339 998       | 695 024     | 13 741   | 15 695            | 146 233   |
| HG17      | 1 496 426     | 8146     | 7801                | 167 270       | 695 127     | 3457     | 3893              | 48 193    |
| HG18      | 1 496 465     | 8106     | 7753                | 145 209       | 695 127     | 3457     | 3893              | 48 193    |
| HG19      | 1 496 303     | 8153     | 7913                | 227 746       | 694 529     | 3472     | 3919              | 48 294    |
| HG38      | 1 498 065     | 9329     | 9543                | 222 986       | 693 090     | 2743     | 2798              | 78 216    |
| HuRef     | 1 502 946     | 12 680   | 15 582              | 295 338       | 697 946     | 5037     | 5862              | 95 444    |
| KO131     | 1 496 143     | 16 388   | 18 246 <sup>b</sup> | 158 849       | 694 978     | 6169     | 6984 <sup>b</sup> | 75 827    |
| KO224     | 1 496 143     | 14 736   | 16 997 <sup>b</sup> | 158 635       | 694 978     | 5486     | 6543 <sup>b</sup> | 68 173    |
| WGSA      | 1 503 392     | 25 607   | 31 897              | 336 064       | 697 939     | 10 403   | 12 470            | 137 623   |
| YH        | 1 496 143     | 9388     | 8785                | 153 532       | 694 978     | 4063     | 4406              | 49 951    |

<sup>&</sup>lt;sup>a</sup>This total ERGC result is not certain, as ERGC crashed on the pair of sequences: CHM1\_1.0 (reference), CHM1\_1.1 (target).

outperformed by both GDC and iDoComp. Taking a closer look, we observed that in all but  $D_1$  one of the KOREAN genomes is used either as a reference or a target genome. The KOREAN genomes differ from the other ones in that they contain both lower and upper case letters (see Supplementary material). They also contain non-standard symbols, similarly to the YH genome.

To better understand if the potential of ERGC is due to the design of the algorithm itself, or to the handling of the lower and upper case letters and/or the non-standard symbols, we simulated these experiments again  $(D_2, \ldots, D_5)$  with the KOREAN genomes transformed to all upper case. Surprisingly, as demonstrated in Table 2, ERGC achieved the worst compressed size in all cases. The reported results suggest that the gain of ERGC comes from the handling of the lower and upper case symbols.

Since the previous experiments may not be representative enough, we decided to use a wider set of data. Specifically, we chose the human genome datasets from the GDC 2 paper (Deorowicz *et al.*, 2015), which is a superset of the collections from (Deorowicz and Grabowski, 2011) and (Ochoa *et al.*, 2014). To reduce the amount of computations, we

chose only two chromosomes from each genome: chr10 and chr20. Then, to follow the ERGC paper methodology, we tested referential compression in pairs of chromosomes, in a round-robin fashion. In Table 3, we show the sums of compressed sizes; detailed results are presented in the Supplementary material. ERGC is outperformed in all cases by GDC and iDoComp.

#### 3 Conclusion

From the conducted experiments we can draw several conclusions. Firstly, the reported compression results on the  $D_1$  and  $D_5$  cases in the cited paper are not correct for both GDC and iDoComp. Secondly, ERGC seems to get the main advantage due to specific handling of lower and upper case difference between the target and the reference genome. We believe that using the KOREAN genomes is fair, however, the authors should have provided simulations on a wider set of data where the gain does not come from the lower and upper case difference. Finally, after running several experiments on different datasets, we can conclude that ERGC performs in general poorly when compared with GDC and iDoComp.

<sup>&</sup>lt;sup>b</sup>These total iDoComp results are taken from its bug-fix version (https://github.com/mikelhernaez/iDoComp, v1.2) as the earlier version v1.1 crashed on five pairs of sequences.

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## **Funding**

This work has been supported by a fellowship from the Basque Government, a grant from the Center for Science of Information (CSoI), the 2014-07364-01 NIH grant and the 1157849-1-QAZCC NSF grant. It is also supported by the Polish National Science Centre under the project DEC-2011/03/B/ST6/01588. The infrastructure supported by POIG.02.03.01-24-099/13 grant: 'GeCONiI-Upper' Silesian Center for Computational Science and Engineering'.

Conflict of Interest: none declared.

## References

Deorowicz, S. et al. (2015) GDC 2: Compression of large collections of genomes,. Sci. Reports, 5, Article no. 11565.

Deorowicz, S. and Grabowski, S. (2011) Robust relative compression of genomes with random access, *Bioinformatics*, 27, 2979–2986.

- Kuruppu,S. et al. (2011) Optimized relative Lempel-Ziv compression of genomes. In: Reynolds,M. (ed.) Proceedings of the ACSC Australasian Computer Science Conference. Australian Computer Society, Inc., Sydney, Australia, pp. 91–98.
- Ochoa, I. et al. (2015) iDoComp: a compression scheme for assembled genomes. Bioinformatics, 31, 626–633.
- Saha,S. and Rajasekaran,S. (2015) ERGC: An efficient referential genome compression algorithm. *Bioinformatics*, 31, 3468–3475.
- Wandelt,S. and Leser,U. (2013) FRESCO: Referential compression of highly similar sequences,. IEEE/ACM Trans. Comput. Biol. Bioinformatics, 10, 1275–1288.