Algorithmic and computational comparison of metagenome assemblers

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ABSTRACT

Assembly of genome sequences of a microbial community is computationally challenging and complex than its single genome counterparts. Keeping in view the volume, diversity and varied abundance of different microbes, number of metagenome assemblers have been developed addressing specific associated computational issues mainly following De Bruijn Graph (DBG) and Overlap Layout Consensus (OLC) approaches. It is very pertinent to understand different computational approaches and issues of metagenomic assembly to further improve them with respect to time and computational resource requirements. Therefore, the main objective of this article is to discuss various metagenomics assemblers with respect to their development addressing major computational issues. Initially the computational perspective of single genome assemblers based on OLC and DBG graph construction approaches was described. This is followed by review of metagenomic assemblers with respect to the algorithm implemented for addressing issues in metagenome assembly. Further, performance of some of the popular metagenome assemblers were empirically evaluated with respect to their run time and memory requirements by taking diversified benchmark metagenomics data at ICAR-IASRI, New Delhi in 2019. It was concluded that performance of assemblers varied considerably on these datasets and there is further need to make an effort to develop new tools or to modify the existing ones using efficient algorithms and data structures.

Key words: Assembler, Algorithm, Computation, Metagenome

Metagenomics is emerging as a promising technology for structural and functional profiling of microbial communities. With the advent of new sequencing techniques, the sequencing of the genome of many environmental and human microbial communities have been done as these were difficult to culture in the laboratory. This information overload has given rise to the development of many tools and software for analysis and assembly of metagenomics data. Sufficient literature is available on assemblers which are capable in assembling sequencing data of single genome, but in case of assembling mixed genomic sequences, i.e. metagenomics data of microbial community of an environment have number of computational issues and challenges. Attempts have been made to analyze the assembly challenges and accordingly reviews were published (El-Metwally et al. 2013, Ghurye et al. 2016). Miller et al. (2010) have compared genome assemblers for single genome based on DBG and OLC approaches. A classification of existing assemblers has been provided considering the common features of these along with complete description of layout and DBG graph theory. Klefogiannis et al. (2013) have compared the memory efficiency of single genome assemblers in terms of memory usage, execution time and quality metrics.

It may be noted that the above reviews were confined to the genomic assemblers capable of assembling sequence data of single species. A comprehensive review of metagenomics assemblers was done by Vollmers et al. (2017) in terms of biological/microbiologist perspective. Recently in a review, methods and tools for metagenome read classification, metageome assembly tools and contents of existing metagenome databases with quality aspects is done (Breitwieser et al. 2017). Therefore, hardly there is any review article which compares metagenomics assemblers from computational and algorithmic perspective. Therefore, in this article an attempt has been made to compare metagenomic assemblers from computational and algorithmic perspective, i.e. (i) variants of DBG and OLC graph construction, (ii) algorithmic approaches and (iii) computational requirements with the help of benchmark datasets. This article will be highly useful to the computational biologists/scientists, scholars and students working in the area of metagenomics and related fields.

Current challenges in assembly of metagenome

Assembly of metagenome consists of three main steps namely (i) sequence cleaning, (ii) fragment assembly and
(iii) Binning/taxonomic assignment. Sequence cleaning deals with the filtering of duplicated, short, low quality, contaminated reads and reads containing ambiguous bases. Fragment assembly of metagenomic sequences is performed following two approaches namely OLC and DBG. De novo /Fragment assembly results in the generation of contigs/scaffolds. Binning/taxonomic assignment is needed to classify/associate these data with the organisms from which they were originated. This process of association between sequence data to their related species is called binning or classification. Existing binning methods can be classified into two categories, namely taxonomy dependent and taxonomy independent. In case of taxonomy dependent methods, the extent of ‘similarity’ of the reads with sequences (in reference databases) or pre-computed models helps in this assignment process. Further, based on the strategy used for comparing sequence reads with existing sequences/ pre-computed models, taxonomy-dependent methods can be further classified into three categories, i.e. alignment-based, composition-based and hybrid methods (Mande et al. 2012). Alignment based algorithms use existing alignment and read mapping algorithms to align the reads to known and characterized genomes. This approach is used in MGRAST, CAMERA, MEGAN, MetaPhyler and MARTA etc. Faster methods based on composition features use GC content, codon usage, oligonucleotide usage patterns for comparing reads to reference databases. Under this method, specific models are built for various genomes using various statistical and data mining approaches. Some binning tools based on this method are Phylophthia, TACOA, Phymm, ClaMS and RAIdphy etc. Hybrid approaches combine the alignment and composition based methods for classification of metagenomic data. In case of taxonomy independent methods, simply group/bin of sequence reads are made from given dataset based on their mutual similarity and it does not involve any database comparison. Tools following this approach are TETRA (Teeling et al. 2004), CompostBin (Chatterji et al. 2008), AbundanceBin (Wu et al. 2011) and MetaCluster (Leung et al. 2011).

Initially, single genome assemblers were used for assembly of metagenomic data but due to computational and biological issues specific to this field, effective and accurate assembly could not be done. Some of the major challenges with assembly of metagenomes are: (i) uncertainty about population size, (ii) composition of species and (iii) noise model of sequencing technology. Also, coverage across species is uneven and affected by the species abundance in the sample. Therefore, an alternative approach was required which may also construct the contigs even for species having low coverage data. It was further observed that the performance of existing DBG based approaches degrade with increase in number of errors in the reads. This demands additional efforts for detecting, correcting and filtering the reads before assembly. Most of existing techniques in this regard implemented in single genome assemblers assumes the low coverage as a criterion to filter out the reads. This is not desirable as well as applicable in case of metagenomics data. Also, identification of repeat region in case of metagenomic data needs to be extended to multiple genomes. Separation of Single Nucleotide Polymorphism (SNPs) is difficult to separate from sequencing errors in metagenome. Due to the above facts, application of single genome assemblers for metagenome assembly leads to the formation of chimeric and shorter contigs. Therefore, number of metagenomic assemblers are developed in due course of time by addressing above issues. In the next section computational developments in DBG and OLC approaches are described briefly.

Computational developments

Assemblers have been developed for assembling metagenomic data generated through various sequencing platforms. The algorithmic approach of different assemblers varies in: (i) ways of handling type of reads (i.e. long reads to short reads), (ii) type of graph construction, (iii) sequencing error correction methods, and (iv) ability to deal with different length of fragments. Mainly, two types of assembling algorithms were developed following OLC and DBG construction approach according to the size of reads. The DBG is a graph data structure suitable for representing the overlap relationship of short read sequences, whereas, OLC is generally used for assembling long read sequences. It was observed that, the overlap graphs work well if there is small number of reads with significant overlap. However, this method is computationally expensive for large genomes. Complexity of pairwise sequence alignment is quadratic in terms of number of reads. Efforts have been made by researchers for improving the computational efficiency of OLC and DBG based approaches.

Developments in computational framework of OLC approach

The first graphical representation of sequence assembly as an overlap graph was introduced by Kececioglu and Myers in 1995. In an overlap graph, a vertex represents a read and an edge represents an overlap. Optimal path through each vertex was used to find the layout of reads to covert to larger fragments of reads. Transitive reduction approach was proposed to reduce the redundancy of the overlaps. In 2000, Celera assembler was developed for best assembly of short-gun reads (Myers et al. 2000). A more accurate and efficient assembly program for better handling of overlap reads using multiprocessors was developed in 2003 (Huang et al. 2003). Further, an efficient and better tool for detection and classification of single nucleotide polymorphisms for expressed sequence tags using OLC approach was developed by Chevreux et al. (2004). Also, Myers in 2005 has proposed a string graph OLC based assembler. In order to apply this approach for short read sequences, FM-index was applied to find overlaps in OLC graph (Simpson and Durbin 2010). This was followed by two faster and efficient overlap finder proposed by Dinh and Rajasekaran (2011) and Gonnella and Kurtz (2012).
Developments in computational framework of DBG approach

DBG based approach is more computationally efficient as compared to OLC approach. The DBG approach also resolves repeat region problem of OLC based approach with polynomial time solution (Pevzner et al., 2001; Pevzner and Tang, 2001). Further, the problem of repeat region is addressed efficiently through double-barrel data or by additional PCR experiments following this approach. A parallel version of DBG decomposition algorithm was given by Bokhari and Sauer (2005). This algorithm made an attempt to resolve some important issues of genome assembly like reads length, orientation, complementarity and errors in a DNA sequence. Further, the computational efficiency was improved through parallelized DBG implementation in C language. Alternative DBG representation has provided better error elimination and sequence resolution by using more efficient algorithms which was implemented in Velvet assembler (Zerbino and Birney, 2008). Velvet was implemented in C and tested on a 64-bit Linux machine. Issue of scaling up the available memory for storing DBG graphs especially for large genomes was addressed by Simpson et al. (2009). For reducing the memory requirements, a new distributed DBG structure using hash tables instead of pointers was proposed which was implemented in an assembler for short sequences named as ABySS. In this, sequence data is partitioned and distributed over a number of nodes in Linux Cluster to run it parallel. Message Passing Interface (MPI) protocol was used for communication between the nodes.

In order to further reduce memory requirements, Li et al. (2010) proposed an approach of recording minimum-information in DBG instead of read locations and pair-end information. Another effort in this direction of reducing large memory space requirements for DBG construction was undertaken by Ye et al. (2011). They proposed sparse DBG structure which requires less memory and space through skipping some intermediate k-mers. This proposed structure also takes care of substitution errors introduced during the sequencing process.

Storing of nodes in a DBG using hash tables and pointers was further modified by Conway and Bromage (2011). They proposed a new storage configuration having more efficient data structure using information theory concepts. In this case, DBG was represented as entropy compressed succinct DBG. This configuration is ten times more memory efficient as compared to the existing methods. Further, Chikhi and Risk (2013) introduced a new, space-efficient representation of the DBG by implicitly encoding this as a Bloom filter. In this case, separate data structure is used to store each of the false and true positive nodes. Besides this, a new data structure was also proposed which stores the information of visited nodes during traversal. Therefore, this structure stores only a subset of k-mers leading to more efficient space utilization. An efficient tool for de novo assembly namely Minia was developed by combining the Bloom filter, critical false positives structure and the above marking structure. The memory requirements of the data structure proposed by Chikhi and Rizk (2013) was further reduced up to 40% through using cascading Bloom filters and storing k-mers in less number of bits (Salikhov et al., 2014). Many popular assemblies of genome and metagenome requires to construct the DBG graph with multiple values of k. This hampers their performance with the increase in genome size. In order to resolve this problem, generalization to DBG has been proposed by Lin and Pevzner (2014) with the introduction of the manifold de Bruijn (M-Bruijn) graph theory. In this, multiple k-mers sizes are considered in single iteration for selection of optimal k-mer size instead of varying k-mer size in each iteration as in Iterative de bruijn Graph Assembly (IDBA) approach.

Boucher et al. (2015) described a method for efficiently representing multiple DBG of different orders in a single succinct data structure. The proposed data structure allows the order of the DBG to be changed on the fly. This technique considerably improves the memory and space requirements of the existing assemblers. Recent technology such as Single Molecule Real Time (SMRT) sequencing platform generates longer but error-prone reads which is being assembled using OLC based approach. Lin et al. (2016) have applied generalized DBG approach to assemble these reads to generate more accurate genome reconstruction using A-Bruijn graph. A-Bruijn graph is developed by exploiting the benefits from both DBG and OLC approaches.

A computational perspective of metagenome assemblers

Assembly of metagenomic data poses many challenges which cannot be addressed by existing single genome assemblers. Some of the existing metagenomic assemblers are Genovo, Xgenovo, Meta-IDBA, Ray Meta, IDBA-UD, MetaVelvet, MetaVelvet-SL, MegaHit, MegaHit1.0, PRICE, Omega etc. There is a need to understand the algorithmic approaches of existing assemblers to develop more efficient algorithm to address the computational challenges. Therefore, in this section the important algorithms used by above assemblers are given in brief.

OLC based approaches for metagenome assembly

Genovo is a de novo assembler developed by Laserson et al. (2011) for assembling metagenome. In order to discover the population structure, sample is initially subjected to Chinese restaurant process to generate prior probabilities to classify reads. This process accounts for the unknown number of genomes in the sample prior to their assembly. Likely assemblies are made by applying a series of hill-climbing algorithms steps iteratively until convergence. This Bayesian based approach offers better sensitivity for assembly in highly diverse environment. Genovo’s reconstruction procedures are comparatively more efficient than their competitive procedure which consequently yields a higher assembly score. Genovo was further improved as Xgenovo by incorporating paired-end information to get higher quality assemblies (Afshahayati et al., 2013).

Haider et al. (2014) proposed an assembler named Omega (Overlap Graph Metagenome Assembler) for
assembling and scaffolding metagenomics data obtained from Illumina sequencing platform at a deeper coverage. In this, hash table is used as data structure to find overlaps between reads. Also, simplification to the overlap graph was done by deleting transitive edges and trimming short branches. Further, mate-pair information is used to get scaffolds after merging contigs. It is reported that Omega outperforms its competitors in terms of computing time with respect to existing OLC based Celera assembler on the MiSeq dataset. Also, it was found that the performance of Omega is at par on a HiSeq 100-bp dataset and superior on a MiSeq 300-bp dataset when compared to DBG based popular assemblers such as SOAPdenovo, IDBA-UD and MetaVelvet.

DBG based approaches for metagenome assembly

The Meta-IDBA algorithm was proposed by Peng et al. (2011) for assembling reads of metagenomic data for handling the issue of polymorphism. This assembler is based on the biological observation that the genome of sub species of same species share more common regions than the genome of sub-species from different species. Therefore, the major task is partitioning of the DBG graph into different components such that each component represents a consensus contig of a species. The process of DBG partition is based on the concept that if, path of a given length from one k-mer can be traversed to another k-mer in DBG starting from each of its out-going edges then two k-mers are assumed to be originated from the same species. Otherwise, the given two k-mers may belong to genomes of different species/sub-species and should be separated. Further, larger components are broken down into the smaller components and each component represents a consensus contig of sub species from a single species. Paired-end information was used by Meta-IDBA to combine the contigs into scaffolds. This was followed by performing multiple alignment of each component with small variants of the genomes of sub-species. Finally, consensus sequence was used to represent genome of one species. Meta-IDBA was shown by Peng et al. (2011) to construct longer contigs with accuracy similar to Velvet and Abyss for different metagenomic datasets. One of the major limitation of Meta-IDBA is that it cannot reconstruct the contigs of each single sub-species due to presence of common regions in the genomes of sub-species. Another, drawback is that the accurate separation of components in some case for some species is not possible. Also, in some situations many small components may be obtained after graph partitioning like presence of false positive edges, more number of species (>1000). It may be noted that Meta-IDBA does not depend upon the coverage for the construction of DBG graph, so the change in abundance ratio does not affect its performance. Subsequently, Peng et al. (2012) developed an algorithm (IDBA-UD) based on DBG approach considering uneven sequencing depths for metagenomic assembly. In this assembler, instead of using single threshold value for k-mer removal multiple values are used depending on the sequencing depth. Also, the branching problem of low-depth short repeat regions is solved with the help of local assembly using paired-end information. In order to speed up the process, an error correction step is incorporated to correct reads of high-depth regions which can be aligned to high confident contigs.

MetaVelvet assembler is the extension of single genome assembler named Velvet for the assembly of genome of various species in microbial communities. It was developed by Namiki et al. (2012) for mixed short read sequences of multiple species. The process of assembly in MetaVelvet is done in four phases. In the first phase, it constructs the DBG from the input reads. In second phase, it detects the multiple peaks based on empirical distribution of node coverage assuming that the expected frequencies of k-mer occurrences follow a Poisson distribution in a single-genome assembly and the expected k-mer frequencies in metagenome assembly were shown to follow a mixture of Poisson distributions. MetaVelvet distinguishes a sub-graph composed of nodes belonging to a same peak from the other sub-graphs in the main DBG. In phase three, it identifies shared nodes (chimeric nodes) between two sub-graphs and disconnects two sub-graphs by separating the shared nodes with one in and out degree. MetaVelvet uses the number of incoming and outgoing nodes to classify a chimeric node constructed due to repeats in the same species or due to similar sequence in different species. Lastly in phase four, it builds contigs and scaffolds based on the decomposed subgraphs using Velvet functions.

The major issue in MetaVelvet is the correct identification of chimeric nodes as this prevents generation of longer contigs and scaffolds due to their misclassification among species. In order to address this problem a new tool called MetaVelvet-SL was developed (Afiahayati et al. 2015). The performance of this tool is improved by applying supervised machine learning for classification of chimeric node. The first step of MetaVelvet-SL is the construction of DBG followed by identification of chimeric candidate nodes along with their features as a second step. Further, a Support Vector Machine is used for learning the classification model based on features extracted from candidate nodes. It also prepares training sample using MetaPhlAn based on prior knowledge about the taxonomic profile of the target microbial community. This training samples are generated using various profiling methods such as: (i) collection of reference genome of closely related species, (ii) collection of genomes to simulate sequence reads, and (iii) alignment of nodes with reference genome. Finally, species for each node is identified. The major drawback of low accuracy and sensitivity in detecting the chimeric nodes in the DBG generated by MetaVelvet is overcome by this approach. Further, DBG is decomposed into sub-graphs for identification of unique nodes in each sub-graph. Finally, MetaVelvet is used for generation of contigs and scaffolds.

The problem of assembling metagenome of species having large genomes was addressed by Li et al. (2015) by developing MEGAHIT based on succinct DBG (SDBG), i.e.
COMPARISON OF METAGENOME ASSEMBLERS

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Recently, an assembler named as metaSPAdes has been developed which is useful in case of single cell and highly polymorphic diploid genomes (Nurk et al. 2017). This tool is useful in efficient assembling of metagenomic data from application of single cell (Kashtan et al. 2014) and TruSeq Synthetic Long Reads, i.e. TSLRs (Sharon et al. 2015) technologies. Therefore, it is capable of assembling vast micro diversity strains which shares genomic sequences having variations due to mutations, insertions of mobile elements, genome re-arrangements, and gene transfer. metaSPAdes initially constructs DBG using SPAdes and simplifies using various graph simplification algorithms. The main objective of metaSPAdes is to generate consensus assembly of strain mixture. Genomic differences between related strains and other sequencing errors leads to formation of large buldges and longer tips which are detected and removed following the algorithmic approaches suggested by Pevzner et al. (2004), Zerbino and Birney (2008). In order to preserve the information about rare strains, this assembler first detects the filigree edges formed due to rare strain variants and then disconnect them from their predominating vertices in the assembly graph instead of removing. Further, metaSPAdes algorithm also resolves repeats through application of exSPAnder module of SPAdes after modification in its decision rules by making it more conservative (Prijibelski et al. 2014, Vasilinetc et al. 2015, Antipov et al. 2016). Subsequently, undesirable path extensions are filtered out using coverage estimate of region being constructed. The various computational features of metagenomic assemblers are provided in Table 1.

Empirical evaluation and discussion

Computational comparison of recent assemblers namely MEGAHIT, Meta-IDA, MetaVelvet and metaSPAdes has been done for assessing the memory and time complexity on a sever machine with four processors (each with 8 cores), 256 GB RAM and hard disk of 3 TB approximately. For the purpose of performance evaluation of these assemblers, two benchmark datasets (Vollmers et al. 2017) representing bacterial community of varying complexity are selected and downloaded from NCBI. The dataset samples are chosen from Kelp BioFilm (KBF) and Marburg Forest Soil (MFS) samples. KBF dataset comprises low community diversity but high read redundancy, whereas, MFS dataset is highly community diversified but is with low read redundancy. Raw sequences from chosen dataset are pre-processed for adapter clipping, quality and length trimming of sequences using

![Running time of different Metagenome assemblers](image)

Fig 1 Comparison of time of assembly for selected assemblers for KBF and MFS datasets.
Table 1  Computational features of metagenome assemblers

<table>
<thead>
<tr>
<th>Name of assembler</th>
<th>Data structure</th>
<th>Novelty in approach</th>
<th>Parallelism</th>
<th>Limitation</th>
<th>Language developed</th>
<th>Read size/platform</th>
</tr>
</thead>
<tbody>
<tr>
<td>Genovo</td>
<td>OLC</td>
<td>Extracts more information from datasets and thus is suitable for low abundance</td>
<td>Single Server</td>
<td>Assessment with Illumina dataset</td>
<td>C++</td>
<td>454 sequencing</td>
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<td></td>
<td></td>
<td>sequences.</td>
<td></td>
<td>-Quality of assembly depends on the quality of DBG.</td>
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<td></td>
<td></td>
<td>-Change in abundance ratio does not affect the performance</td>
<td></td>
<td>-Many species leads to many false positive edges which leads to many small</td>
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<td></td>
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<td>components.</td>
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<tr>
<td>Meta-IDBA</td>
<td>DBG</td>
<td>-Handled the polymorphism in similar species.</td>
<td>Single Server</td>
<td></td>
<td>C++</td>
<td>NGS</td>
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<tr>
<td></td>
<td></td>
<td>-Change in abundance ratio does not affect the performance</td>
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<tr>
<td>MetaVelvet</td>
<td>DBG</td>
<td>Decomposition of DGB graph constructed from the mixed short reads into sub-</td>
<td>Single Server</td>
<td>Accuracy identification of chimeric reads</td>
<td>C++</td>
<td>Short Read</td>
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<tr>
<td>PRICE</td>
<td>DBG</td>
<td>Contig length extension using paired read</td>
<td>Multi-threading on</td>
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<td></td>
<td></td>
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<td>multicores single CPU.</td>
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<tr>
<td>XGenovo</td>
<td>OLC</td>
<td>-Additional parameter for solving chimera contig case</td>
<td>Single Server</td>
<td>-Implementation of probabilistic model with DBG graph</td>
<td>C++</td>
<td>454 paired end</td>
</tr>
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<td></td>
<td></td>
<td>-Use of modified sampling process for the location of read in a contig</td>
<td></td>
<td>-high computational cost</td>
<td></td>
<td>reads</td>
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<tr>
<td>Omega</td>
<td>OLC</td>
<td>Uses less computing time as compared to existing OLC based assemblers at higher</td>
<td>Multi-threaded</td>
<td>Less efficient as compared to DBG based assemblers</td>
<td>C++</td>
<td>Illumina Long</td>
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<td></td>
<td></td>
<td>coverage</td>
<td>and multiprocess</td>
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<td>Reads</td>
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<td></td>
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<td></td>
<td>distributed memory</td>
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<tr>
<td>MetaVelvet-SL</td>
<td>DBG</td>
<td>Better detection of chimeric nodes by supervised learning.</td>
<td>Single Server</td>
<td>Classification model for chimeric reads may be improved.</td>
<td>C++</td>
<td>Short Read</td>
</tr>
<tr>
<td>MEGAHIT</td>
<td>Succinct</td>
<td>-No need for normalization and partitioning</td>
<td>GPU based and 3-5 times</td>
<td>Parallelization is restricted by small size of on-board GPU memory.</td>
<td>C++</td>
<td>NGS</td>
</tr>
<tr>
<td>v1.0</td>
<td>DBG</td>
<td>-Efficient dynamic removal of edges</td>
<td></td>
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<tr>
<td>metaSPAdes</td>
<td>DBG</td>
<td>-Long bubble merging</td>
<td>Runs on Single Sever</td>
<td></td>
<td>C++</td>
<td>Single cell and</td>
</tr>
<tr>
<td></td>
<td></td>
<td>-Faster and uses less memory than MEGAHIT</td>
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<td></td>
<td></td>
<td>TSLRs</td>
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<td></td>
<td></td>
<td>Detection and removal of large bulges and longer tips</td>
<td>Single Server</td>
<td>Incorporation of recent high quality jumping mate-pair libraries</td>
<td>C++</td>
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</table>

Trimmomatic tool. Running time and peak memory usage of selected assemblers for complete assembly process has been observed and shown in Fig 1 and Fig 2, respectively.

Following points can be observed from this empirical analysis

The assembly of KBF dataset takes less time as compared to MFS irrespective of assemblers. However, peak memory usage for KBF is more in case of MEGAHIT and Meta-IDBA, whereas it is less in case of metaSPAdes and Meta-Velvet.

The performance of MetaVelvet with respect to running time is best which is followed by MEGAHIT, whereas the performance of Meta-IDBA is comparable with metaSPAdes specially for KBF dataset. However, for MFS dataset, Meta-IDBA performs poorly in terms of running time than other assemblers.

The peak memory requirement for MEGAHIT is the least followed by metaSPAdes, MetaVelvet and Meta-IDBA. The runtime requirement for KBF dataset is less in
comparison to MFS dataset across all assemblers is less due to the fact that this dataset is less diverse and have high read redundancy as comparison to MFS dataset. The runtime requirement for metagenome assembly for both datasets is less in case of MetaVelvet, as it partition the graph into sub-graph with the help of statistical distribution of k-mers. Although, Meta-IDA also makes sub-graphs from DBG graph based on identification of particular graph pattern of k-mers which may be computationally less efficient.

The performance of MEGAHIT in terms of runtime assembly is likely to be much better than MetaVelvet on GPU machine. Run time performance of metaSPAdes is on expected lines, as this assembler is specifically developed for single cell and sequencing data based on TSLR technology. Peak memory requirements for assembly in MEGAHIT is least due to the reason that it dynamically removes the edges and normalization as well as partitioning of graph is not required. This has been followed by less memory requirement of metaSPAdes which may be due to removal of large bulges and longer tips during the construction of graph.

Conclusion and future research directions

In general, it may be difficult to identify the computationally best assembler from the existing assemblers as their performance may vary based on characteristic of dataset as well as architecture of the computational resources available for metagenome assembly. However, from this study it can be concluded that the performance of MetaVelvet is considerably better than its counterparts. Although, the performance of MEGAHIT is likely to be better on GPU environment. The computational performance of metagenome assembly process can be further improved in future by (i) computationally efficient graph partitioning to handle highly diversified multi-genomic data, (ii) application of better time and memory management data structures, (iii) implementation of novel parallelization approaches and Big Data technologies, and (iv) incorporating recently introduced auxiliary information such as Nextra Mate Pair libraries with existing recent TSLR technology in existing metagenome assemblers.

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