Uncovering the topology of the temporal region in Alzheimer's disease.

Philip Hartout

November 24, 2020

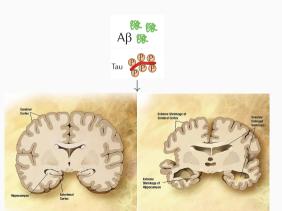
Alzheimer's disease:



- Nearly 40 million people live with AD
- Cost in US alone \$ 2 trillion by 2030
- Among leading causes of death in EU

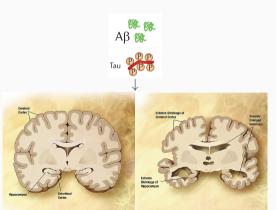
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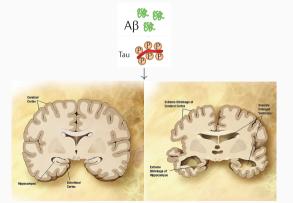
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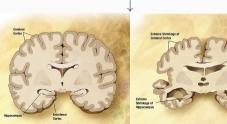
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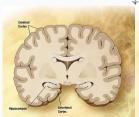


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- Concerned with "properties of a geometric object that are preserved under continuous deformations, such as [...] crumpling."
- Recently, persistent homology has emerged to quantify (differences in) the shape of data.
- How can we apply persistent homology to quantify changes in shape due to Alzheimer's disease?

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1. Diagnosis (classification)

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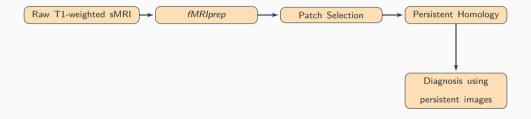
- 1. Diagnosis (classification)
- 2. Subtype identification
- 3. Progression & forecasting
- \rightarrow Some findings in these directions will be presented today

Raw T1-weighted sMRI









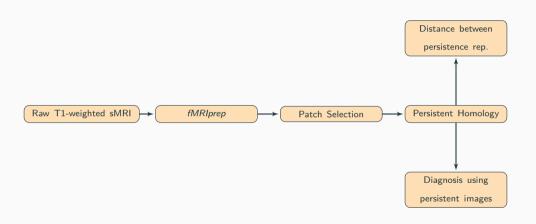




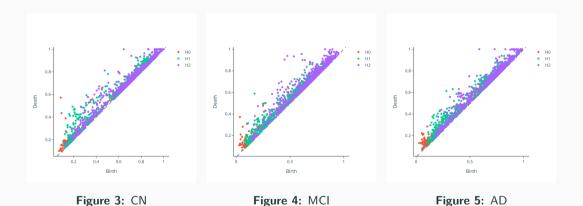
Figure 2: AUCPRC on each patch, achieved using a model described in earlier work. Chosen patch for analyses is boxed in red (patch with highest accuracy).

Computed the cubical filtration to obtain persistence image for each patch

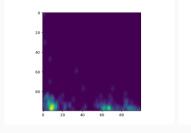
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- Simple CNN to classify AD/CN patients.

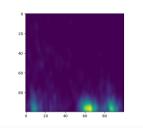
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- Simple CNN to classify AD/CN patients.
- Use same partition as NeurIPS submission, train ANN 3 times on each partition

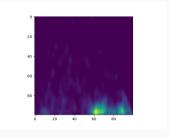
I - Persistent homology



I - Diagnosis prediction - Persistence Images







I - Diagnosis prediction - Persistence Images

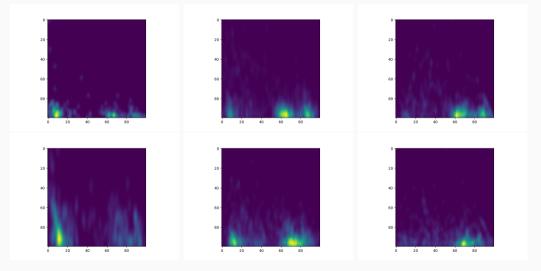
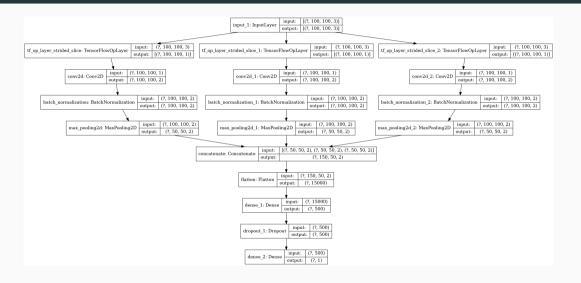


Figure 6: Columns: hom. dimension (0,1,2); Row: Diagnostic category (CN top, AD bottom).

I - Diagnosis prediction - Network architecture



I - Diagnosis prediction - Performance

Model trained on PIs from patches
0.81 ± 0.01
0.78 ± 0.03
0.81 ± 0.04
0.77 ± 0.03
0.85 ± 0.03

Table 1: Performance metrics of the model.

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Table 1: Performance metrics of the model.

 \rightarrow using a normal CPU, training takes 1 minute!

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- Compute a persistence landscape (allows for statistical analysis)
- Compute a median persistence landscape with one layer for each category (CN, MCI, AD)
- Compute L¹ norm from median landscape for each image within category

Median persistence landscape.

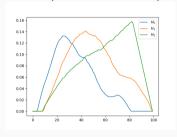


Figure 7: CN

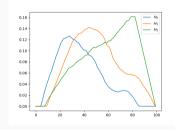


Figure 8: MCI

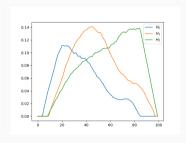


Figure 9: AD

Question: How topologically heterogenous is the data?

	Mean	Median	Standard deviation	Q3	Max	Skewness
CN H ₀	2.16	2.00	0.78	2.50	7.41	1.78
$CN H_1$	2.61	2.27	1.17	2.93	9.47	1.92
$CN H_2$	2.38	2.23	0.88	2.79	7.19	1.39
$MCIH_0$	2.24	2.04	0.82	2.55	6.21	1.71
$MCIH_1$	2.57	2.19	1.29	2.80	11.87	2.57
$MCIH_2$	2.40	2.27	0.83	2.82	6.55	1.18
AD H_0	2.40	2.18	0.96	2.77	7.77	1.97
AD H_1	2.47	2.13	1.15	2.77	9.28	2.10
AD H ₂	2.36	2.20	0.80	2.75	8.39	1.64

Table 2: Summary statistics of the distribution of distances

III - Distance analysis among patients who deteriorate vs. those who don't

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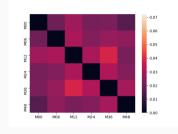
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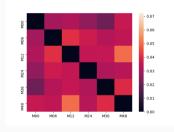
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- The data is a longitudinal dataset (multiple timepoints are available for each patient)
- Some patients deteriorate (transition from CN→MCI or from MCI→AD)
- Compute pairwise distance between patients (L¹ PL distance, Wasserstein distance and bottleneck distance), and average for each patient.

Example of L^1 norm between PLs of a deteriorating patient.





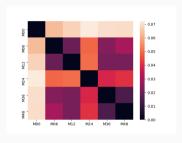
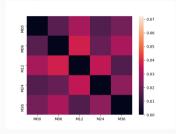


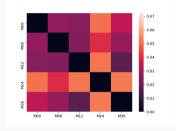
Figure 10: H_0

Figure 11: H_1

Figure 12: H_2

Example of L^1 norm between PLs of a subject who does *not* deteriorate.





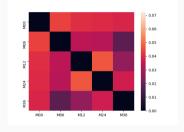
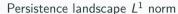


Figure 13: H_0

Figure 14: H_1

Figure 15: H_2



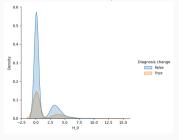


Figure 16: H_0

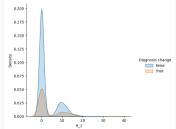


Figure 17: H_1

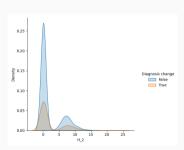


Figure 18: H_2

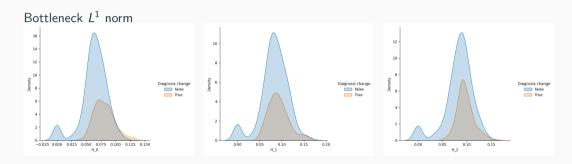
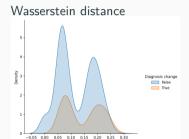
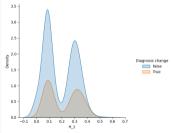


Figure 20: H_1

Figure 19: H_0

Figure 21: H_2





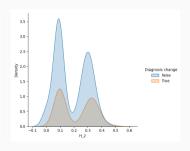


Figure 22: H_0

Figure 23: H_1

Figure 24: H_2

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2. Distances

- In general, very coarse analysis (averages do not pick out subtypes or use individual features to differentiate between images), sensitive to noise.
- Could learn better (topology-based) embeddings to better distinguish between people who
 progress versus those who do not and more finegrained subtype identification resilient noise.

Summary

GitHub repository of the project

github.com/pjhartout/TDA_ADNI_MLCB

With thanks to Bastian Rieck for the supervision and Sarah Brueningk, Felix Hensel, Catherine Jutzeler, Merel Kuijs and Louis Lukas for insightful discussions, code, and data.

