



Vidjil, a platform for the interactive analysis of immune repertoires

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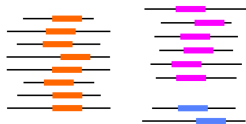
Vidjil Platform

High-throughput Repertoire Sequencing (RepSeq) analysis

Web Application

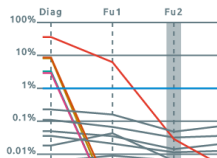
Patient database
Server

Algorithm

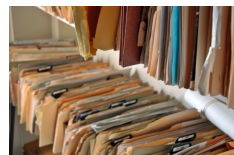


C++

Client



Javascript, d3.js



Python, web2py,
AJAX

- ▶ all the Vidjil components are open-source (GPL v3)
- ▶ code on <http://www.vidjil.org/> and on GitHub
- ▶ continuous integration, > 2,000 unit and functional tests

Duez et al., PLOS One, 2016

Jane Austen patients import/export cluster color by filter

patient from Winchester hospital, LAL-B

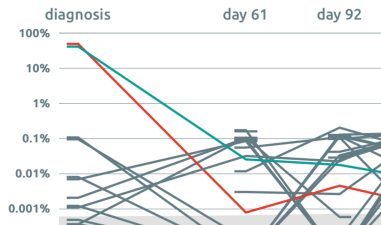
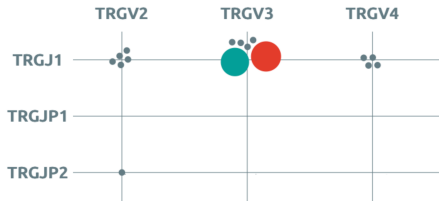
H IGH

G TRG

<i>i</i>	L45601-R1.fastq	analyzed reads	1 798 170 (75.23%)
	1789-12-25	selected locus	1 797 117 (75.18%)



Diagnosis sample, TRG/IGH BIOMED-2 primers

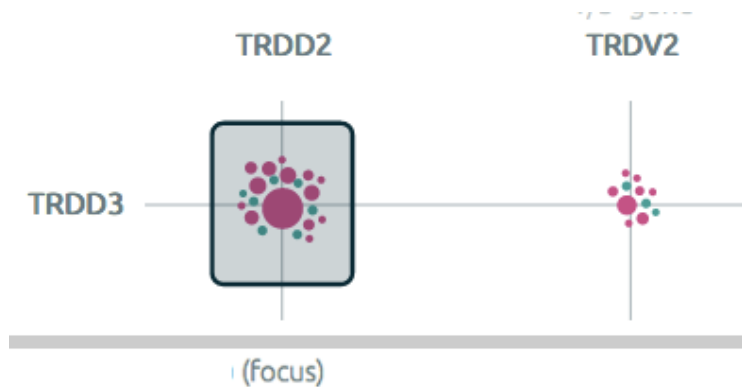


merge align > to IMGT/V-QUEST > to IgBlast > to Blast

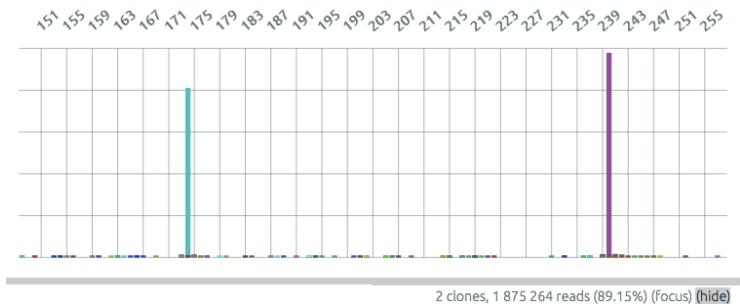
4 clones, 738 494 reads (90.53%)

× TRGV3 1/11/2 J1	49.30%	★ i	GCCACCTGGGACAGCTCCC-TT-GTTC--ATTATAAGAAACTCTTTGGCAGTG
× TRGV3 4/1/2 J1	41.23%	★ i	GCCACCTGGG--A--T--A--T--T--ATTATAAGAAACTCTTTGGCAGTG
× TRGV3 3/16/3 J1	0.0021%	★ i	GCCG-CTTGGG-ACCCCAATTTGGTACGGGTTATAAGAAACTCTTTGGCAGTG
× TRGV3 5/4/2 J1	+	★ i	GCCACCTGGG---GC--CA-A-T--T--A-TA--AGAAACTCTTTGGCAGTG

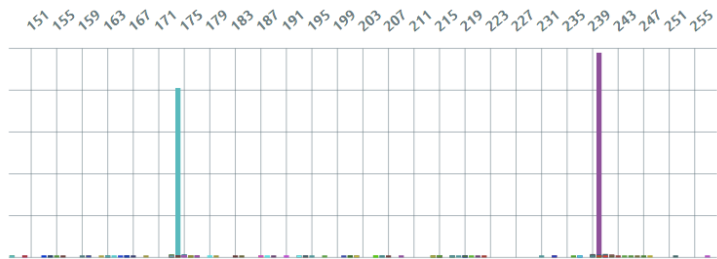
Web application – Browsing and filtering clones



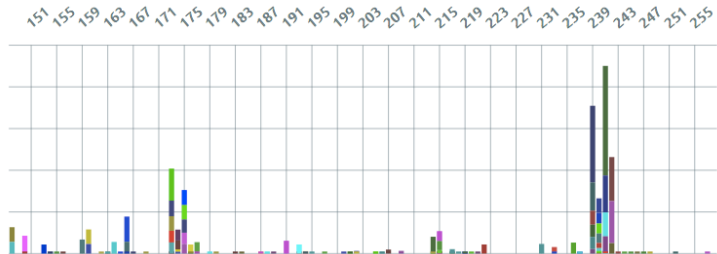
Web application – Browsing and filtering clones



Web application – Browsing and filtering clones



2 clones, 1 875 264 reads (89.15%) (focus) [\(hide\)](#)



Web application: traceability– Reports for medical records

Sample info (IonXpress_086)

Filename: IonXpress_086.L1413629.fastq
 Sample date: 2014-12-19
 Software used: vidjil dev 0cf35de (2015-02-17)
 Parameters: -i -g germline -c clones -z 100 -r 1
 Analysis date: 2015-02-17 21:50:09

User note

PCR avec 250ng d'ADN, Ion Pgt+ 50µL

Reads distribution

total
 segmented

- TRG
- IGK+
- VdJa
- IGH
- TRD+

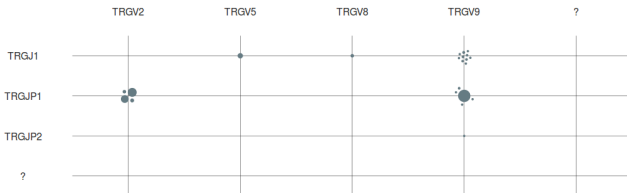
IonXpress 086

632 686

88%



IonXpress_086 – TRG

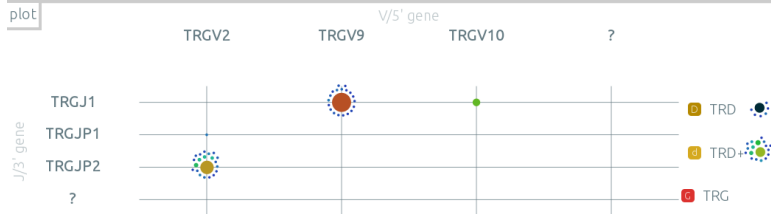
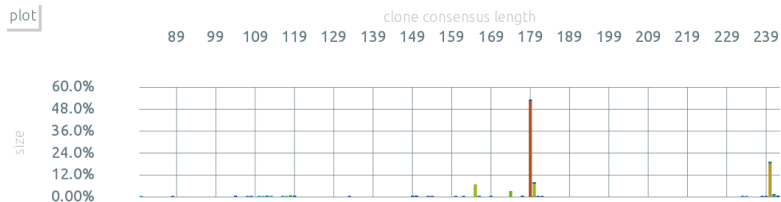


>TRGV9*01 -8/10/-3 TRGJP1*01 | 13.44% (45.99% of TRG)
 CGGCACCTGTCAGAAAGGAATCCGGCATTCCGTCAGGCAAAATTGAGGTGGATAGGATACCTGAAACGTCTACATCCACTCTCACCATTCACAATGTAGAGAAACAGGACATAGCT
 ACCTACTACTGTGCTCTGT CTCAACGAGGG CCACTGGTGGTTCAAGATATTTGGCTGAAGGGACTAAGCTCATAGTAACTTCGCTGGTAA

>TRGV2*01 -9/6/-1 TRGJP1*01 | 5.062% (17.32% of TRG)
 GGAAGGCCCCACAGCGTCTTCAGTACTATGACTCCTCAACTCCAAGGTTGTGTGGAAATCAGGAGTCAGTCAGGGAAGTATTATACTTACGCAAGCACAGGAACAACCTGGAG
 ATTGATACTCGGAAATCTAAATTGAAAATGACTTTGGGGTCTATTACTGTGCCACC CCTGG TACCCTGGTGGTTCAAGATATTTGGCTGAAGGGACTAAGCTCATAGTAA
 CTTCCGCTGGTAA

2017 News – Single samples displayed with two plots

Both Genescan-like view and V/J genes grid, or any other plot



2017 News – Germline Sequences (experimental)

Compare clone sequences to the original germlines

IGHV4-34 1/AC/4 D6-19 6/7/1 J3 GGCTGTATATTACTGTGCGAAAG|AC|ATAGCAGTGGC|ACTCCCC|GATG
IGHV4-34*01 GGCTGTGTATTACTGTGCGAGAG - - - - -

2017 News – More clustering options, including undo (was previously “merge”)

cluster	align	> to IMGT/V-QUEST	> to IgBlast	> to Blast
×	IGHV1-24 0/CGACCC/3 D6-13 1/TAAC/1	1.409%	★	i
×	IGHV1-24 0//3 D6-6 0/GAGG/3 J6*03	1.208%	★	i
×	IGHV1-24 0//3 D3-10 6/11/7 J6*02	1.170%	★	i
×	IGHV1-24 0/GCCCCG/0 D3-3 6/CGGA/6	1.069%	★	i

import/export cluster color by tag filter

The screenshot shows a software interface with a dropdown menu open over the 'cluster' button. The menu contains the following options:

- revert to previous clusters
- cluster selected clones
- cluster by V/5'
- cluster by J/3'
- cluster by locus
- cluster by similarity
- break selected clusters
- break all clusters

In the background, there is a table with columns for 'IGH' and 'IGH', and a row with 'analyze' and 'select' buttons. Below the table are several colored squares (orange, yellow, blue, purple, green, pink) and a vertical bar with a histogram-like scale.

2017 News – Improved permanent URLs

Saving elements of the client state: set, plot, clones...

Discuss data with colleagues or with us

```
http://app.vidjil.org/?set=3241&config=39
```

```
http://app.vidjil.org/?set=3241&config=
39&plot=v,size,bar&clone=11,31
```

More elements to come

Exchanging data between RepSeq software

Encoding a set of clones with V(D)J recombinations, possibly with user annotations

```
{
  "id": "clone-001",
  "sequence": "CTCATACACCCAGGAGGTGGAGCTGGATATTGATACTACGAAATCTAATTG(...)",
  "reads" : [ 243241 ],
  "germline": "TRG",
  "top": 1,
  "seg":
  {
    "5": {"name": "TRGV5*01", "start": 1, "stop": 86},
    "3": {"name": "TRGJ1*02", "start": 89, "stop": 118},
    "cdr3": { "start": 77, "stop": 104, "seq": "gccacctgggccttatt(...)" }
  }
}
```

This .json format is described in doc/format-analysis.org. We are open to make evolve this format to better match other software needs.

Upcoming: encoding distributions.

Exchanging data between RepSeq software

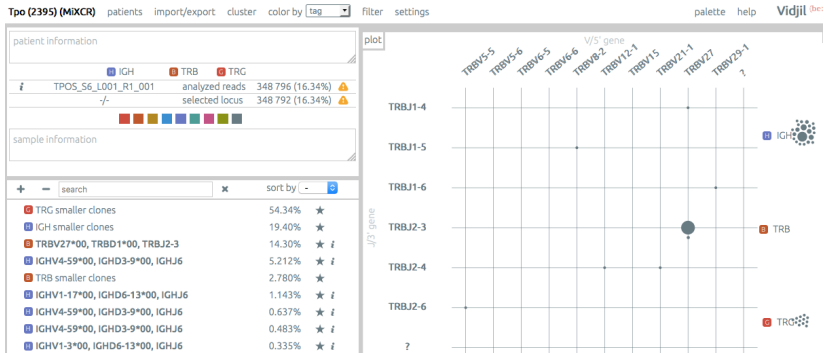
Encoding a set of clones with V(D)J recombinations, possibly with user annotations

What is a clone?

There are several definitions of what may be a clonotype, depending on different RepSeq software or studies. The .json format (and the Vidjil web application) takes a string identifier that may correspond to any kind of clone definition.

- ▶ CGAGAGGTTACTAT...GGTAGGGCAGTACTAC (Vidjil, 50 nt window centered on the CDR3)
- ▶ CARPRDWNTYYYYGMDVW (CDR3 AA sequence)
- ▶ CARPRDWNTYYYYGMDVW IGHV3-11*00 IGHJ6*00 (CDR3 AA sequence with additional V/J gene information, MiXCR)
- ▶ clone sequence as computed by the ARReST
- ▶ IMGT clonotype (AA) or (nt)

MiXCR output on Vidjil



2018: We plan to display results from other software.

app.vidjil.org/analyze – analyzing few sequences

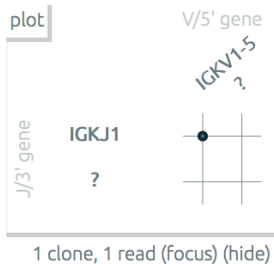
```
>seq-1043-igk
```

```
CTCCTGCTACTCTGGCTCCCAGGTGCCAAATGTGACATCCAGA  
TGACCCAGTCTCCTTCCACCCTGTCTGCGTCTGTAGGAGACAG  
AGTCACCATCACCTGCCGGGCCAGTCAGAGTATTAATAACAAC  
TTGGCCTGGTATCAGGAGAAGCCAGGGAAAGCCCCTAAGGTC
```

Submit

Export results to FASTA

✘ IGKV1-5 9/ACTT/1 J1



No login, no connection to patient/sample database

Next: Add sequences/clones to one sample

2 103 505 (95.33%)
2 103 503 (95.33%)
100%

TRGJ1

sort by
95.87
0.473
0.331
0.190
0.181
0.159
0.138
0.122
0.109% ★ ⓘ
0.103% ★ ⓘ
0.076% ★

Add clones from sequences

>Sanger 2017-03-01
GGAAGGCCCCACAGCGTCTTCTGTA CTACTATGACTCCTACA ACTCCAGGGTTGTGTTGGAATCAGGAATCAGTCG
AGAAAAGTATCATACTTATGCAAGCACAGGGAAGAGCCTTAAATTTATACTGGAAAATCTAATTGAACGTGAC
TCTGGGGTCTATTACTGTGCCACCTGGAAGATACC ACTGGTTGTTCAAAGATATTTGCTGAAGGGACTAAGCT
CATAGTA ACTTCGCCTGGTAA

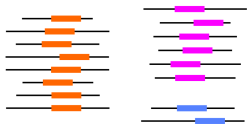
Add

Vidjil Platform

High-throughput Repertoire Sequencing (RepSeq) analysis

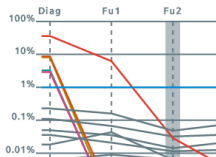
Web Application

Algorithm



C++

Client



Javascript, d3.js

Patient database
Server



Python, web2py,
AJAX



Patient database and server

Autonomous RepSeq analysis pipeline in a clinical/research lab



Patient database and server

Autonomous RepSeq analysis pipeline in a clinical/research lab



Upload

upload list

L1413893.fasta



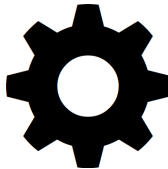


Patient database and server

Autonomous RepSeq analysis pipeline in a clinical/research lab



Upload



Process

upload list

L1413893.fasta



last processing status

2015-02-09

RUNNING

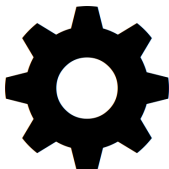


Patient database and server

Autonomous RepSeq analysis pipeline in a clinical/research lab



Upload



Process



Analyze

upload list
L1413893.fasta

last processing	status
2015-02-09	RUNNING

see the result:

[multi](#)



Patient database and server – Hosting

app.vidjil.org

Test server: research use only

Secure data center in Lille

Supported by U. Bristol, SIRIC OncoLille and U. Lille



Patient database and server – Hosting

app.vidjil.org

Test server: research use only

Secure data center in Lille

Supported by U. Bristol, SIRIC OncoLille and U. Lille

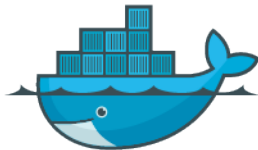
In your lab/hospital

A standard server is enough

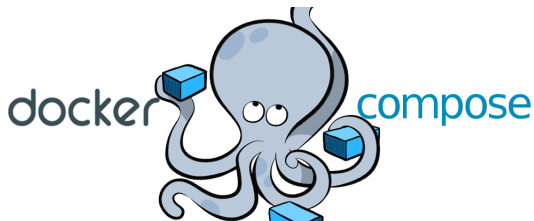
Supported by local IT staff

2017 News – Improvements to **Docker** images

Stability and usability



docker



2017 News – Classify and filter your data with tags

search

name	birth	info
Austeen Jane (43)	1775-12-16	#T-ALL #diagnosis #cr Winchester hospital
de Cervantes Miguel (42)	1547-09-29	#relapse #pre-SCT #post-SCT
von Schiller Friedrich (41)	1759-01-01	#B-ALL #FR1 blood sample, 10µM primer
Alighieri Dante (40)	1265-01-01	#B-ALL #diagnosis #relapse #KDE
Sand Geroge (39)	1804-02-01	#WM
Goethe Johann Wolfgang (38)	1749-08-28	#T-ALL
Bilderdijk Willem (37)	1756-09-07	
خلدون ابن (36)	1332-05-27	#T-ALL #diagnosis 60% blasts
Němcová Božena (35)	1820-02-04	#T-ALL
Aarestrup Emil (34)	1800-12-04	
Držić Marin (33)	1508-01-01	#CLL 98.4%
Brenner Sophia Elisabet (32)	1659-04-29	#diagnosis #B-ALL #BCR-ABL
Alecsandri Vasile (31)	1801-07-21	#B-ALL #IKAROS
Σολωμός Διονύσιος (30)	1798-04-08	#T-ALL #diagnosis #pre-SCT
Collett Camilla (29)	1813-01-23	
Wilkońska Paulina (28)	1815-01-01	#CLL
Espanca Florbela (27)	1894-12-08	#T-ALL 70% blasts
老子 Laozi (26)		

2017 News – Classify and filter your data with tags

search	#ALL		
name	ALL	birth	info
Austee		1775-12-16	#T-ALL #diagnosis #cr Winchester hospital
de Cen	B-ALL	1547-09-29	#relapse #pre-SCT #post-SCT
von Sch	T-ALL	1759-01-01	#B-ALL #FR1 blood sample, 10µM primer
Alighieri Dante (40)		1265-01-01	#B-ALL #diagnosis #relapse #KDE
Sand Geroge (39)		1804-02-01	#WM
Goethe Johann Wolfgang (38)		1749-08-28	#T-ALL
Bilderdijk Willem (37)		1756-09-07	
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Collett Camilla (29)		1813-01-23	
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老子 Laozi (26)			



2017 News – Classify and filter your data with **tags**

search

name	birth	info
von Schiller Friedrich (41)	1759-01-01	#B-ALL #FR1 blood sample, 10 μ M primer
Alighieri Dante (40)	1265-01-01	#B-ALL #diagnosis #relapse #KDE
Brenner Sophia Elisabet (32)	1659-04-29	#diagnosis #B-ALL #BCR-ABL
Alecsandri Vasile (31)	1801-07-21	#B-ALL #IKAROS

2017 News – Download auxiliary **output files**

May include bigger lists of clones, statistic analysis...

last processing	status		
2017-10-05 15:43:39	COMPLETED	dl out	 run ...
2017-10-05 16:07:02	COMPLETED	dl out	 run ...

file	size
000078.edges	0
000078.vidjil.log	58.8 kB
000078.vidjil	178 kB
000078.windows.fa	6.33 kB
000078.vdj.fa	36.9 kB
seq	4.10 kB
000078-34.fuse.log	1.09 kB
000078-26.fuse.log	1.72 kB

Working collaboratively on shared data

Lille hospital



Working collaboratively on shared data

Lille hospital



Upload data
Create patient

Working collaboratively on shared data

Lille hospital



Upload data
Create patient

Annotate results
Freeze/sign (upcoming)

Working collaboratively on shared data

Lille hospital



Upload data
Create patient

Annotate results
Freeze/sign (upcoming)

Read-only access

Working collaboratively on shared data

Lille hospital



Upload data
Create patient

Annotate results
Freeze/sign (upcoming)

Read-only access

They can also belong to other organizations

Track updates to patient records

Lille hospital



Track updates to patient records

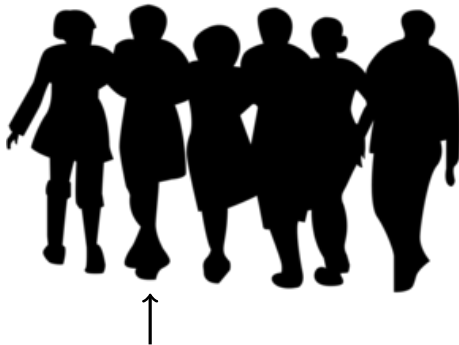
Lille hospital



2016-09-29 10:15 – **Léa** created patient Jane Doe

Track updates to patient records

Lille hospital

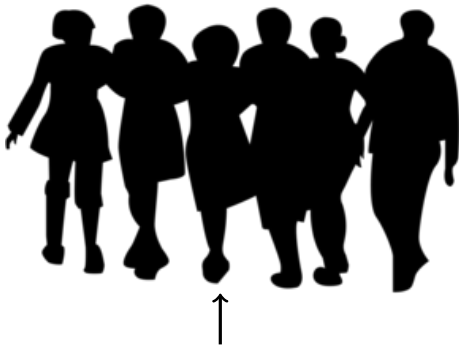


2016-09-29 10:15 – **Léa** created patient Jane Doe

2016-09-29 11:45 – **Guy** uploaded seqs.fastq (15 MB)

Track updates to patient records

Lille hospital



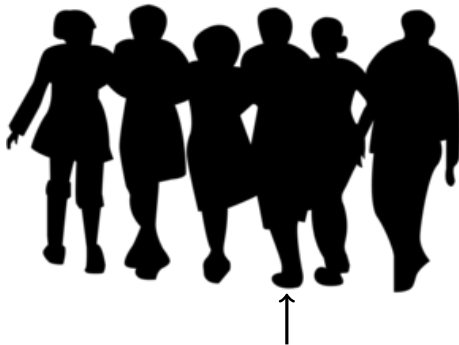
2016-09-29 10:15 – **Léa** created patient Jane Doe

2016-09-29 11:45 – **Guy** uploaded seqs.fastq (15 MB)

2016-09-29 12:12 – **Élodie** annotated the results

Track updates to patient records

Lille hospital



2016-09-29 10:15 – **Léa** created patient Jane Doe

2016-09-29 11:45 – **Guy** uploaded seqs.fastq (15 MB)

2016-09-29 12:12 – **Élodie** annotated the results

2016-09-30 10:47 – **Cindy** froze and signed the report