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STUDYING YEAST PROVIDES NEW INSIGHT TO GENOME EVOLUTION

- Researchers at the Centre for Genomic Regulation have proposed a new theory about the origins of the yeast lineage 100 million years ago.
- By studying the whole-genome duplication in yeast, the scientists have determined that it actually represents a hybrid between two distinct species. Their new proposal differs radically from the currently accepted theories within the scientific community.
- The results describe the mechanism that led to the duplication of the yeast genome and offers new insight to the origin and evolutionary consequences of whole-genome duplications, which have occurred not only in yeast but also in plants and vertebrates.

Researchers from the Centre for Genomic Regulation (CRG; Barcelona, Spain) have proposed a new theory to explain the origin of whole genome duplication at the beginning of the yeast lineage. Yeasts are single-celled fungi that originated over 100 million years ago. The ability of these organisms to ferment carbohydrates is widely used for food and drink fermentation. Yeasts are also one of the most commonly used model organisms in research. For example, the yeast *Saccharomyces cerevisiae*, which is used to make bread, wine and beer, was the first eukaryotic organism to be sequenced (in 1996) and is a key model organism for studying molecular and cellular biology.

Once the yeast genome sequences were available, researchers were able to determine that the yeast genome contains more than 50 repeated fragments. Since then, the scientific community has accepted the theory that yeast underwent a whole genome duplication, a phenomenon that is not isolated and can also be found in other species. For instance, we know that whole genome duplications were important in the early evolution of vertebrates and that it is a very common phenomenon in plants, especially cultivated ones.

The CRG scientists Marina Marcet-Houben and Toni Gabaldón (CRG group leader and ICREA Research Professor) have now studied the origins of the whole genome duplication in yeast to gain a more thorough understanding of this phenomenon, which is thought to have played a key role the evolution and adaption of the species. Their results were published today in the journal PLoS Biology. Unexpectedly, they show that the appearance of duplicated genes was not caused by a simple duplication of the whole genome but rather by a hybridization of two different species. Their proposal, which is at odds with the currently most widely accepted theory in the scientific community, provides new insight into this key process during genome evolution and the origins of species.



"When we first saw the results of our study, we thought there had been some kind of mistake. Honestly, when the results are not what you expect and contradict what is established, the first thing you do is think that they were affected by some kind of problem. But once all the potential problems have been discarded, you begin to interpret the data objectively, without preconceived ideas, and to do real science. That's when we started to consider the different possible explanations and to work on a new idea," explained Toni Gabaldón, the lead investigator of the study and head of the Comparative Genomics Group. He added, "It's one of those magical moments of research when, once you open your mind enough, you can surrender to the evidence in the data and discard what you had considered as a proven fact to adopt an entirely new paradigm, no matter how implausible it seems at first. Afterwards you reel in your new paradigm and see that it also explains other independent observations.

Looking at the past to understand the present and predict the future

Once genomics entered into evolutionary studies, scientists were able to show that the phenomenon of whole genome duplication is common to several species, which leads us to believe that it is one of the key mechanisms in the evolution and adaptability of species. Duplicating the genome would give an organism multiple copies of genes, so that it could vary some of these copies over time without loosing the original and thus produce new proteins with novel functions, which could give it an evolutionary advantage by permitting it to adapt more readily and to cope with different conditions.

However, it would seem intuitively to make sense that incorporating something different through hybridization, rather than duplicating what is already present, would lead more rapidly to a higher degree of diversity and to better adaptation. The problems with this are that most hybrid crosses are not viable, and if viable, the resulting hybrids are sterile and therefore unable to pass on their new genetic information to their offspring. Because of this, and because of the similarities of the repeated genes, the idea of duplicating the whole genome previously seemed more plausible.

The work of Marcet-Houben and Gabaldón has now revealed that, for yeast, hybridization was indeed behind the duplication of some genes. The researchers analyzed genomic data with computational tool, based on cutting-edge computational methods, and designed by the Gabaldón group, to study the phylogenetic trees of yeast families. This tool allowed the researchers to reconstruct gene duplications and to determine what happened in evolutionary time, making it a computational equivalent of carbon-14 dating for fossils.

To their surprise, they found that the age of some duplicated genes seemed to be much greater than that predicted by the theory of whole genome duplication. Rather than supporting a genome duplication event at the time when yeast evolved to have twice the number of chromosomes, their data indicated that the duplicated genes had begun to diverge long before. This result suggested the possibility of hybridization between species. In this case, the genes that have been duplicated still differ from each other, so that their divergence preceded the duplication of the chromosome.



The hybridization hypothesis has strong implications on how we interpret the origin and evolution of duplicated genomes. For instance, we no longer need to imagine that massive and rapid changes are necessary to generate new functions from duplicated genome regions, since hybridization combines the properties of the two parental lines from day one and opens the door to new ecological and evolutionary opportunities.

The new proposal is strengthened when we see that the number of genome duplications is much higher in cultivated plants. "We should consider whether genome duplications in other species also hide hybridizations since, for example, it seems clear that we could be facing a similar situation in cultivated plants. Traditionally, growing a plant and improving its production will lead to forced hybridizations" commented Gabaldón.

Being able to "look" at 100 million-year-old eukaryotic genomes will now allow us to deepen our knowledge about genomes as well as about the evolutionary mechanisms that lead to diversifying and acquiring new function. This challenging and innovative work will have major implications on how we interpret the functional and evolutionary consequences of genome duplication. It also highlights the importance of basic research for understanding genomes, evolution and diversity.

<u>Reference paper</u>: Marina Marcet-Houben and Toni Gabaldón "Beyond the whole genome duplication: phylogenetic evidence for an ancient inter-species hybridization in the baker's yeast lineage" PLOS Biology. 7th August 2015.

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