Xinghua Lou, Bernhard X. Kausler, Ullrich Koethe, Fred A. Hamprecht HCI, IWR, University of Heidelberg, Heidelberg 69115, Germany December 24, 2012

# BOT: A C++ Library for Biomedical Object Tracking Design Document and User Guide

1 Abstract

Biomedical Object Tracking (BOT) is a C++ library for tracking a massive number of objects from time lapse image sequences. It features (I) object association based pairwise tracking, (II) interactive structured learning for optimal feature parametrization, (III) generic and extensible feature design, and (IV) configurable workflow for different applications (e.g. tracking cells and tracking worm). In this report, we provide the technical details on the implementation of the aforementioned algorithms, features and detailed guide to installing and using this library.

#### 1 Introduction

Reliable multiple object tracking is fundamental to the interpretation of time-lapse microscopic image data in biomedical research. Recently, object association based on mathematical programming for object tracking has drawn much attention due to its high efficiency in solving large-scale, complex tracking problems such as lineage-tree reconstruction [2] and cell culture study [4, 3]. We recently developed structured learning for cell tracking, an approach for learning the optimal parameterization of the high-dimensional association features from a training set of ground truth associations [1]. Experimental results show that this approach not only provides superior tracking performance but also enables biologists to contribute their expertise in an intuitive fashion.

Despite the popular use of object association in many tracking problems, there has not been an open implementation of this method, let alone a well-designed library with high extensibility and standard interfaces. We attempt to address this situation by introducing BOT, a extensible C++ library for biomedical object tracking. Though initially designed for cell tracking, we hope to make BOT a generic algorithmic library adaptable to various biomedical tracking applications. Briefly, BOT features:

- Solver An efficient pairwise object association solver based on integer linear programming (ILP);
- 23 Feature A rich set of generic features and extensible feature design using factory pattern;
- Learning Structured learning for high-dimensional feature parametrization;
- Diversity Configurable workflow for supporting diverse applications;
- GUI A user-friendly GUI by integration with the Interactive Learning and Segmentation Toolkit (ilastik, http://www.ilastik.org/) [5].

# 28 2 Implementation Details

## 29 2.1 Prerequisites

BOT has two prerequisites: data and solver system.

Regarding data, BOT assumes that the objects of interest have been segmented or detected from the raw images. The corresponding segmentation or detection algorithms are usually problem-specific. The result is a labeling, in which BOT accepts value zero as background and considers all pixels with identical label (other than zero) as a single object.

Regarding solver systems, BOT relies on at least an integer linear programming (ILP) solver for predicting the tracking. To perform structured learning for parameter optimization, quadratic programming (QP) 36 or linear programming (LP) solver is required. By default, BOT uses IBM ILOG CPLEX<sup>1</sup> which provides 37 solvers for ILP, QP and LP problems. It is nevertheless possible and easy to interface BOT with alternative solvers such as lpsolve<sup>2</sup> and Gurobi<sup>3</sup>. More details on installing CPLEX and on using customized solver systems will follow in Section 3.1.2 and Section 2.5.3, respectively.

#### Concepts 2.2 41

31

33

51

61

### 2.2.1 Object, Singlet and Multiplet

An object literally refers to an object of interest whose behavior is to be tracked. Object is an abstract 43 concept and has two concrete instances: singlet and multiplet. A singlet is exactly a segment or detection which consists of pixels with the same label (nonzero). It represents the very rudimentary entity but is 45 not sufficient for explaining sophisticated biomedical events. We therefore introduce multiplet which is a 46 combination of several singlets (usually restricted to a neighborhood system). Multiplets capture events in which more than one singlet are involved, such as the two daughter cells from a cell division and the 48 multiple parts from an over-segmented worm. In its current implementation, BOT only considers multiplet 49 consisting of two singlets, see Section 2.7 for more details.

In BOT, class Object is the base class for class Singlet and class Multiplet. It employs a sparse representation (i.e. a point cloud) of the pixels within the object, i.e. their coordinates, intensity values and labels. It also stores the object features (see Section 2.2.3) as a vector of Matrix2D (i.e. std::vector<Matrix2D >). Class Singlet and class Multiplet instantiate class Object. To generate all singlets and multiplets in a frame with configurable neighborhood constraints, use class SingletGenerator and class MultipletGenerator, respectively. These generators assign an id to each singlet/multiplet that is unique within this frame.

#### 2.2.2 Event and Hypothesis 57

An event is one particular type of association between at least two and possibly more objects respective-58 ly from a pair of neighboring frames. An event has a particular biological or technical interpretation. For example, cell division is a biological event and the involved objects are the father cell from one frame and the two daughter cells from the next frame. For another example, a sudden over-segmentation (i.e. split) is an event cause by a technical limitation. 62

Multiple events can be defined to handle more complex tracking problems. For example of cell tracking with imperfect segmentation, we can define six events as in Table 1. The first four events express the usual cell behaviors and the last two capture segmentation errors, i.e. split for over-segmentation and merge for under-segmentation. In the header of the table, Pairing specifies the types of objects involved, e.g.  $1 \rightarrow 1$  for singlet to singlet,  $1 \to 2$  for singlet to multiplet, and  $1 \to \emptyset$  for singlet to *nothing* (considered as a special type

<sup>&</sup>lt;sup>1</sup>http://www-01.ibm.com/software/integration/optimization/cplex-optimizer/

<sup>&</sup>lt;sup>2</sup>http://sourceforge.net/projects/lpsolve

<sup>&</sup>lt;sup>3</sup>http://www.gurobi.com

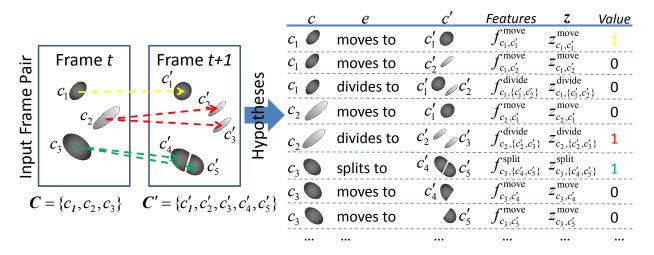


Figure 1: Toy example: two frames of objects and a list of the possible associations as hypotheses. One particular interpretation of the scene is indicated by colored arrows (left) or equivalently by a configuration of binary indicator variables z (rightmost column in table).

of "object"). BOT uses class EventConfiguration to import the definition of events from an INI file<sup>4</sup> and manages them. Please refer to Section 2.4 for more details on configuring the workflow and on preparing the INI file.

Name	Pairing	Feature 1	Feature 2	Feature
Move	$1 \rightarrow 1$	Change of size	Spatial displacement	• • •
Division	$1 \rightarrow 2$	Angle pattern	Father cell intensity	
Appearance	$\emptyset \to 1$	Distance to the border	Overlap with the border	
Disappearance	$1 \to \emptyset$	Distance to the border	Overlap with the border	
Split	$1 \rightarrow 2$	Shape compactness	Mass evenness	
Merge	$2 \rightarrow 1$	Shape compactness	Mass evenness	

Table 1: Event definition for cell tracking (six types of events).

Given the definition of events, hypotheses are essentially their instantiations. As shown in the toy example in Fig. 1, hypotheses are generated throughout the entire sequence for each type of event, subject to a pre-defined neighborhood constraints including the k nearest neighbor and the spatial distance threshold. BOT uses class HypothesisSpace to generate and manage all hypotheses throughout the entire sequence. For a particular pair of frames, its hypotheses and the corresponding joint features (see Section 2.2.3) are represented using class FramePair.

#### 2.2.3 Object Features and Joint Features

78

79

80

81

An object feature characterizes the object only and is extracted from the object itself (e.g. position, size and principal components) or together with the global context information (e.g. distance to the border and overlap with the border). Object features are normally generic and primitive.

A joint feature, on the other hand, characterizes a hypothetical association. Extracted from the object features, it is a measure of the compatibility of the raw information and the hypothetical association. For example, spatial displacement is a joint feature computed from the position of the two objects and it is

<sup>&</sup>lt;sup>4</sup>http://en.wikipedia.org/wiki/INI\_file

intended to respect the speed limitation of movement (e.g. cell migration). Appropriate distance measure should be selected according to the object feature, such as using earth mover's distance (EMD) for comparing two intensity histograms (see class MeasureEarthMoversDistance). Note that joint features are not always extracted from both objects in the hypothesis. For example of cell division, the father cell usually appears brighter than the usually ones so that we directly take the mean intensity of the father cell as a "joint feature" for the division event.

BOT has already included a rich set of generic object features and a handful of joint features. Furthermore, it adopts the factory method pattern to allow for easy extension of the feature set and dynamic feature loading. More details are provided in Section 2.5. To avoid possible confusion of these two concepts, in BOT we refer to the joint feature as *measure* since it essentially provides a measure of the compatibility. We refer to the object feature simply as *feature*. Class JointFeatureExtractor extracts the object features and class MeasureExtractor extracts the measures.

### 2.3 Data Structure and Representation

#### 7 2.3.1 Primitive Data Types

90

92

93

108

To enforce consistency throughout the library, we always use int32 for indices and object ids. Furthermore, we define Matrix2D, a data type for representing the raw images, segmentations and feature matrices. The advantage of such representation is that we can use powerful linear algebra operations to facilitate the feature computation such as those in VIGRA (see namespace vigra::linalg).

```
// data type definition for indices, ids, etc.

typedef int int32;

// data type definition for images, segmentations, features, etc.

typedef double MatrixElem;

typedef vigra::MultiArray < 2, MatrixElem > Matrix2D;
```

#### 2.3.2 Objects, Hypotheses and Features

We use point cloud to sparsely represent the raw pixels that belongs to an object such as their coordinates, intensity values and labels. The id (int32 type) of a singlet is unique within a frame. The same applies to multiplet as well.

```
class Object {
112
113
   protected:
114
        /* id of this object */
115
        int32 id_;
116
117
        /* 2D matrix of the coordinates of the pixels */
118
        Matrix2D pixels_;
119
120
        /* 2D matrix of the intensity values of the pixels */
121
        Matrix2D values_;
122
```

```
/* 2D matrix of the labels of the pixels */
Matrix2D labels_;

126 };
```

A hypothesis is represented by a pair of ids of the objects involved and a list of hypotheses is thus a vector of such pairs:

```
typedef std::vector < std::pair < int 32 >> Hypotheses;
```

Features are always represented using type Matrix2D.

#### 2.4 Configurable Workflow

#### 132 2.4.1 Event Definition

130

148

149

150

151

152

154

155

157

158

BOT allows the definition of the events to be tailored to a specific problem. For example, division is a particular important event for cell culture study but not needed for worm tracking. As already shown in Table 1, an event definition consists of a name, a pairing and a list of joint features. The event's name must be unique. The feature setting will be introduced immediately in Section 2.4.2. Class EventConfiguration imports the event definitions from an INI file and manages them, such as:

```
[Division]
Pairing = 1 2
Feature1 = ObjectFeatureIntensitySum MeasureNormalizedEuclideanDistance -0.4680
Feature2 = ObjectFeatureCenters MeasureAnglePattern -1.3055
Feature3 = ObjectFeatureIntensityMean MeasureAppointmentLeft 0.9591
Feature4 = ObjectFeatureEccentricity MeasureAppointmentLeft -0.9441
Feature5 = ObjectFeatureVolumeEvenness MeasureAppointmentRight 1.6322
Feature6 = ObjectFeatureShapeCompactness MeasureAppointmentRight -0.7221
Feature7 = ObjectFeatureMassEvenness MeasureAppointmentRight -0.8504
```

#### 147 2.4.2 Dynamic Feature Loading

Each event definition should be associated with a list of joint features (or measures). Each measure consists of three elements: the underlying object feature, the measure on this object feature and the weight. The object feature can be an instance of any class with a name similar to ''ObjectFeature\*\*\*'' (except class ObjectFeatureFactory and class ObjectFeatureExtractor). The same holds for the measures, that is ''Measure\*\*\*'' (again, except class MeasureFactory and class MeasureExtractor). The weight can be manually set or learned from some training examples (see Section 2.6 for more details).

BOT adopts the factory method pattern<sup>5</sup> to dynamically create extractors by their class names. Two respective factories are implemented, namely class <code>ObjectFeatureFactory</code> for object features and class <code>MeasureFactory</code> for measures. Class <code>ObjectFeatureExtractor</code> wraps the necessary functionalities for extracting multiple object features, just like class <code>MeasureExtractor</code> for measures. Only a list of class names are required as inputs. More details on the extensibility of this factory method pattern will follow in Section 2.5.

<sup>&</sup>lt;sup>5</sup>http://en.wikipedia.org/wiki/Factory\_method\_pattern

#### 2.5 Extensibility

#### 2.5.1 Object Features

161

163

164

188

189

191

192

BOT has already included a rich set of generic object features but we encourage the developers to design new features that better suit their applications, and it can be efficiently accomplished. Following the factory method pattern, an virtual class ObjectFeatureFactory serves as the base class for object features, who has several virtual functions that need be implemented in the derived class:

```
class ObjectFeatureFactory {
   public:
167
       /*! Default constructor
168
169
         */
170
       ObjectFeatureFactory() {};
172
       /*! A virtual function that extract the object feature
173
            @param feature_mat The object feature to be returned
174
            @param obj The input object
175
            @param context The global context
176
         */
       virtual void extract (Matrix2D& feature_mat, const Object& obj,
178
            const Context& context) = 0;
179
180
       /*! Return the shape (size) of the feature matrix
181
            @param dim The input data dimension
182
            @return A Matrix2D:: difference_type object as the shape
         */
184
       virtual Matrix2D:: difference_type shape(int dim = 2) = 0;
185
187
```

To add a new object feature, one simply has to

- 1. derive a new class from ObjectFeatureFactory and implement the virtual functions;
- 2. implement a static function getClassName() that returns an std::string as an identifier of this class;
- 3. in the source file ObjectFeatureFactory.cxx, add your newly implemented class in the following function (within the big if-else block) that identifies the target feature class by std::string, creates an instance of the corresponding class and returns a pointer to the instance.

```
ObjectFeatureFactory *ObjectFeatureFactory::make(const std::string& name) {
ObjectFeatureFactory *fea = 0;

if (name.compare(ObjectFeatureVolume::getClassName()) == 0) {

fea = new ObjectFeatureVolume();

}
else if (name.compare(ObjectFeaturePosition::getClassName()) == 0) {
```

Furthermore, class ObjectFeatureExtractor creates and manages multiple object feature extractors such as deleting the extractors to avoid memory leek when necessary.

#### 206 2.5.2 Joint Features

The procedure for extending the joint features (measures) is very much alike. The relevant classes/files are class MeasureFactory, class MeasureExtractor and file MeasureFactory.cxx.

#### 2.5.3 Solver System

210

212

213

As introduced in Section 2.1, BOT allows for customized solver systems. To use one's own solver system, simply derive a new class from class SolverSystem and implement its two virtual functions for solving ILP and QP problems. These virtual functions are very much standardized interfaces for mathematical programming and are compatible with most state-of-the-art solver systems.

```
class SolverSystem {
   public:
215
            Solve a binary integer linear programminig (binary ilp) problem with
216
            given equality and inequality constraints. In particular, it solves
217
            a problem as follows:
218
                    max
                             f' * x
219
                    s . t .
                             Aineq*x <=
                                          bineg
                             Aeq*x
                                          beg
                                      =
221
                             each variable in x is binary
222
            @param f The coefficient of the objective function
            @param Aineq The inequality constraints: Aineq *x \le bineq
224
            @param bineq The inequality constraints: Aineq *x \le bineq
225
            @param Aeq The equality constraints: Aineq *x = bineq
226
            @param beg The equality constraints: Aineg * x = bineg
227
            @param x0 The initial solution
228
            @param x The solution
229
            @param msg The return message
230
231
        virtual std::string solve_bilp(
            const Matrix2D& f,
233
            const Matrix2D& Aineq, const Matrix2D& bineq,
234
            const Matrix2D& Aeq, const Matrix2D& beq,
            const Matrix2D& x0, Matrix2D& x) const = 0;
236
237
       /*! Solve a quadaratic programminig (qp) problem with given equality and
```

```
inequality constraints and bounds. In particular, it solve:
239
                         0.5 * x' * H * x + f * x
                min
240
                st.
                         Aineg*x <= bineg
241
                         Aeq*x
                                   = beq
242
                         lb \ll x \ll ub
243
            @param H Double matrix for objective function (quadratic term)
244
            @param f Double matrix (vector) for objective function (linear term)
245
            @param Aineq The inequality constraints: Aineq *x \le bineq
246
            @param bineq The inequality constraints: Aineq *x \le bineq
247
            @param Aeq The equality constraints: Aineq *x = bineq
248
            @param beg The equality constraints: Aineg * x = bineg
            @param lb The lower bound
250
            @param ub The upper bound
251
            @param x0 The initial solution
            @param x The solution
253
            @return A std::string as the message of the solution status
254
255
        virtual std::string solve_qp(
256
            const Matrix2D& H, const Matrix2D& f,
257
            const Matrix2D& Aineq, const Matrix2D& bineq,
            const Matrix2D& Aeq, const Matrix2D& beq,
259
            const Matrix2D& lb, const Matrix2D& ub,
260
            const Matrix2D& x0, Matrix2D& x) const = 0;
261
   };
262
```

#### 2.6 Tracking Prediction and Parameter Learning

#### 264 2.6.1 Methodology

263

275

To learn more about the underlying methodology of tracking prediction and structured learning for parameter optimization, please refer to our paper [1]. In BOT, they are implemented in class TrackingPredictor and class TrackingLearner, respectively. To learn more, simply follow the example in Section 3.3.

#### 2.6.2 Two Representations of the Tracking Result

We use different representations of tracking result for computation and storage. For computation, the tracking result is merely a matrix of binary values (technically a vector) in which one means the corresponding hypothesis is accepted and zero otherwise. This representation is more of an intermediate solution of the ILP formulation or an input to the learning procedure. We therefore refer to it as Solution:

```
// a vector of ILP solutions (for multiple events)
typedef std::vector<Matrix2D > Solution;
```

Obviously, Solution is dependent on the hypothesis space and is therefore not suitable for storage. This is simply because one usually does not want to store the entire hypothesis space and the result should

be fully interpretable using the images and segmentations as its raw information. We therefore use another representation termed LabelAssociation which directly stores the labels from the segmentation or detection in a pair of matrices as the source and the target:

```
// associating the segmentation labels
struct LabelAssociation {
    std::string name;
    Matrix2D source;
    Matrix2D target;
};

// a vector of label associations (for multiple events)
typedef std::vector<LabelAssociation > LabelAssociations;
```

#### 289 2.7 Limitations

290

291

292

293

294

300

301

302

306

310

We declare two major methodological limitations of BOT:

- 1. BOT is currently restricted to pairwise object association. We are now investigating a global object association approach that hopefully will bring better tracking performance.
- 2. A multiplet is limited to only two singlets. We are investigating possible improvement such as replacing the singlet/multiplet generation with hierarchical segmentation.

#### ₃₃ 3 User Guide

#### 3.1 Installation and Compilation

BOT and several of its dependencies use CMake<sup>6</sup> as the building system. Make sure CMake is correctly install (minimum version 2.6) before processing any further.

#### 299 3.1.1 Install and Compile Dependencies

- 1. Download and install VIGRA from http://hci.iwr.uni-heidelberg.de/vigra/.
  - The installation instructions can be found at
  - http://hci.iwr.uni-heidelberg.de/vigra/doc/vigra/Installation.html.
- 2. Download and install HDF5 from http://www.hdfgroup.org/ftp/HDF5/current/src/. The installation instructions can be found inside the package.
  - If you don't want to compile from source code, you may also use the pre-compiled binary package that is available here: http://www.hdfgroup.org/HDF5/release/obtain5.html.
    - Or, you can choose to use .tiff format instead. If so, download and install libtiff. Instructions can be found here: http://www.libtiff.org/. For windows, a pre-compiled binary package for Win32 is available at http://gnuwin32.sourceforge.net/packages/tiff.htm.

<sup>&</sup>lt;sup>6</sup>http://www.cmake.org/

```
🧬 xlou@LouVB: ~/projects/BOT/build
                                                       Page 1 of 1
 CMAKE_BUILD_TYPE
 CMAKE INSTALL PREFIX
                                    /usr/local
 CPLEX BASE PATH
                                   /home/xlou/local/CPLEX_Studio125
 CPLEX_LIBFORMAT
                                   static pic
 CPLEX_SYSTEM
                                   x86-64_sles10_4.1
 HDF5_INSTALL_PATH
 LIBTIFF_INSTALL_PATH
                                   /home/xlou/local/tiff
                                   OFF
USE HDF5
                                   ON
USE TIFF
 VIGRA_INSTALL_PATH
                                   /home/xlou/local/vigra
WITH_TESTS
                                   OFF
CMAKE_BUILD_TYPE: Choose the type of build, options are: None(CMAKE_CXX_FLAGS or CMAKE_C_FLAGS us
Press [enter] to edit option
                                                                                 CMake Version 2.8.9
Press [c] to configure
 ress [h] for help
                              Press [q] to quit without generating
Press [t] to toggle advanced mode (Currently Off)
```

Figure 2: Configuration using ccmake GUI.

3. To access IBM ILOG CPLEX, a license is required. Detailed guidelines can be found at http://www-01.ibm.com/software/integration/optimization/cplex-optimizer/.

You need a valid license to run CPLEX. For academic organization, you can check out this link: http://www-01.ibm.com/software/websphere/products/optimization/academic-initiative/.

#### 315 3.1.2 Install and Compile BOT

316

- 1. Download the BOT source code from https://github.com/xlou/BOT/.
- 2. Open a console, go to the root directory of BOT and run command **ccmake** ..
- 3. Specify variables in Table 2. The values for CPLEX\_SYSTEM and CPLEX\_LIBFORMAT can be found under [CPLEX root]/cplex/lib/. For example of the x86-64 architecture, the CPLEX library locates at [CPLEX root]/cplex/lib/x86-64\_sles10\_4.1/static\_pic/, which gives x86-64\_sles10\_4.1 to CPLEX\_SYSTEM and static\_pic to CPLEX\_LIBFORMAT. An example is shown in Fig. 2.
- 4. Download a sample dataset using this link:
  http://hci.iwr.uni-heidelberg.de/Staff/xlou/export/dcelliq-sequence-training.h5.
  This file is too huge to be hosted on github. Put the file in ./data.
- 5. Run make. After the compilation, two applications will be built: ./app/TrackingTrainer for training and ./app/TrackingPredictor for prediction.
- For example, you can try the training of the structured model for cell tracking by calling:

  ./apps/TrackingTrainer ./data/dcelliq-sequence-training.h5 ./data/event-configuration-cell.ini

  Training on this sample dataset usually takes 18 iterations to converge and the resulting learned feature weights will be displayed.

Variable	Description	
CPLEX_BASE_PATH	The root path of CPLEX.	
CPLEX_LIBFORMAT	The library format of the CPLEX library.	
CPLEX_SYSTEM	The system type of the CPLEX library.	
VIGRA_INSTALL_PATH	The root path of VIGRA.	
USE_TIFF	Set <b>ON</b> to use tiff library.	
LIBTIFF_INSTALL_PATH	The root path of libtiff. Needed if <b>USE_TIFF</b> is <b>ON</b> .	
USE_HDF5	Set <b>ON</b> to use hdf5 library.	
HDF5_INSTALL_PATH	The root path of HDF5. Needed if <b>USE_HDF5</b> is <b>ON</b> .	

Table 2: Description of CMake variables for BOT.

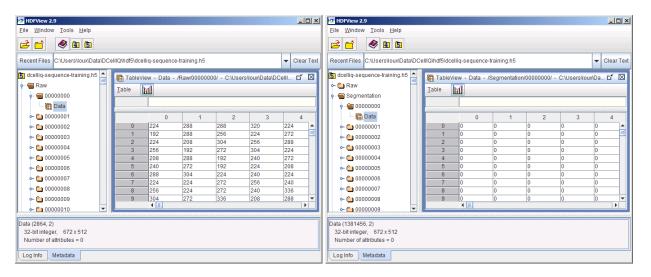


Figure 3: Save raw images and segmentation in HDF5.

#### 3.2 Data Preparation and Event Configuration

#### 3.2.1 Data and Results in HDF5 Format

333

334

335

336

337

338

339

341

343

344

345

The two applications (./app/TrackingTrainer and ./app/TrackingTrainer) accept hdf5 file as input. The raw images and the segmentations are stored in a format shown in Fig. 3. For example, directory 00000000 indicates time point zero.

The tracking result and the training data are stored in the same format as in Fig. 4. The directory name (e.g. Division) must be identical to the one in the event definition ini file. Each event has a pair of subdirectories, namely Source and Target, which corresponds to associated cells from the first frame and the second frame, respectively. Note that for division one cell in the first frame is associated with two cells in the second frame (Fig. 4, left). And, for disappearance, there is no associated cell in the second frame so the id is -1 (Fig. 4, right).

The easiest way to understand this file structure is to explore the sample ./data/dcelliq-sequence-training.h5 using hdfview (freely available at http://www.hdfgroup.org/hdf-java-html/hdfview/).

#### 3.2.2 Data and Results in TIFF and Txt Format (Recommended)

Using hdf5 file has the advantage that everything can be stored in a single file. But it accompanies two disadvantages. Firstly, hdf5 file is not directly viewable, particularly for image data. Secondly, it introduces

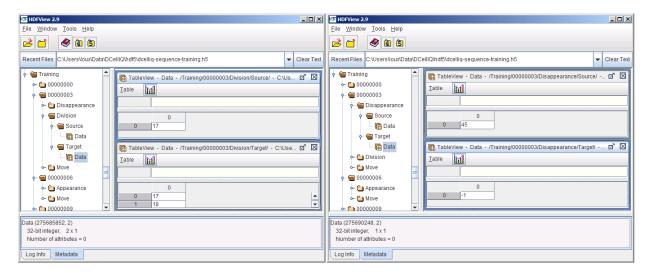


Figure 4: Save training data and tracking results in HDF5.

additional dependency on the huge hdf5 library which can be difficult to compile and install.

We therefore added another approach for data importing: .tiff for image and segmentation, and .txt for annotations. Make sure

- Images and segmentations are stored in tiff format in separate folders.
- Annotations are stored as the following example:

```
[Move]
352
353
          57 -> 57
           58 -> 58
355
          59 -> 60
356
          60 -> 59
357
           61 -> 61
358
          62 -> 62
359
           63 -> 63
360
           64 -> 64
361
362
           . . .
363
           [Division]
364
          17 -> 17 18
365
366
          [Disappearance]
367
          45 -> -1
368
```

348

350

351

369

370

371

Obviously, event name is specified in the squared brackets, and associations of cell id (or ids) from the left and right frame are indicated by ->. For example, "17->1718" means cell #17 in the left frame divides into cell #17 and #18 in the right frame.

#### 3.2.3 Event Configuration in INI Format

```
You can using the default event configuration file ./data/event-configuration-cell.ini, or con-
   struct you own. The format is very simple:
374
   [Global]
375
   k_neighbors = [how many nearest neighbors to be included]
   distance_threshold = [maximum allowed distance as hypotheses thresholding]
377
378
   [Event name, e.g. Move]
379
   Pairing = [one to one association (put 1 1) or one to two association (put 1 2) or ...]
   Feature1 = [the basic object feature] [joint feature, e.g. the measure] [feature weight]
   Feature2 = [the basic object feature] [joint feature, e.g. the measure] [feature weight]
383
384
   [Event name, e.g. Division]
385
```

#### 3.3 A Complete Cell Tracking Example

The following code gives a concrete example of learning the parameter of high-dimensional features. It is extracted from the ./apps/TrackingTrainer.cxx.

```
#include "InputOutput.hxx"
391
   #include "HypothesisSpace.hxx"
392
   #include "ObjectFeatureExtractor.hxx"
393
   #include "AverageObject.hxx"
   #include "SingletsGenerator.hxx"
   #include "MultipletsGenerator.hxx"
396
   #include "CPLEXSolverSystem.hxx"
397
   #include "TrackingPredictor.hxx"
   #include "SolutionCoder.hxx"
   #include "TrainingData.hxx"
   #include "TrackingTrainer.hxx"
401
   #if defined(USE_TIFF)
402
        #include "TIFFReaderWriter.hxx"
403
   #else
404
        #include "HDF5ReaderWriter.hxx"
   #endif
406
407
   using namespace bot;
408
409
   int main(int argc, char* argv[])
410
   #if defined(USE_TIFF)
412
        std::string root(argv[1]);
413
        std::vector<Matrix2D > images, segmentations;
414
        TrainingData training;
415
```

```
std::vector<std::string > references;
416
417
        // load raw images and segmentations
418
        TIFFReaderWriter::loadTiffDir(root + "/raw", images, references);
419
        TIFFReaderWriter::loadTiffDir(root + "/seg", segmentations);
420
        if (images.size() != segmentations.size()) {
421
            std::cerr << "Error:_number_of_raw_images_does_not_match_number_of_segmentations"
422
            << std :: endl :
423
            return EXIT_FAILURE;
        }
425
        // load trainig data from txt files, and each file is
427
        // matched against the references (raw image files)
428
        TIFFReaderWriter::loadAnnotationDir(root + "/training", references, training);
   #else
        std::string filename(argv[1]);
431
        // load the image sequence
432
        std :: vector<Matrix2D > images , segmentations;
433
        TrainingData training;
434
        HDF5ReaderWriter::load(filename, images, segmentations);
                                                                      // load images and segmentations
       HDF5ReaderWriter::load(filename, training);
                                                                        // load trainig data
   #endif
437
438
        // get the context
439
        Context context(images);
440
        std::cout << "****Computing_the_Context****" << std::endl << context
           << std::endl << std::endl;
442
443
        // load the configuration
444
        HypothesisSpace space(argv[2]);
445
        EventConfiguration conf = space.configuration();
447
        // create singlets/muliplets and extract object features
448
        std::cout << "****Extracting_singlets_and_multiplets****" << std::endl;
449
        SingletsSequence singlets_vec;
450
        SingletsSequence avg_singlet_vec;
        MultipletsSequence multiplets_vec;
452
        SingletsGenerator singletsGenerator;
453
        MultipletsGenerator multipletsGenerator(conf.k(), conf.d_max());
454
        ObjectFeatureExtractor extractor(conf.get_feature_names(), context);
455
        for (int32 indT = 0; indT < images.size(); indT ++) {</pre>
            // generate singlets and multiplets
457
            Singlets singlets = singletsGenerator(images[indT], segmentations[indT]);
458
            Multiplets multiplets = multipletsGenerator(images[indT], segmentations[indT], singlets);
459
460
            // extract features for them
461
            extractor(singlets);
```

```
extractor (multiplets);
463
            // save
464
            singlets_vec.push_back(singlets);
465
            avg_singlet_vec.push_back(AverageObject::average(singlets));
            multiplets_vec.push_back(multiplets);
467
468
            std::cout << "#T=="" << indT
469
                << ": _#singlets _= _" << singlets . size ()
470
                << ": "#multiplets == " << multiplets . size() << std :: endl;</pre>
        }
473
474
        // generate hypotheses and extract joint features
475
        space(singlets_vec, avg_singlet_vec, multiplets_vec);
476
        const std::vector<FramePair >& framepairs = space.framepairs();
477
478
        // parse the training data
479
        std::cout << "****Parsing_the_training_data****" << std::endl;
480
        SolutionCoder coder;
481
        int32 nTr = training.times().size();
        for (int32 ind = 0; ind < nTr; ind ++) {
483
            int32 time = training.times()[ind];
484
            std::cout << "****time == " << time << "****" << std::endl;
485
            const LabelAssociations& association = training.associations()[ind];
487
            const std::vector<Event >& events = framepairs[time].events();
            const Singlets& singlets1 = singlets_vec[time];
489
            const Singlets& singlets2 = singlets_vec[time+1];
490
            const Multiplets& multiplets1 = multiplets_vec[time];
491
            const Multiplets& multiplets2 = multiplets_vec[time+1];
492
            Solution solution;
494
            coder.decode(
495
                association,
496
                events,
497
                singlets1, singlets2,
                multiplets1, multiplets2,
499
                solution);
500
            training.solutions().push_back(solution);
501
        }
502
        // start the training
        TrackingTrainer trainer;
505
        const std::vector<Matrix2D > null_vector;
506
        std::vector<Matrix2D > weights = conf.weights(0.5);
507
        std::string msg = trainer(training, framepairs, weights, true);
508
        std::cout << "Training_returns:_" << msg << std::endl;
```

```
conf.weights() = weights;
510
511
       // printe intermediate results: weights, epsilons, losses
512
       513
       std::cout << "Weights: " << std::endl << trainer.weights() << std::endl << std::endl;
514
       std::cout << "Epsilons:_" << std::endl << trainer.epsilons() << std::endl << std::endl;
515
       std::cout << "Losses:_" << std::endl << trainer.losses() << std::endl << std::endl;
516
517
       // print the final weights
518
       std::cout << "Learned_weights:_" << std::endl;</pre>
       conf.print();
520
521
522
       return EXIT_SUCCESS;
523
   }
524
```

## 25 Acknowledgement

We especially thank Bjoern Andres (University of Heidelberg), Christoph Straehle (University of Heidelberg) and Jan Funke (ETHZ) for their constructive comments on our manuscript and the code. We also thank Julian J. McAuley (The Australian National University) and Choon Hui Teo (Yahoo! Labs) for their suggestion on the implementation of the bundle method.

We are very grateful for the following software/code packages that BOT depends on:

- 1. The IBM Academic Initiative allows us to access IBM ILOG CPLEX free of charge.
- 2. The SimpleIni library<sup>8</sup> allows us to efficiently read and write INI-style configuration files.
- 3. The FastEMD library<sup>9</sup> provides fast earth mover's distance computation.

### References

527

528

531

532

- [1] X. Lou and F. A. Hamprecht. Structured Learning for Cell Tracking. In *Neural Information Processing Systems (NIPS)*, 2011
- [2] X. Lou, F. O. Kaster, M. S. Lindner, B. X. Kausler, U. Koethe, H. Jaenicke, B. Hoeckendorf, J. Wittbrodt, and F. A. Hamprecht. DELTR: Digital Embryo Lineage Tree Reconstructor. In *IEEE International Symposium on Biomedical Imaging: From Nano to Macro (ISBI)*, 2011. (Oral).
- [3] T. Kanade, Z. Yin, R. Bise, S. Huh, S. E. Eom, M. Sandbothe, and M. Chen. Cell Image Analysis: Algorithms, System
   and Applications. In *IEEE Workshop on Applications of Computer Vision (WACV)*, 2011.
- [4] D. Padfield, J. Rittscher, and B. Roysam. Coupled Minimum-Cost Flow Cell Tracking. In *Information Processing in Medical Imaging*, 2009.
- [5] C. Sommer, C. Strähle, U. Köthe, and F. A. Hamprecht. ilastik: Interactive Learning and Segmentation Toolkit. In *IEEE International Symposium on Biomedical Imaging: From Nano to Macro (ISBI)*, 2011.

<sup>&</sup>lt;sup>7</sup>https://www.ibm.com/developerworks/university/academicinitiative/

<sup>8</sup>http://code.jellycan.com/simpleini/

<sup>9</sup>http://www.cs.huji.ac.il/ofirpele/FastEMD/code/