

Dept. of Computer Science, University of Copenhagen

Anatomical tree-shape analysis

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In collaboration with!



Starting point: What does the average human airway tree look like?



Wanted: Parametric statistical model for trees, allowing variations in branch count, tree-topological structure and branch geometry



Airway shape modeling



- Smoker's lung (COPD) is caused by inhaling damaging particles.
- Likely that damage made depends on airway geometry
- Reversely: COPD changes the airway geometry, e.g. airway wall thickness.
- ► ~→ Geometry can help diagnosis/prediction.

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Airway shape modeling

Properties of airway trees:

- Topology, branch shape, branch radius
- Somewhat variable topology (combinatorics) in *anatomical* tree
- Substantial amount of noise in *segmented* trees (missing or spurious branches), especially in COPD patients, *i.e. inherently incomplete data*



Airway shape modeling

Wanted properties:

 $T = T_1$

 T_2

Figure: Tolerance of structural noise.

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 T_3

Airway shape modeling

Wanted properties:

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Figure: Handling of internal structural differences.

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Airway shape modeling

We shall consider airway centerline trees embedded in \mathbb{R}^3 .



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Not just airways....

Data with an underlying tree- or graph-structure appear in all kinds of applications:

- Anatomical structures such as airways, blood vessels and other vascularization systems;
- Skeletal structures such as medial axes;
- Descriptors of hierarchical structures (genetics, scale space)



Figure: Figures from Lo; Wang et al.; Sebastian et al.; Kuijper

A space of tree-like shapes: Intuition

What would a path-connected space of deformable trees look like?



- Easy: Trees with same topology in their own "component"
- Harder: How are the components connected?
- Solution: glue collapsed trees, deforming one topology to another

► ~→ Stratified space, self intersections

A space of tree-like shapes: Intuition

The tree-space has conical "bubbles"



- TED is a classical, algorithmic distance
- tree-space with TED is a nonlinear metric space
- ▶ dist(T₁, T₂) is the minimal total cost of changing T₁ into T₂ through three basic operations:

(a)

Remove edge, add edge, deform edge.

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 Tree-space with TED is a geodesic space, but almost all geodesics between pairs of trees are non-unique (infinitely many).



- Then what is the average of two trees? Many!
- Tree-space with TED has everywhere unbounded curvature.

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• TED is *not* suitable for statistics.

Many state-of-the-art approaches to distance measures and statistics on tree- and graph-structured data *are* based on TED!

- Ferrer, Valveny, Serratosa, Riesen, Bunke: Generalized median graph computation by means of graph embedding in vector spaces. Pattern Recognition 43 (4), 2010.
- Riesen and Bunke: Approximate Graph Edit Distance by means of Bipartite Graph Matching. Image and Vision Computing 27 (7), 2009.
- Trinh and Kimia, Learning Prototypical Shapes for Object Categories. CVPR workshops 2010.

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The problems can be "solved" by choosing specific geodesics. OBS! Geometric methods can no longer be used for proofs, and one risks choosing problematic paths.



Figure: Trinh and Kimia (CVPR workshops 2010) compute average shock graphs using TED with the simplest possible choice of geodesics.

Build a tree-space: Tree representation

How to represent geometric trees mathematically? Tree-like (pre-)shape = pair (\mathcal{T}, x)

- 𝒮 = (V, E, r, <) rooted, ordered/planar binary tree, describing the tree topology (combinatorics)
- ► x ∈ ⊕_{e∈E} A, each coordinate in an attribute space A describing edge shape

 $= \sqrt[3]{4} \sqrt[4]{6} + (1, \mathbf{1}, \mathbf{1}, \mathbf{1}, \mathbf{1}, \mathbf{1})$

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Build a tree-space: Tree representation

We are allowing collapsed edges, which means that

- we can represent higher order vertices
- \blacktriangleright we can represent trees of different sizes using the same combinatorial tree $\mathcal T$



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(dotted line = collapsed edge = zero/constant attribute)

Build a tree-space: Tree representation

- Edge representation through landmark points:
- Edge shape space is $(\mathbb{R}^d)^n$, d = 2, 3.
- (For most results, this can be generalized to other vector spaces)

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The space of tree-like preshapes

First: \mathscr{T} an infinite, ordered (planar), rooted binary tree Definition Define the space of tree-like *pre*-shapes as the direct sum



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where $(\mathbb{R}^d)^n$ is the edge shape space. This is just a space of *pre-shapes*.

From pre-shapes to shapes

Many shapes have more than one representation



From pre-shapes to shapes

Not all shape deformations can be recovered as natural paths in the pre-shape space:



Shape space definition

- Start with the pre-shape space $X = \bigoplus_{e \in E} (\mathbb{R}^d)^n$.
- Define an equivalence ~ by identifying points in X that represent the same tree-shape.



- ▶ This corresponds to identifying, or gluing together, subspaces $\{x \in X | x_e = 0 \text{ if } e \notin E_1\}$ and $\{x \in X | x_e = 0 \text{ if } e \notin E_2\}$ in X.

Shape space definition

Remark

- Tree-shape definition a little unorthodox: we do not factor out scale and rotation of the tree.
- Our data (segmented airway trees) are incomplete; the number of segmented branches is unstable and depends on the health of the patient.



Definition of metric on tree-space

Given a metric d on the vector space $X = \bigoplus_{e \in E} (\mathbb{R}^d)^n$ we define the quotient pseudometric \overline{d} on the quotient space $\overline{X} = X / \sim$ by setting

$$\bar{d}(\bar{x},\bar{y}) = \inf\left\{\sum_{i=1}^k d(x_i,y_i)|x_1\in\bar{x},y_i\sim x_{i+1},y_k\in\bar{y}\right\}.$$
 (1)



 $\bar{d}(\bar{x},\bar{w}) = d_1 + d_2 + d_3$

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Theorem The quotient pseudometric \overline{d} is a metric on \overline{X} .

Definition of metric on tree-space

• Two metrics on \bar{X} from two product norms on $X = \bigoplus_{e \in E} (\mathbb{R}^d)^n$:

11 norm:
$$d_1(x, y) = \sum_{e \in E} ||x_e - y_e||$$

12 norm: $d_2(x, y) = \sqrt{\sum_{e \in E} ||x_e - y_e||^2}$

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- \bar{d}_1 = Tree Edit Distance
- Terminology: d
 ₂ = QED (Quotient Euclidean Distance) metric.

Theorem

Let $\bar{d} = \bar{d}_1$ or \bar{d}_2 . Then (\bar{X}, \bar{d}) is a geodesic space.

Unordered trees

- Give each tree a random order
- ▶ Denote by G the group of reorderings of the edges (in 𝒮) that do not alter the connectivity of the tree.
- The space of spatial/unordered trees is the space $\bar{X} = \bar{X}/G$
- Give \overline{X} the quotient pseudometric \overline{d} .
- $\overline{d}(\overline{x}, \overline{y})$ chooses the order that minimizes $\overline{d}(\overline{x}, \overline{y})$.

Theorem

For the quotient pseudometric \overline{d} induced by either \overline{d}_1 or \overline{d}_2 , the function \overline{d} is a metric and $(\overline{X}, \overline{d})$ is a geodesic space.

Distances between airways?

Evaluation of metric:

MDS based on approximate geodesic distances between 30 airways of healthy individuals and individuals with moderate COPD.



Complexity of computing tree-space geodesics?

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Assume edge attributes have dimension > 1 (for dim = 1, Scott Provan).
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Theorem

Computing QED geodesics is NP complete.



A little metric geometry – geodesics

• Let (X, d) be a metric space. The length of a curve $c: [a, b] \rightarrow X$ is

$$l(c) = sup_{a=t_0 \le t_1 \le \dots \le t_n = b} \sum_{i=0}^{n-1} d(c(t_i, t_{i+1})).$$



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- A geodesic from x to y in X is a path $c: [a, b] \to X$ such that c(a) = x, c(b) = y and l(c) = d(x, y).
- ► (X, d) is a geodesic space if all pairs x, y can be joined by a geodesic.

Curvature in metric spaces



A CAT(0) space is a metric space in which geodesic triangles are "thinner" than for their comparison triangles in the plane; that is, d(x, a) ≤ d(x̄, ā).

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Curvature in metric spaces



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• A space has non-positive curvature if it is locally CAT(0).

Curvature in metric spaces



- A CAT(0) space is a metric space in which geodesic triangles are "thinner" than for their comparison triangles in the plane; that is, d(x, a) ≤ d(x̄, ā).
- ► A space has non-positive curvature if it is locally CAT(0).
- (Similarly define curvature bounded by κ by using comparison triangles in hyperbolic space or spheres of curvature κ.)



Theorem (see e.g. Bridson-Haefliger)

Let (X, d) be a CAT(0) space; then all pairs of points have a unique geodesic joining them. The same holds locally in $CAT(\kappa)$ spaces, $\kappa \neq 0$.

Statistics in metric spaces?

Theorem (Sturm)

Frechet means are unique in CAT(0) spaces.



Other midpoint-finding algorithms also converge in CAT(0) spaces:

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- centroid
- Birkhoff shortening
- circumcenters

Curvature of shape space?

> This space has everywhere unbounded curvature!







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Bad news for statistics?

Regularize via extra assumptions on the trees:

- Restrict to: all representations of certain restricted tree topologies.
- **Example 1:** Restrict to the set \bar{X}_N of trees with N leaves.



• Example 2: Restrict to all topologies occuring in airway trees.

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Curvature of shape space

Theorem

► Consider (X̄, d̄₂) and (X̄, d̄₂), ordered/unordered tree-shape space.

A (1) × A (2) × A (2) ×

- At generic points, the space is locally CAT(0).
- Its geodesics are locally unique at generic points.
- At non-generic points, the curvature is unbounded.

Curvature of shape space

In fact, curvature is one of:



Figure: Space of ordered/unordered trees with at most 2 edges

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How much better did it get?

 Reasonably large CAT(0) neighborhoods are now larger containing several top-dimensional tree topologies.

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Computational complexity? Still NP complete.

We can compute means! Leaf vasculature data:

Figure: A set of vascular trees from ivy leaves form a set of planar tree-shapes.



Figure: a): The vascular trees are extracted from photos of ivy leaves. b) The mean vascular tree.

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We can compute means!

The mean upper airway tree¹



Figure: A set of upper airway tree-shapes along with their mean tree-shape.

We can compute means!





Figure: The QED and TED (algorithm by Trinh and Kimia) means.

¹Feragen et al, *Towards a theory of statistical tree-shape analysis*, submitted one aasa@diku.dk,

Dealing with NP - Useful property of airways

The first 6-8 generations of the airway tree are "similar" in different people.



NB!: Not all present in all people; not all present in all segmentations.

Regularize via second set of assumptions

- Label the "leaves" of your trees and insist that all trees have the same leaf label set.
- Polynomial time distance algorithms (Owen, Provan)
- Also: Factor out leaf labels via leaf permutation group ~> NP complete.

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Statistics on larger trees: Mean airway



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Joint with Megan Owen.

Idea:

 Generate leaf label configurations and the corresponding tree spanning the labels



 Evaluate configuration in comparison with training data using geodesic deformations between leaf-labeled airway trees (Owen, Provan)

²Feragen, Petersen, Owen, Lo, Thomsen, Dirksen, Wille, de Bruijne, *A hierarchical scheme for geodesic anatomical labeling of airway trees*, MICCAI 2012. aasa@diku.dk,

Idea:

Make tractable using a hierarchical labeling scheme



²Feragen, Petersen, Owen, Lo, Thomsen, Dirksen, Wille, de Bruijne, *A hierarchical scheme for geodesic anatomical labeling of airway trees*, MICCAI 2012.

Performance:

- 40 airway trees from 20 subjects with different stages of COPD, hand labeled by 3 experts in pulmonary medicine.
- All 20 segmental labels were assigned (segmental = most distal branches) at an average success rate of 72.8%.
- Performance: as good as the performance of an expert in pulmonary medicine.

(measured in terms of ability to agree with the two other experts)

²Feragen, Petersen, Owen, Lo, Thomsen, Dirksen, Wille, de Bruijne, *A hierarchical scheme for geodesic anatomical labeling of airway trees*, MICCAI 2012.



COPD stage by GOLD standard (0=healthy, 3=severe)

Spearman: ($\rho = 0.22, p = 0.18$)

²Feragen, Petersen, Owen, Lo, Thomsen, Dirksen, Wille, de Bruijne, *A hierarchical scheme for geodesic anatomical labeling of airway trees*, MICCAI 2012. ass@diku.dk,

Performance:

2 scans per subject, registered for label transfer. Reproducible segmental labels on average:

- 14.0 (expert 1)
- 15.1 (expert 2)
- 15.2 (algorithm)

²Feragen, Petersen, Owen, Lo, Thomsen, Dirksen, Wille, de Bruijne, *A hierarchical scheme for geodesic anatomical labeling of airway trees*, MICCAI 2012.

Conclusion and discussion

- We have introduced a geometric framework for analysis of tree-shapes such as airways
- We have made proof-of-concept statistical experiments
- Distance computations are generally NP hard; we use heuristics
- We have utilized the tree-shape framework to automatically assign labels to anatomical airway trees
- This gives a fast and robust procedure with very few tuning parameters, which performs well in presence of COPD.

Future work

- Development of heuristics for tree geodesic computation
- More extensive statistical analysis of airway trees
- Statistics on individual branches based on branch labeling

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- Kernels on anatomical trees (for speed)
- Kernels on anatomical graphs