

Step 1 – Build DBG



Step 2 – Identify repeat nodes



Step 3 – Output quantified graph objects (components, complex bubbles, etc...)



Step 4 – Graph object differential analysis on a complex events perspective

Step 1 – Build DBG

- 1/ DBG construction;
- 2/ Get $(k+1)$ -mers (edges) counts per read file;

Step 2 – Identify repeat nodes

- Currently done reference-based;
- Identifies the unitigs that are due to **genomic** repeats;
 - Not ideal, we want to find unitigs corresponding to transcriptomic repeats;
 - Or even further: **repeats that are problematic to the sequenced transcriptome only**;
- Pipeline:
 - 1/ Align unitigs to the genome using STAR;
 - 2/ Intersect the .bam with UCSC's repeat masker track;
 - 3/ Unitigs having overlaps with repeats of at least 20 bases (by default) are identified as repeat nodes;
 - 4/ Unitigs mapped to too many loci and unmapped unitigs are also identified as repeat nodes;
- TODO:
 - Do this in a de-novo way;
 - Use machine learning with several features including beta-value, branching concentration, coverage, sequence entropy (what else?) to help on classifying if a node is repeat-induced or not;
 - Features extraction (for the current ones) is almost done;

Step 3 – Output quantified graph objects (components, complex bubbles, etc...)

- 1/ Output the full repeat-free graph (nodes and quantified edges);
- 2/ Print all the uncompressed components of the graph (nodes and quantified edges);
 - This was based on the hypothesis that removing repeat-induced untigs would disconnect the graph into several components, one for each gene, or gene family;
- 3/ Compress the components of the graph in a sequence-based way;
 - Pretty bad algorithm for now:
 - 1/ List all paths from any source to any target in the component;
 - 2/ Transform all paths in sequences;
 - 3/ Cluster the sequences (sequences with edit distance $<1\%$ are clustered together);
 - 4/ For each cluster, get the most k-mer covered sequence as representative;
 - 5/ Build the DBG of the representative sequences;
 - 6/ Output the compressed components;
 - Edge requantification not yet done;
 - Bad algorithm because in some components with some hundreds nodes, we can have millions paths and this would take too, too much time...
 - Fall back to simpler methods like 4 nodes compression? Iterative approach? Is this really needed?
- 4/ We used to also find and output complex bubbles in each component... But not anymore, I don't remember why...

Step 4 – Graph object differential analysis on a complex events perspective

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