# Project IX: Big Data Meets Medical Physics Dosimetry

#### Foundations and Recent Advances

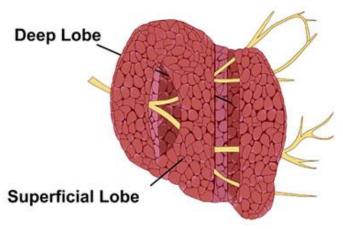
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## Topic

- Goal improve radiotherapy risk assessment through data mining.
- Formal term for risk measure is normal tissue complication probability (NTCP)
- Initial focus xerostomia (dry mouth) due irradiation of the parotid gland.



The Parotid Gland and Facial Nerve

Images courtesy of Dr.Todd McNutt, Dr. Scott Robertson

#### Literature

- Conventional method for computing NTCP, Lyman-Kutcher-Berman (LKB).
- This seminar presents four papers that form the foundation of LKB.
- Seminar includes a more recent paper incorporating PCA.

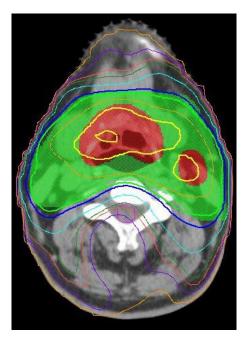


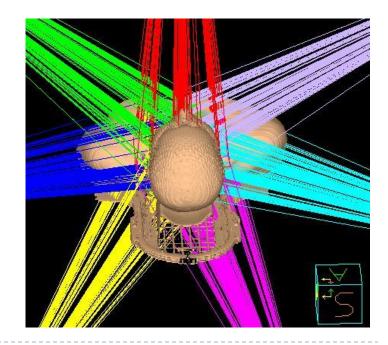
| ROI Statistics |          |        |           |        |        |        | % OI      |    |
|----------------|----------|--------|-----------|--------|--------|--------|-----------|----|
| 1              | Line Typ | e ROI  | Trial     | Min.   | Max.   | Mean   | Std. Dev. |    |
| Ŷ              |          | ptv1   | Conformal | 3568.6 | 7567.6 | 6252.6 | 896.3     | 0  |
| Ŷ              | -        | ptv1   | IMRT      | 3355.2 | 7366.9 | 6078.4 | 743.5     | 0. |
| Ŷ              | •••••    | ptv2   | Conformal | 5382.9 | 7541.1 | 7174.1 | 209.3     | 0. |
| Ŷ              | -        | ptv2   | IMRT      | 5049.4 | 7366.9 | 7016.2 | 128.2     | 0. |
| Ŷ              | •••••    | s.cord | Conformal | 132.1  | 4687.2 | 2449.1 | 1439.0    | 2. |
| <b>^</b>       |          | s.cord | IMRT      | 236,7  | 3957.8 | 1725.1 | 1138.0    | 19 |

Images copied from http://oftankonyv.reak.bme.hu/tiki-index.php?page=The+Clinical+Application+of+IMRT.

## Motivation for LKB

- The adoption of CT led to the emergence of 3D dose planning.
- Previous approach assumed uniform irradiation of the entire organ.
- NTCP assessment used TD<sub>50</sub> and TD<sub>5</sub> "tolerance doses" (Rubin & Casarett 1972).
- Need to calculate NTCP of non-uniform doses over parts of the volume.



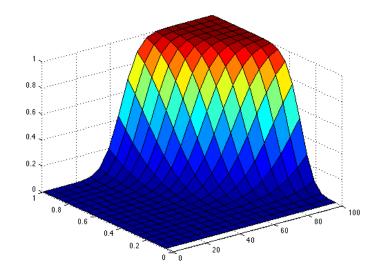


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### Lyman (1985) – The Model

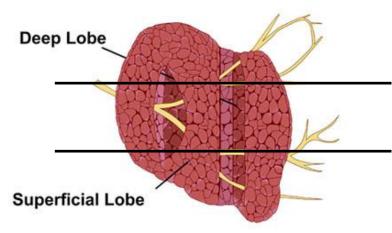
- Assumes a power equation relationship between whole and partial volume.
- Incorporates previous TD<sub>50</sub> approach.
- Requires parameterization of *n* and *m*.
- Accounts for partial volume, but still assumes uniform dose.

$$TD_{50}(V) = \frac{TD_{50}(1)}{V^{n}}$$
  
t ~ N(0, 1), t =  $\frac{D - TD_{50}}{\sigma(V)}$ ,  
where  $\sigma(V) = m[TD_{50}(V)]$ 



#### Emami, et al. (1991) – The Data

- Provides  $TD_{50}$  and  $TD_5$  values (estimates) for  $\frac{1}{3}, \frac{2}{3}$ , and whole volumes.
- Values from combination of literature review, hard data, estimates from clinical experience.
- For parotid, assume minimum 50% volume exposure for xerostomia.
- TD<sub>5</sub> = 3,200,  $TD_{50}$  = 4,600 for  $\frac{2}{3}$  and whole volumes;  $TD_{100}$  = 5,000.

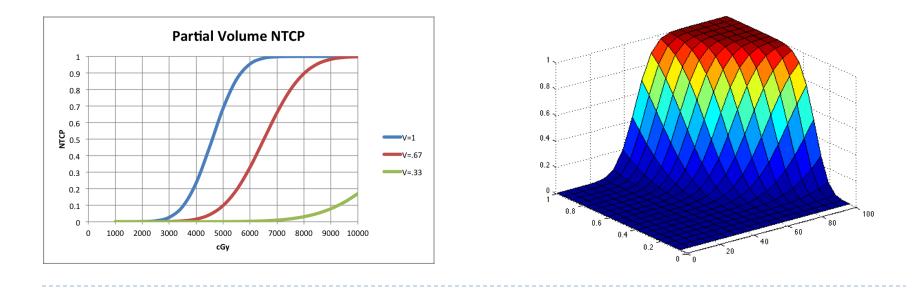


The Parotid Gland and Facial Nerve

Images courtesy of Dr.Todd McNutt, Dr. Scott Robertson

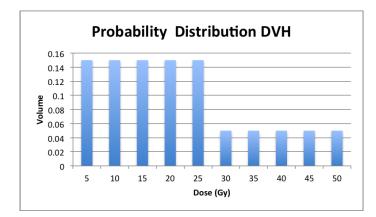
#### Burman, et al. (1991) – Parameters

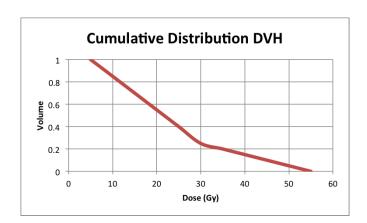
- Uses data from Emani, et. al. to calculate Lyman parameters.
- Values from combination of literature review, hard data, estimates from clinical experience.
- For parotid, assume minimum 50% volume exposure for xerostomia.
- TD<sub>50</sub> = 4,600, TD<sub>5</sub> = 3,200 for  $\frac{2}{3}$  and whole volumes; TD<sub>100</sub> = 5,000.



## Kutcher & Burman (1989) – DVH

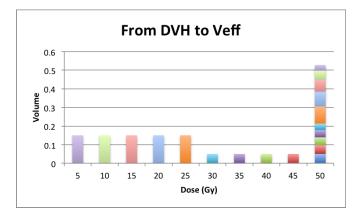
- Replace uniform with dose volume histogram (DVH).
- DVH may represent probability distribution or cumulative distribution.
- DVH removes spatial location.
- Typically visualized as cumulative distribution, "y% of the volume has received at least x dose."

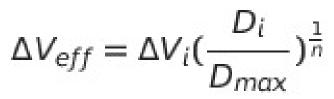




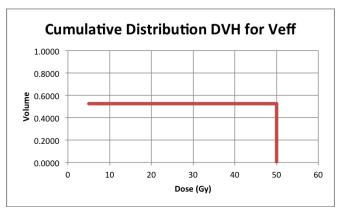
## Kutcher & Burman – $D_{max}$ , $V_{eff}$

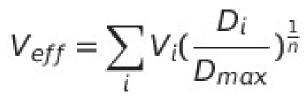
- Reduces DVH to a single pair of  $D_{max}$ ,  $V_{eff}$  values.
- Assumes high doses to small volumes are equivalent to smaller doses to larger volumes.
- Equivalence described by power relationship.
- Results in inputs for Lyman equation.





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## LKB – Summary

#### Pros:

- Adapts single dose whole volume TD<sub>5</sub> and TD<sub>50</sub> dose guidelines to NTCP values for non-uniform doses over a partial volume.
- Makes biological based assumptions.
- Only three parameters.

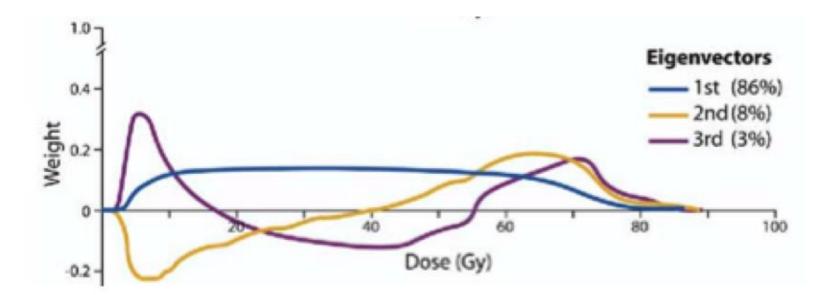
#### Cons:

- Does not account for treatment location.
- Conventional parameterization performed with scarcity of hard data.
- Using  $(D_{max}, V_{eff})$  removes a great deal of shape data from the DVH.

## Dawson et al. (2005) – PCA of DVH

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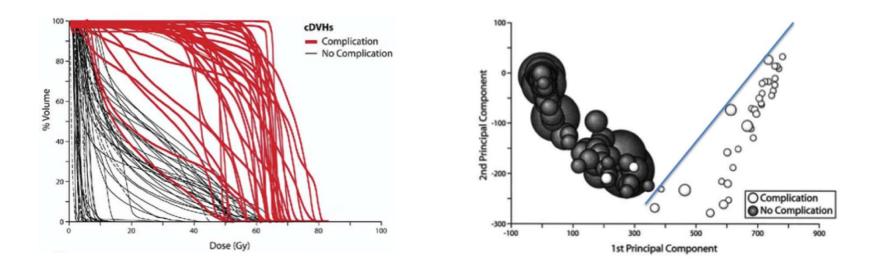
- No need for parameterization.
- For parotid, two PCs described 94% of the variance.
- Three PCs richly capture DVH shape.
- Possible improvement by centering and scaling data.



## Dawson et al. (2005) – PCA of DVH

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- PCA may provide better classification than  $D_{max}$ ,  $V_{eff}$ .
- Treatment based, not biologically based. PCs may not capture new protocols.



## Application to Project IX

- We wish to include dose location in NTCP calculation.
- Divide our regions of interest into 125 equal sized 3D rectangular volumes.
- Calculate DVHs on the whole ROI and each subregion.
- Reduce DVHs to two PCs for data mining task.

| DVHData |                  |  |  |  |
|---------|------------------|--|--|--|
|         | Х                |  |  |  |
|         | Υ                |  |  |  |
|         | Z                |  |  |  |
|         | roiDoseSummaryID |  |  |  |
|         |                  |  |  |  |

| Re | RegionsOfInterest |  |  |  |  |
|----|-------------------|--|--|--|--|
| 8  | ID                |  |  |  |  |
|    | name              |  |  |  |  |
|    | volume            |  |  |  |  |
|    | mask              |  |  |  |  |
|    | patientRepID      |  |  |  |  |
|    | Created           |  |  |  |  |
|    | Modified          |  |  |  |  |
|    | Active            |  |  |  |  |

Images courtesy of Dr.Todd McNutt, Dr. Scott Robertson

#### References

- \*Burman, C., et al. (1991). Fitting of normal tissue tolerance data to an analytic function. International Journal of Radiation Oncology\* Biology\* Physics, 21 (1), 123-135.
- \*Dawson, L. A., et al. (2005). Use of principal component analysis to evaluate the partial organ tolerance of normal tissues to radiation. International Journal of Radiation Oncology\* Biology\* Physics, 62 (3), 829-837.
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  International Journal of Radiation Oncology\* Biology\* Physics, 21 (1), 137-146.
- \*Kutcher, G. J., & Burman, C. (1989). Calculation of complication probability factors for non-uniform normal tissue irradiation: The effective volume method. International Journal of Radiation Oncology\*Biology\* Physics, 16 (6), 1623-1630.

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- \*Lyman, J.T. (1985). Complication probability as assessed from dose-volume histograms. Radiation Research, 104 (2s), \$13-\$19.
- Rubin, P., & Casarett, G. (1972). Direction for clinical radiation pathology. the tolerance dose. Tech. rep., Univ. of Rochester, NY.

## Thank You

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