



Novel Predictors for Type A Dissections: Ascending Aortic Biomechanics

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UNITED STATES
DEPARTMENT OF VETERANS AFFAIRS

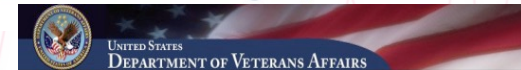


Disclosures

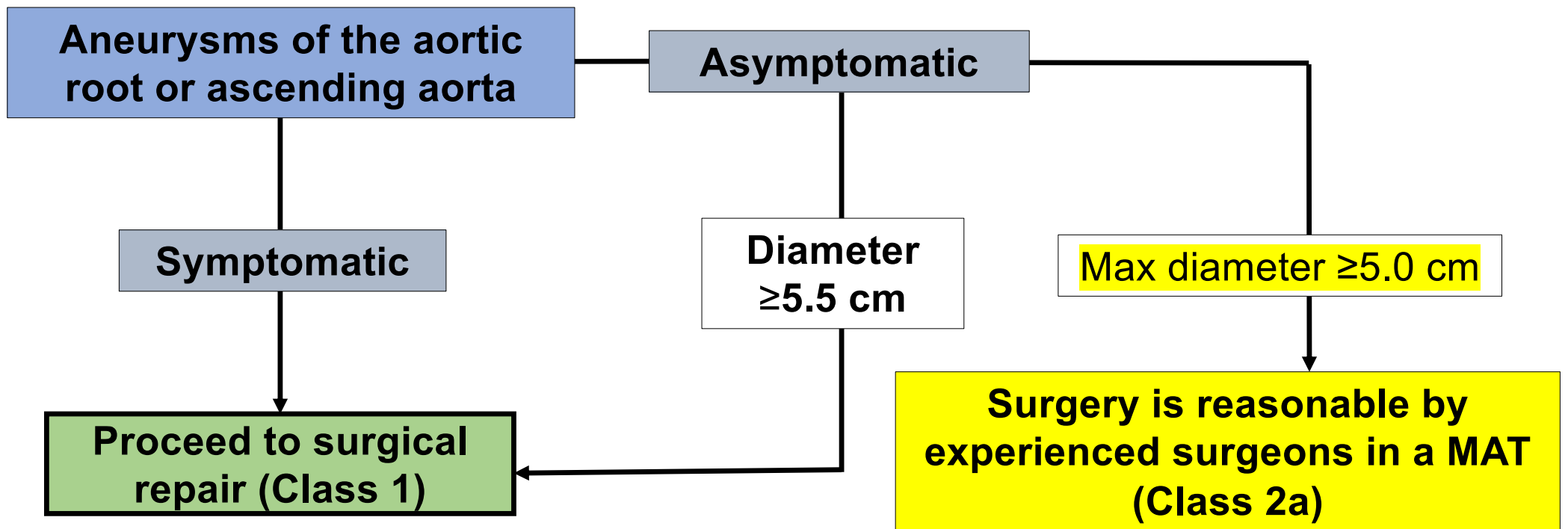
- No disclosures.



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Recommendations for Surgery for Sporadic Aneurysms of the Aortic Root and Ascending Aorta

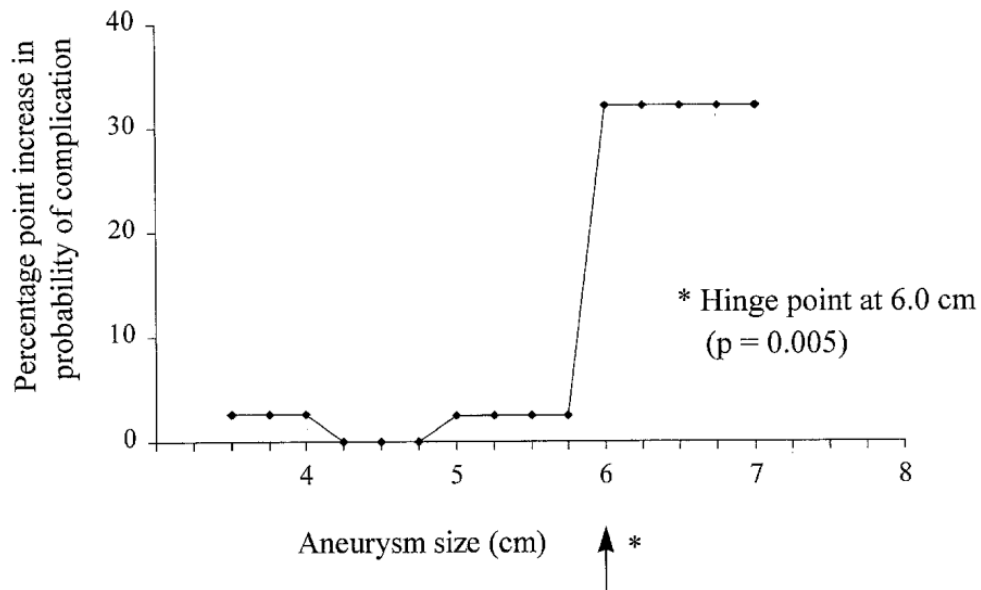


MAT, multidisciplinary aortic team; max, maximal;



Historic Aortic Event Rates

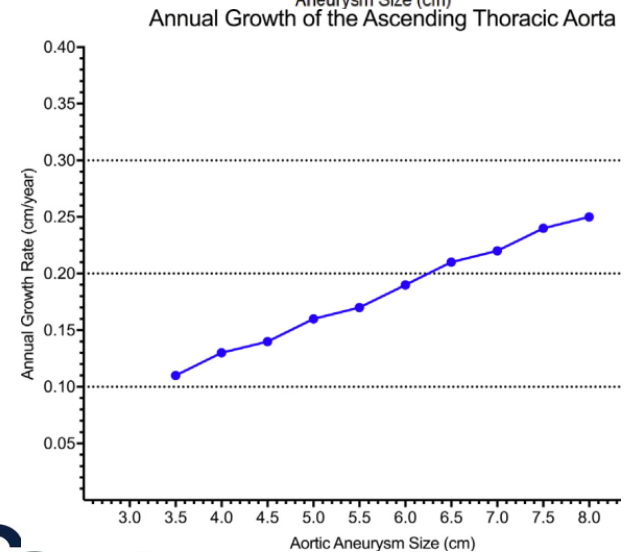
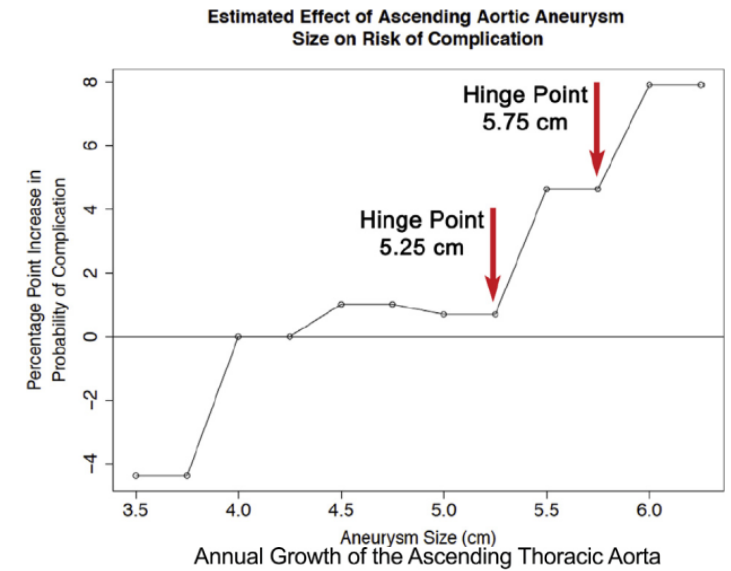
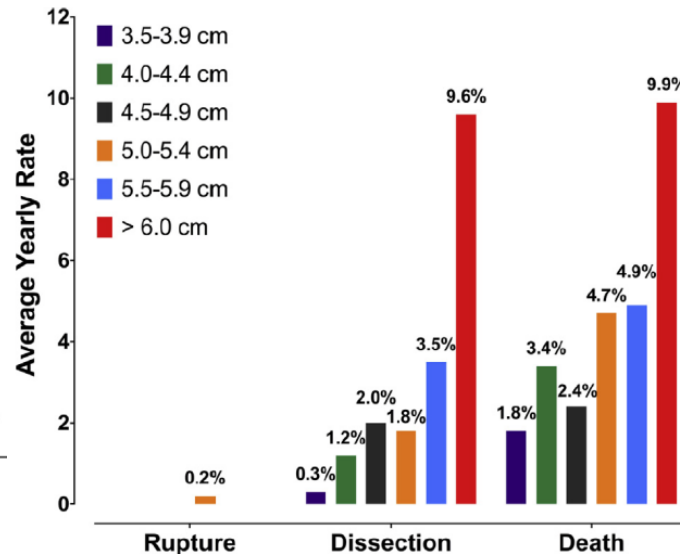
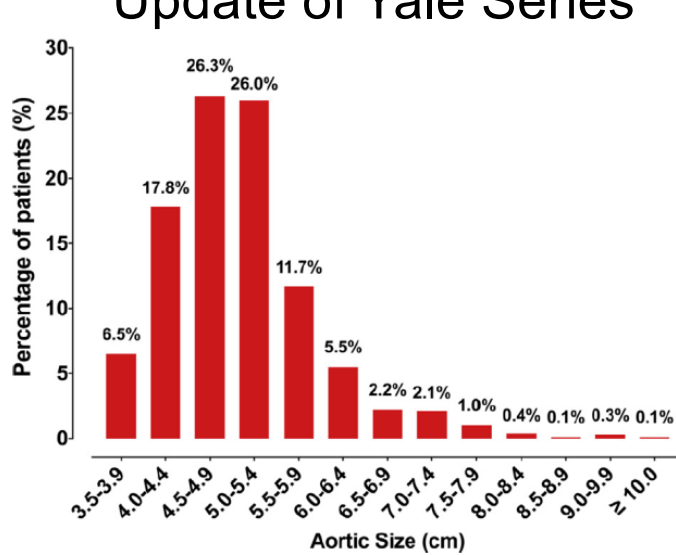
Original Studies: Yale 1985-1997



- 230 pts with TAA
- 111 pts with ascending TAA
- Hinge point at 6.0cm
- Increased probability of dissection or rupture relative to aneurysm size

Yale Aortic Event Rates

Update of Yale Series



- 3349 TAA pts: 780 aTAA pts (1272 size measurements), mean f/u 47.7 mo
- First hinge point 5.25cm, second 5.75cm
- Updated growth rate: 1.4mm/yr (females and Marfan risk factors for growth)



Zafar JTCVS 2018;155:1938-50

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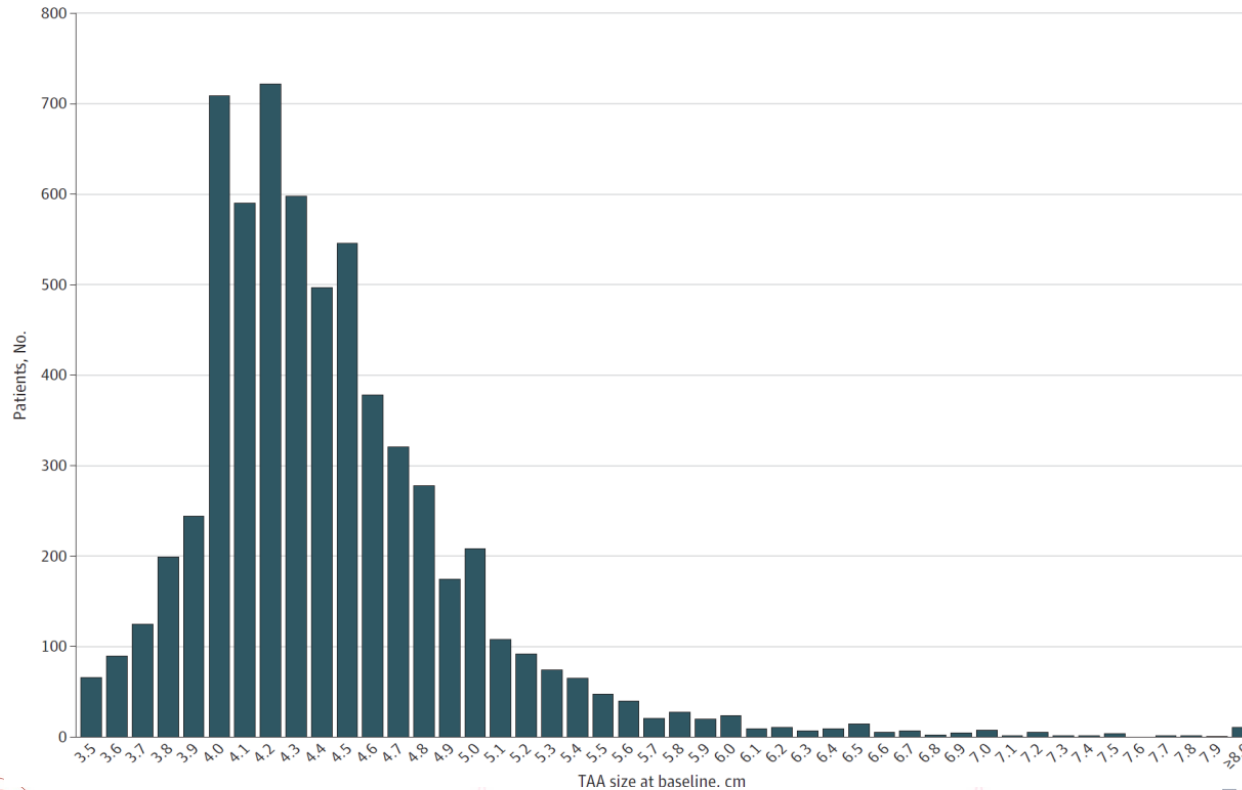
ATAA Size and Long-Term Outcomes

JAMA Cardiology | Original Investigation

Association of Thoracic Aortic Aneurysm Size With Long-term Patient Outcomes The KP-TAA Study

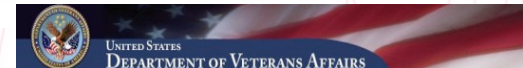
Matthew D. Solomon, MD, PhD; Thomas Leong, MPH; Sue Hee Sung, MPH; Catherine Lee, PhD; J. Geoff Allen, MD; Joseph Huh, MD; Paul LaPunzina, MD; Hon Lee, MD; Duncan Mason, MD; Vicken Melikian, MD; Daniel Pellegrini, MD; David Scoville, MD, PhD; Ahmad Y. Sheikh, MD; Dorinna Mendoza, MD; Sahar Naderi, MD; Ann Sheridan, MD; Xinge Hu, MD, PhD; Wendy Cirimele, BSN, MPA; Anne Gisslow, RN, MSN; Sandy Leung, RN; Kristine Padilla, RN; Michael Bloom, MA; Josh Chung, MD; Adrienne Topic, MD; Paniz Vafaei, MD; Robert Chang, MD; D. Craig Miller, MD; David H. Liang, MD, PhD; Alan S. Go, MD; for the Kaiser Permanente Northern California Center for Thoracic Aortic Disease

- Kaiser-Permanente TAA study: 6372 aTAA pts from 2000-2016
- TAA ≥ 3.5 cm, >18 yrs, no aortic surgery or dissection, no genetic aortic disease
- 32% women, 68% men
- Mean TAA size at dx: 4.3cm
- Mean f/u 3.7 yrs
- BAV 11%

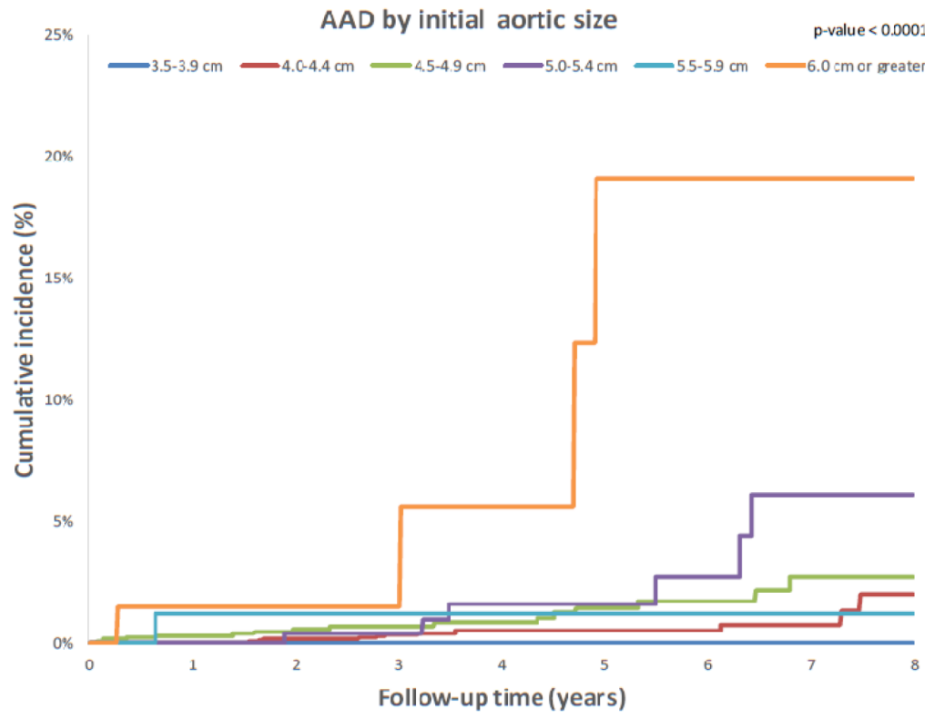


Solomon JAMA Card 2022;7(11):1160-9

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ATAA Size and Long-Term Outcomes

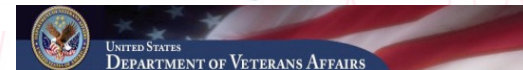
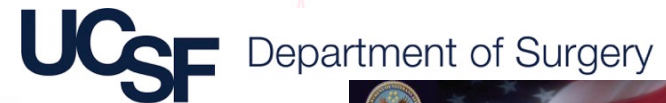


Patients at risk	0	1	2	3	4	5	6	7	8
3.5-3.9 cm	726	630	503	400	304	219	155	76	43
4.0-4.4 cm	3120	2604	1954	1416	990	681	454	222	96
4.5-4.9 cm	1698	1312	996	782	566	400	285	164	95
5.0-5.4 cm	548	323	239	186	136	104	70	40	26
5.5-5.9 cm	156	76	54	40	31	25	22	12	7
6.0 cm or greater	124	45	33	24	16	12	8	4	1

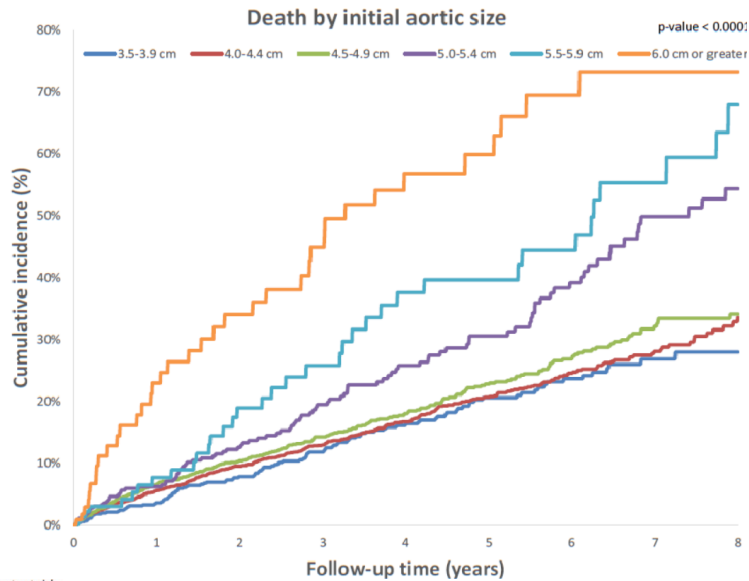
- Aortic dissection 44pts (0.7%)
 - Only 1 BAV (1/747pts)
 - 27 Ascending, 13 Root, 4 Arch
 - 37 dissection, 3 IMH, 4 rupture
- Overall Incidence 0.22/100pt-yrs
- Median age at AD 75.2yrs (63.7-81.0yrs)
- No AD <4.0cm
- Dissection rates:
 - >4cm <6cm <0.5/100pt-yr, <1%/yr
 - ≥6cm 2.19/100pt-yr
- Hinge point 6cm for AD and AD + all-cause mortality



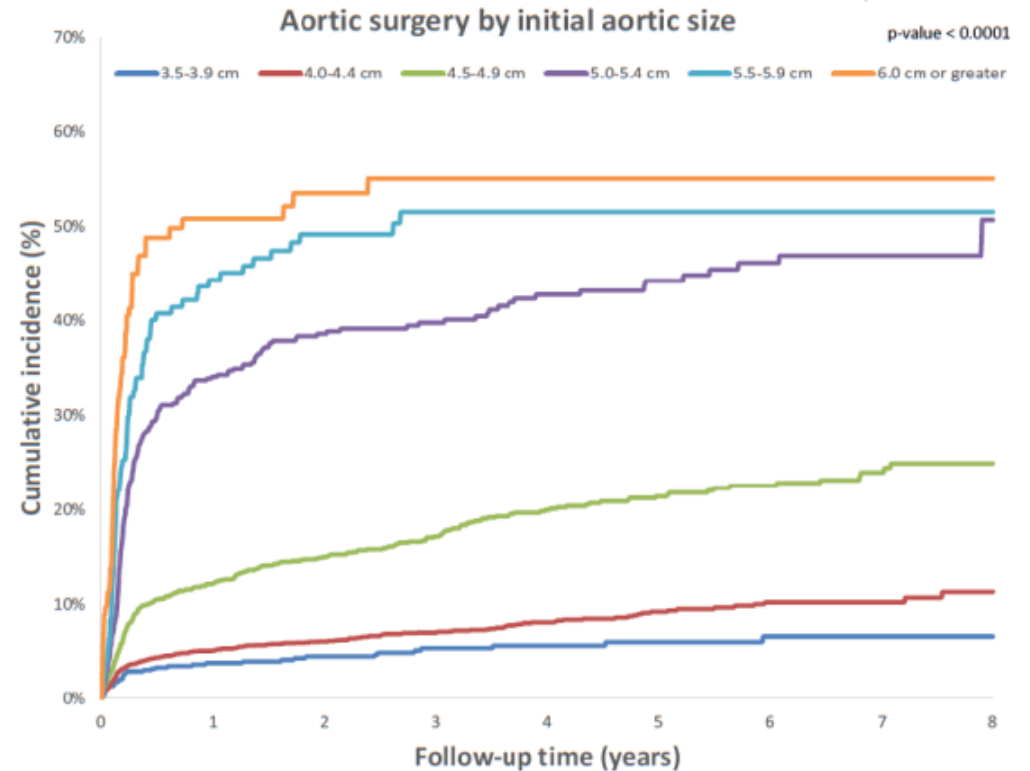
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ATAA Size and Long-Term Outcomes



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5.5-5.9 cm	156	76	54	40	31	25	22	12	7
6.0 cm or greater	124	45	33	24	16	13	9	5	2



- Dissection rates by time-updated ATAA size: Rates doubled for each size category from 4.0-5.5cm

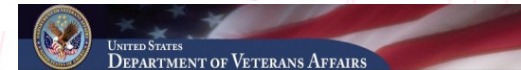
- <5.5cm AD <1%/yr

5.5-5.9cm AD 1.15/100pt-yr (~1% risk/yr)



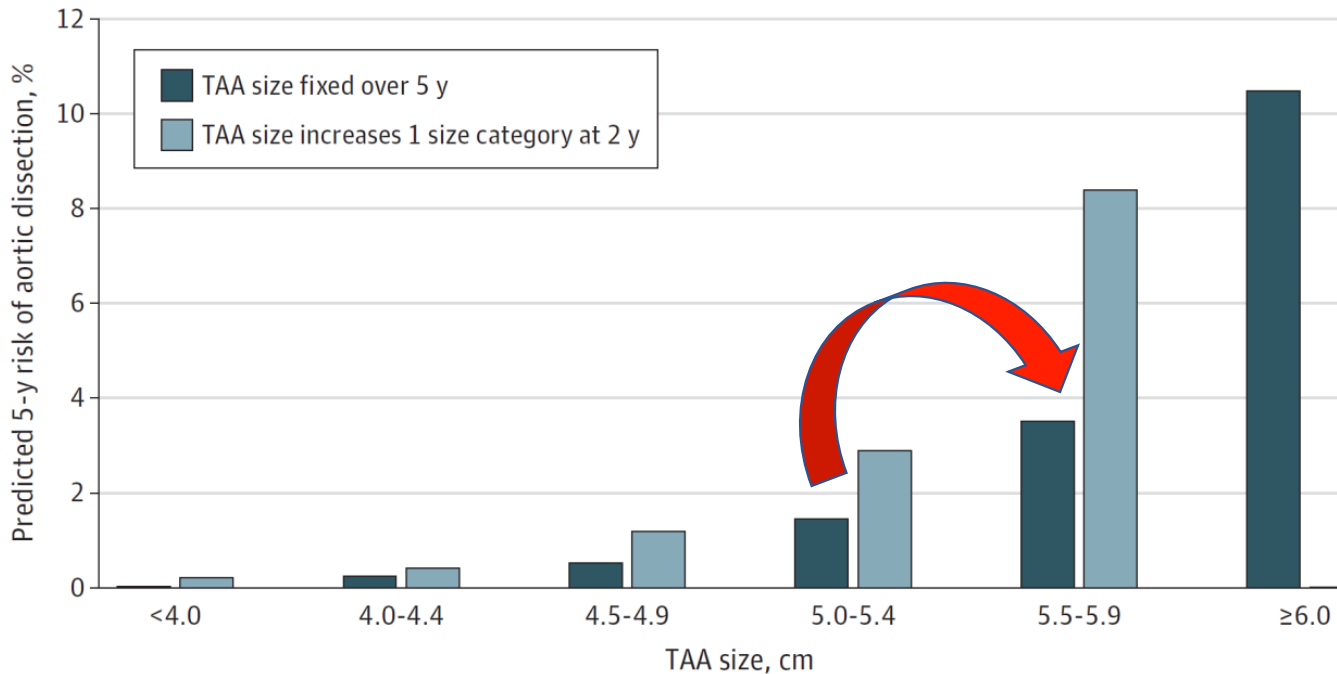
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ATAA Size and Long-term Outcomes

Predicted Risk of Aortic Dissection Over 5 Years Based on Size & Growth

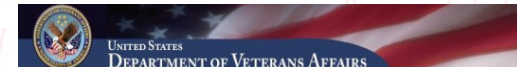
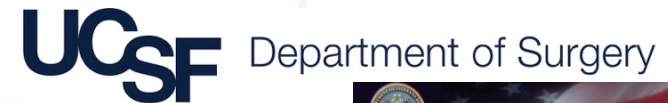


- Predicted risk AD in 5yrs:
 - 4-4.4cm 0.3% (CI 0.3-0.7)
 - 4.5-4.9cm 0.6% (0.4-1.3)
 - 5.0-5.4cm 1.5% (1.2-3.9)
 - 5.5-5.9cm 3.6% (1.2-12.8)
 - ≥6cm 10.5% (2.7-44.3)

- No difference in predictive power: diameter vs aortic size index

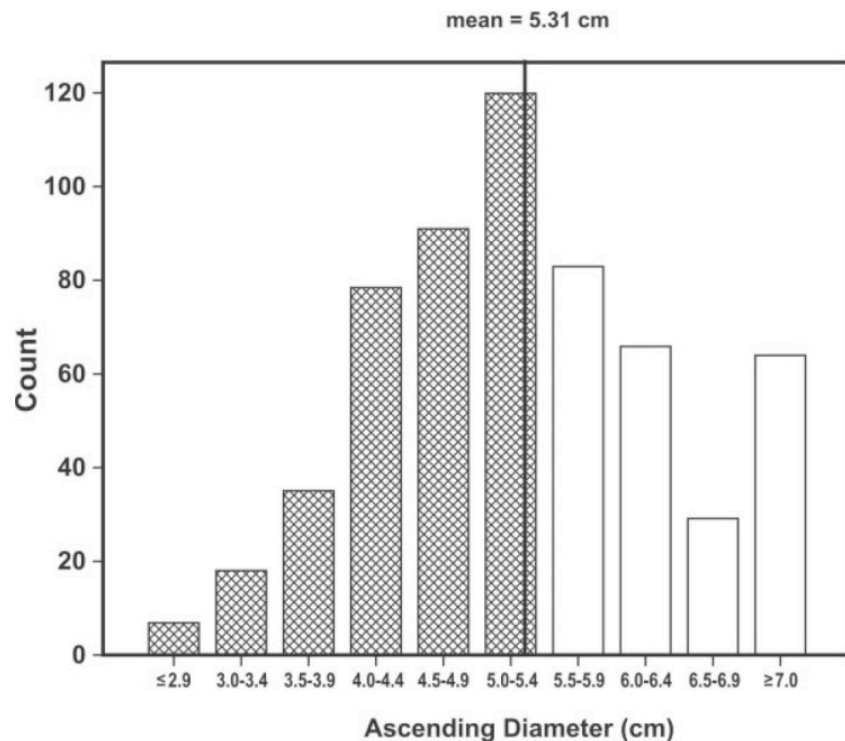


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TAA Size Paradox: 5.5cm Not Good Predictor of Type A Dissection

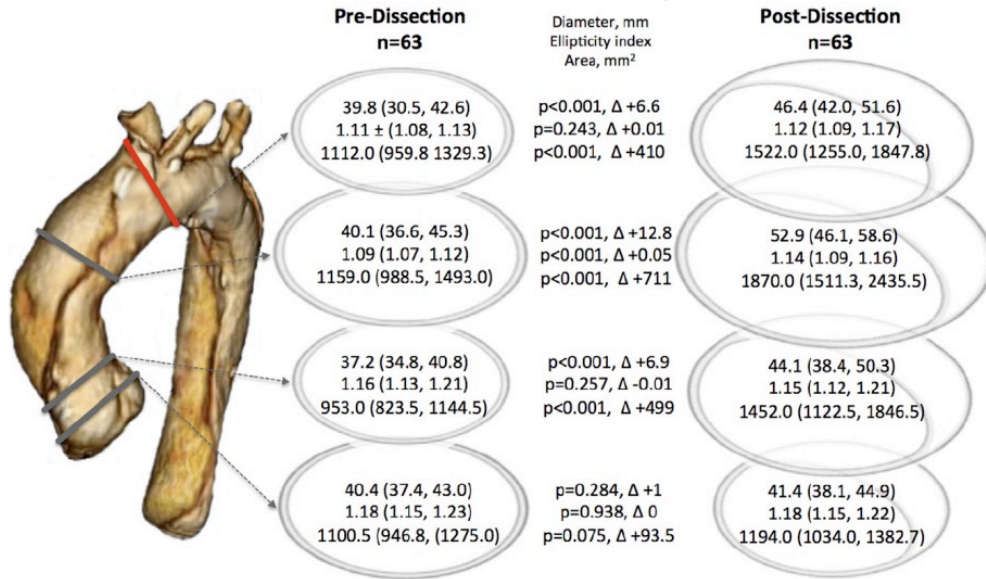
International Registry of Acute Aortic Dissection (IRAD)



- Distribution of aortic size at time of type A presentation (total n=591)
- 59% of pts <5.5cm
- 40% of pts <5.0cm

ATAA Size Paradox: Pre-Dissection Size

How Does the Ascending Aorta Geometry Change When It Dissects?



	All (n = 63)	Spontaneous AADA (n = 27)
Sinus of Valsalva	40.4 (37.4;43.0)	40.5 (36.7;46.2)
Sinotubular junction	37.2 (34.8;40.8)	38.3 (34.9;42.2)
Mid-ascending aorta	40.1 (36.6;45.3)	43.1 (36.8;47.9)
Proximal aortic arch	39.8 (36.1;42.6)	38.8 (36.8;43.9)
Distal aortic arch	33.8 (30.5;40.6)	31.1 (27.9;33.8)
Proximal descending thoracic aorta	36.0 (32.0;43.0)	32.4 (28.0;34.3)
Mid-descending thoracic aorta	32.5 (28.6;40.0)	28.3 (26.0;30.2)

- 6 tertiary centers across 2 continents with n=1821 type A dissection pts
- Excluded Marfan and BAV
- Included CTA <2 yrs before and within 12 hrs after dissection onset
- N=63 (27 spontaneous, 36 retrograde)
- Median age 68
- 54% men
- 62/63 max ascending diameter <5.5cm
- Prior to dissection largest diameter at mid-ascending with median 40.1mm (36.6, 45.3). Increased to 52.9 (46.1, 58.6)
- After dissection, 44% (28/63) ≥5.5cm and 60% (38/62) ≥5.0cm

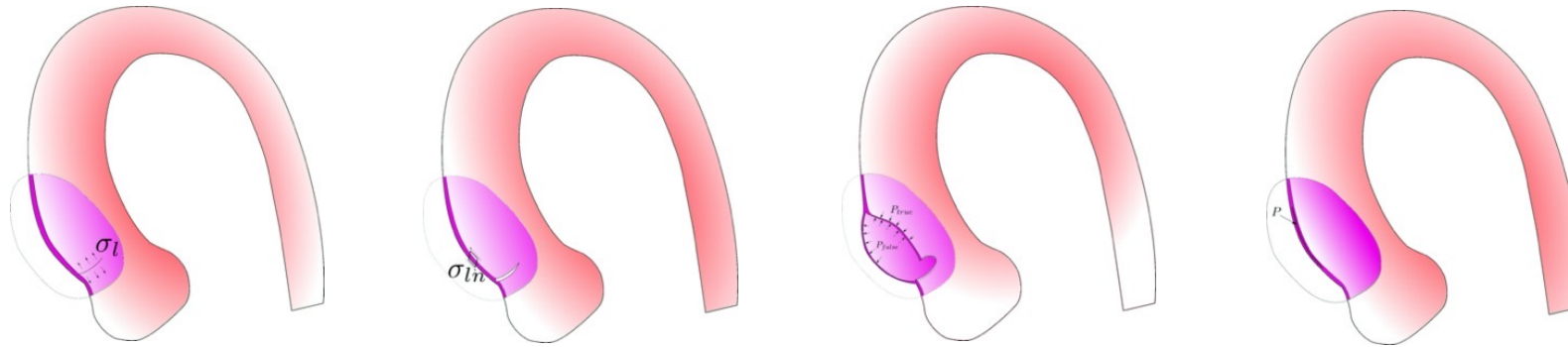


Rylski JACC 2014;63:1311-9



How Can We Better Risk Stratify aTAA Pts?

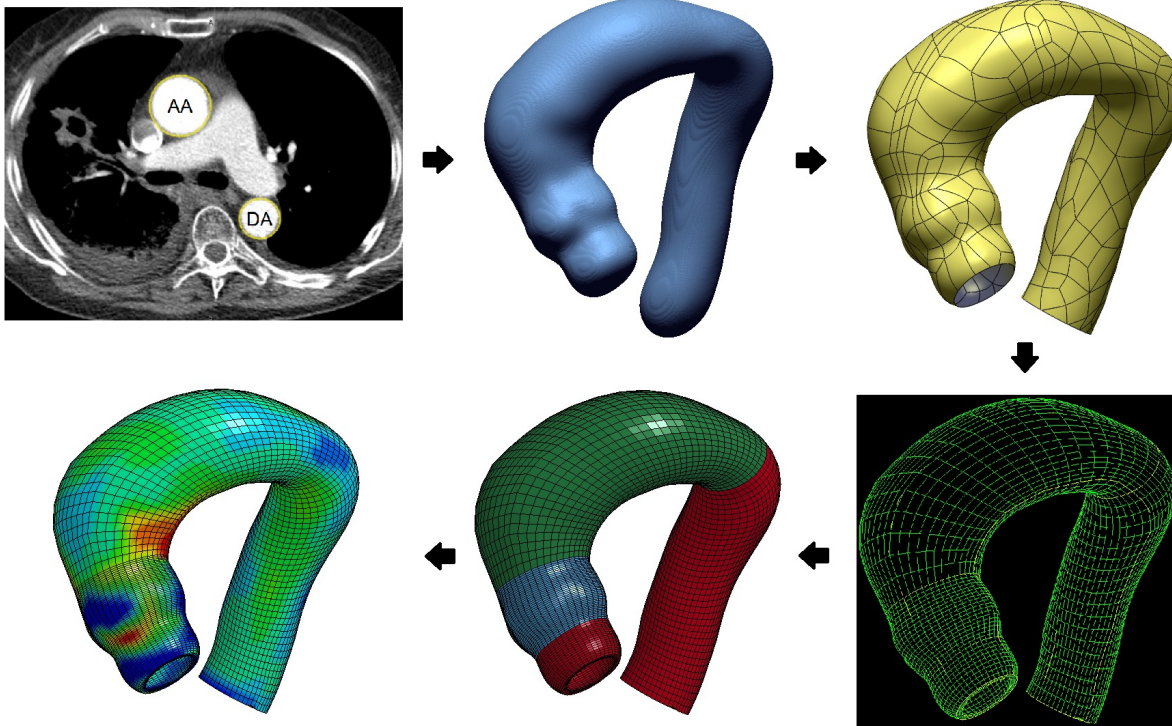
Aortic Dissection as a Biomechanical Failure



- The phenomena of dissection is a biomechanical event
 - Intimal tear in the aortic wall
 - Separation of the aortic layers
 - **Stress on aortic wall exceeds failure strength**
- The biomechanics approach seeks to quantify the physical forces on the aortic wall.

How Do We Better Predict Dissection?

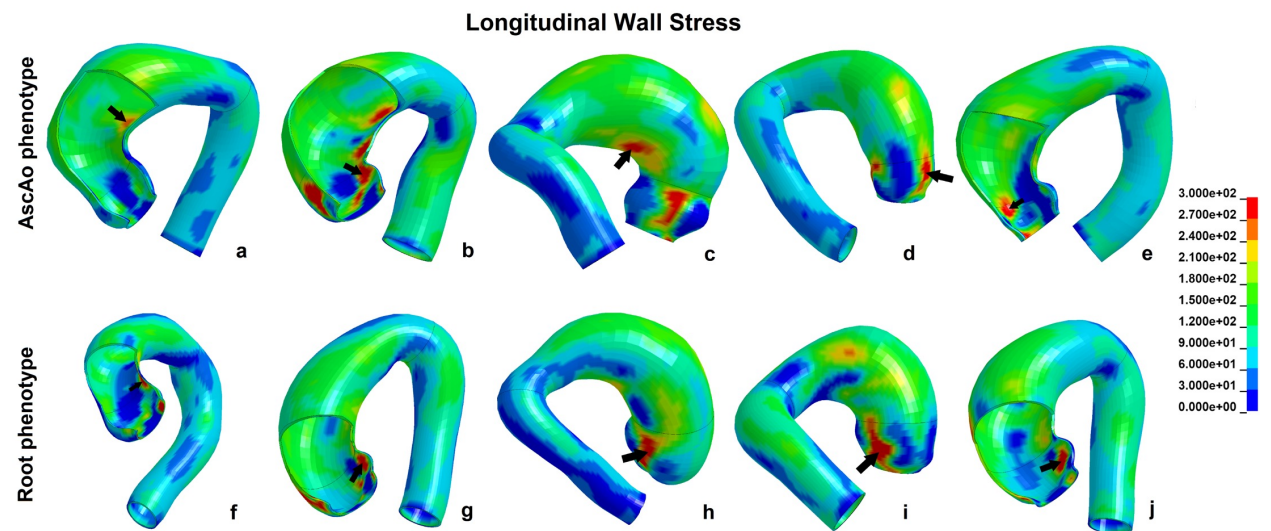
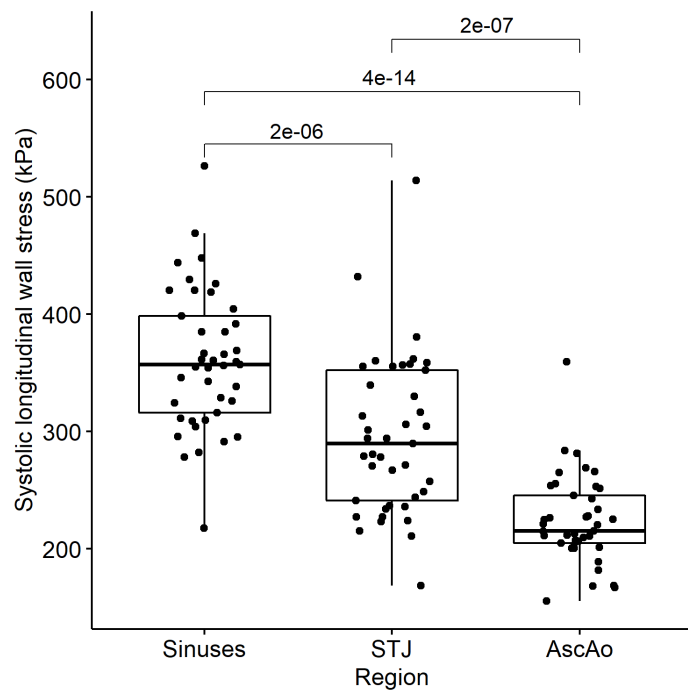
Patient-specific Ascending Thoracic Aortic Aneurysm Wall Stress > Wall Strength



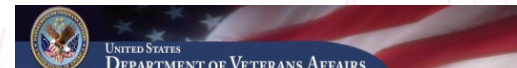
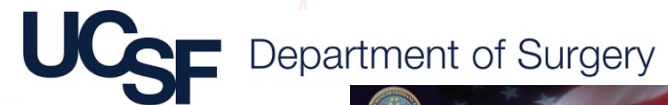
- Computed Tomography Imaging
- Image Segmentation and Geometry Reconstruction
- Geometry meshing
- Cardiac cycle simulated
- Finite element analyses
- Mechanical Failure: Aneurysm wall stress > wall strength

How Is Wall Stress Distributed?

- Bicuspid Aortic Valve (BAV)

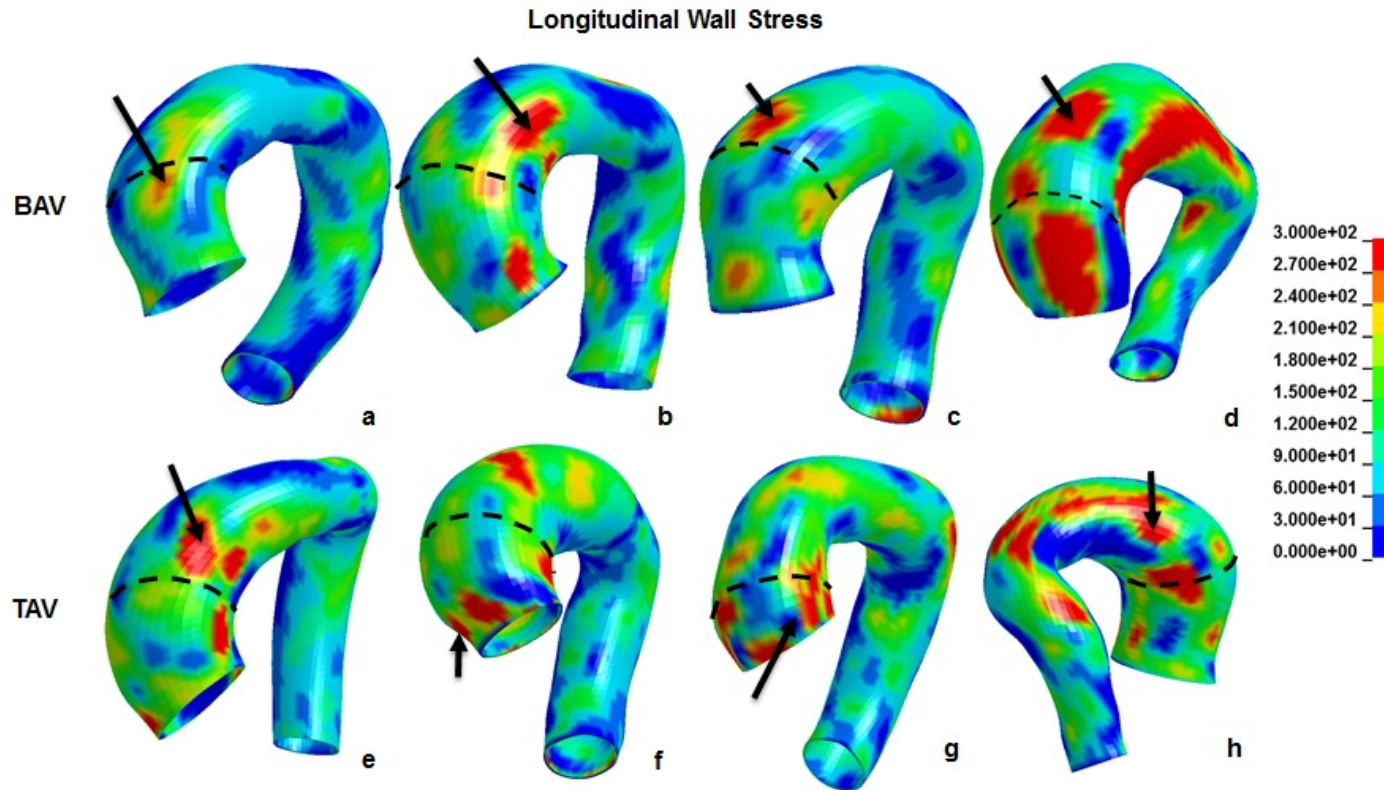


- Wall Stress in Sinuses > Sinotubular Junction > Ascending Aorta

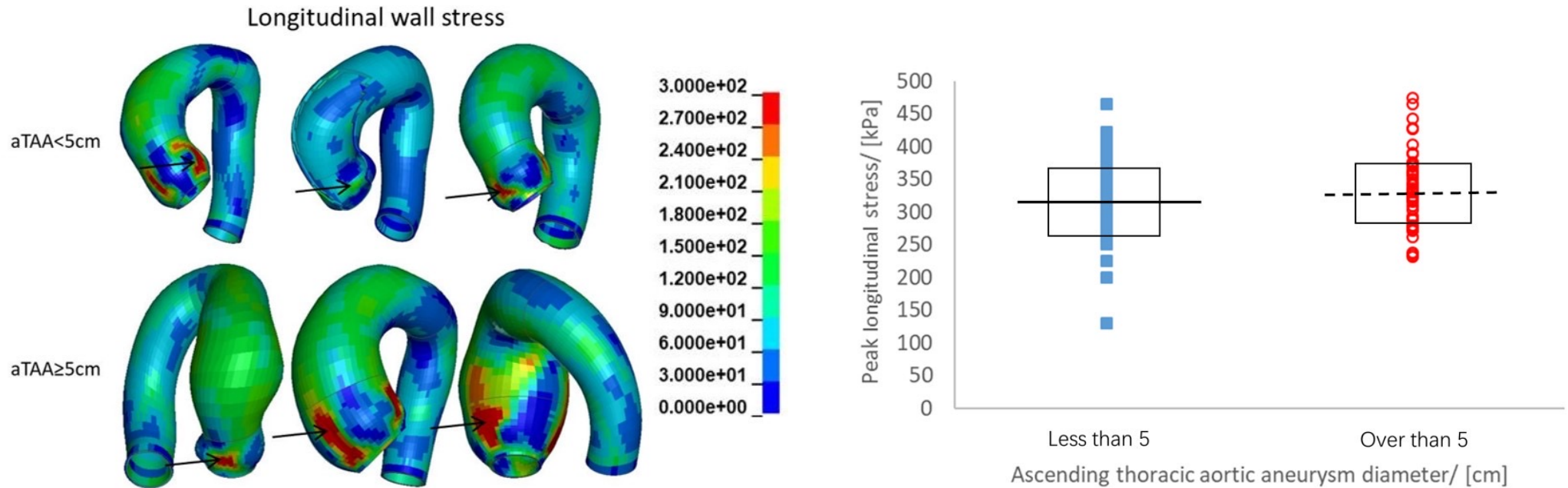


Where is Peak Wall Stress vs Max Diameter?

BAV and TAV Peak Wall Stress Is Not at Location of Max Diameter



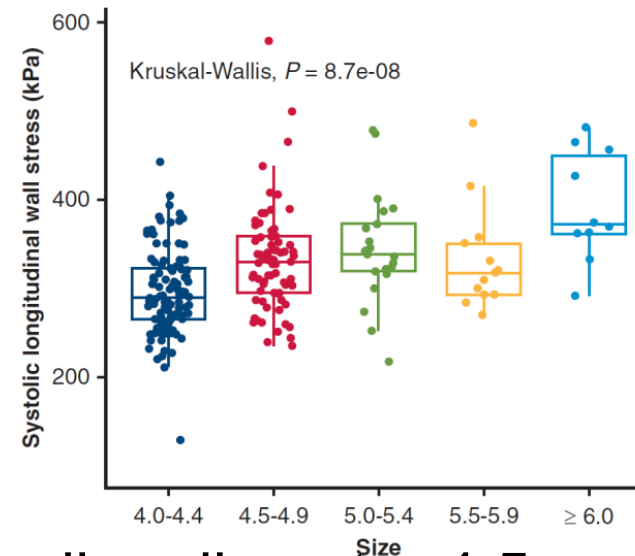
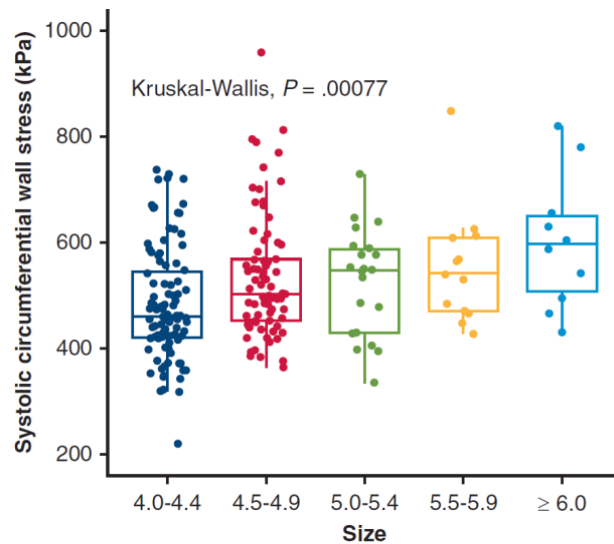
What is Wall Stress in $<5\text{cm}$ vs $\geq 5\text{cm}$ ATAA?



- Peak wall stresses do not differ significantly between $<5\text{cm}$ and $\geq 5\text{cm}$ aTAA.

Does Wall Stress Correlate with Diameter?

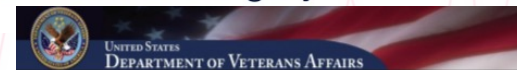
Tricuspid Aortic Valve (TAV) Aneurysm Peak Wall Stress Correlates with Diameter



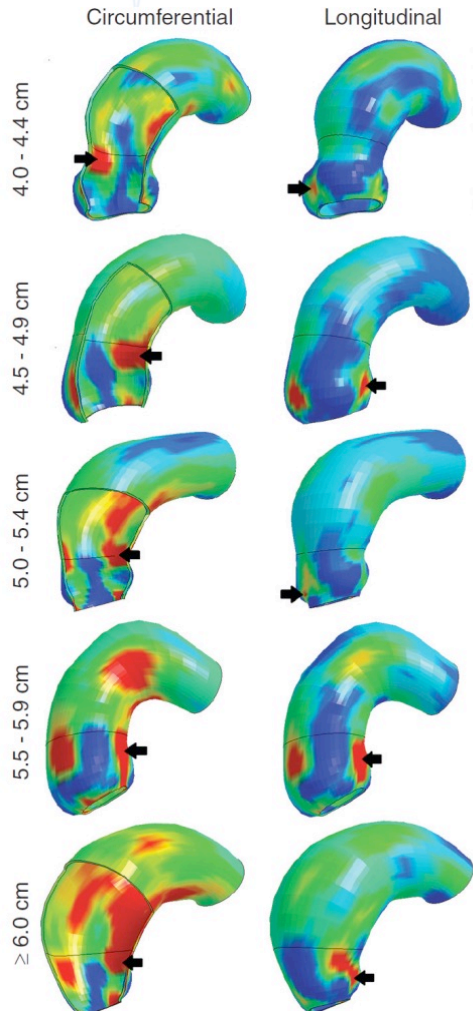
- N=221 veterans TAV aneurysms and median diameter 4.5cm.
- Both peak circumferential and longitudinal wall stresses increased with diameter ($p < 0.001$) in aneurysm as whole and each subregion, sinuses, STJ, and ascending Ao.
- Large overlap of stress ranges among groups.



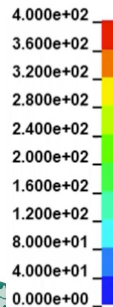
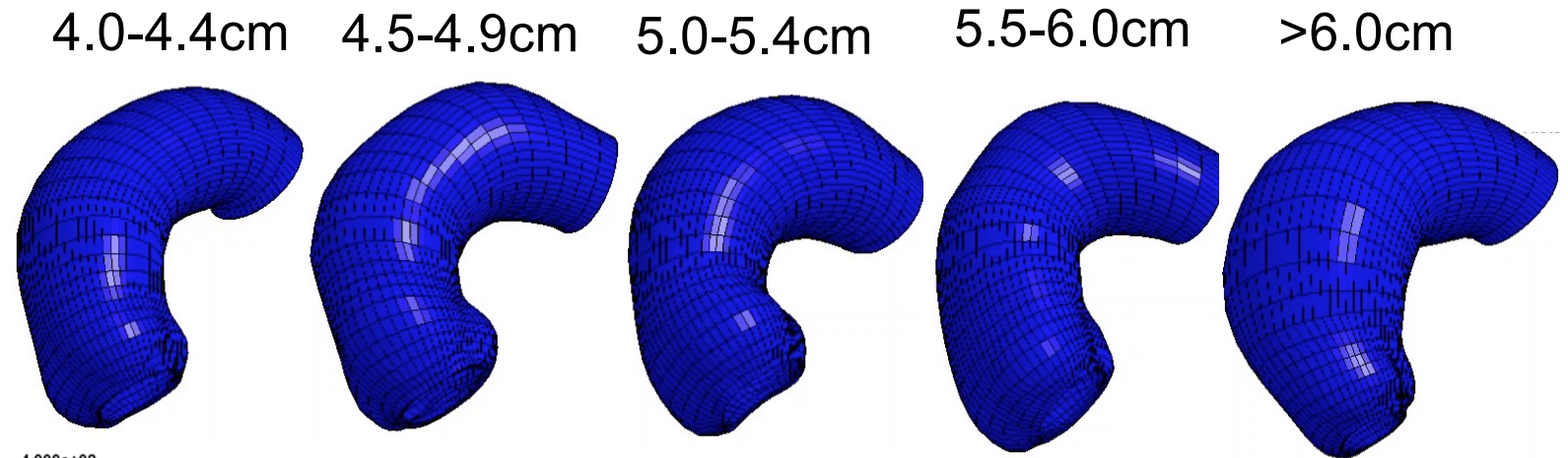
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TAV-ATAA Wall Stress FEA



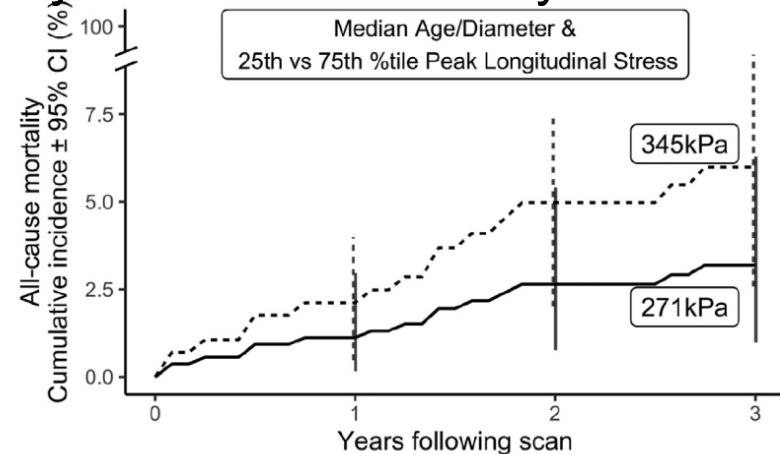
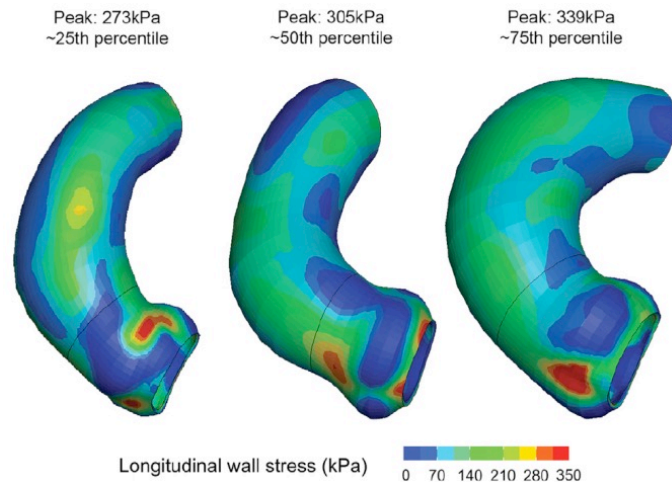
Representative Simulations Based on Diameter Ranges



- Circumferential >> longitudinal stresses.

Peak ATAA Wall Stresses & 3-yr Mortality

Peak Longitudinal Stresses Associated with 3-yr All-Cause Mortality



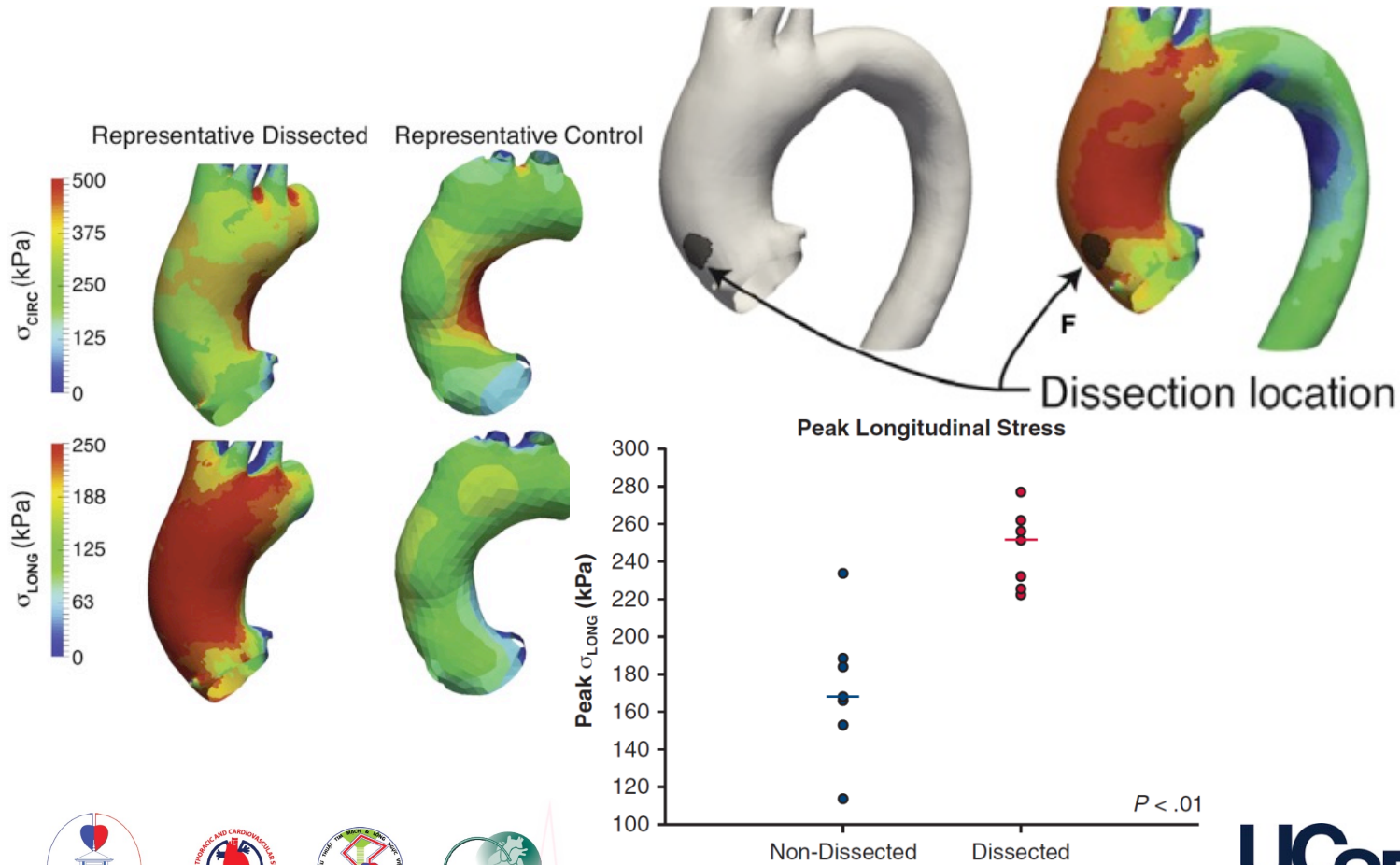
	0	1	2	3
Number at risk	273	214	171	148
Cumulative repairs	–	51	52	53
Cumulative deaths	–	6	13	15

- N=273 veterans, 17% BAV
- Median diameter 4.5 (4.3, 4.9). 9% (25/273 pts) \geq 5.5cm
- Median f/u 3 yrs.
- During f/u 19% (53/273) elective TAA repair with 0% mortality
- All-cause mortality 5.5% (15/273)



Peak Longitudinal Stress & Dissection

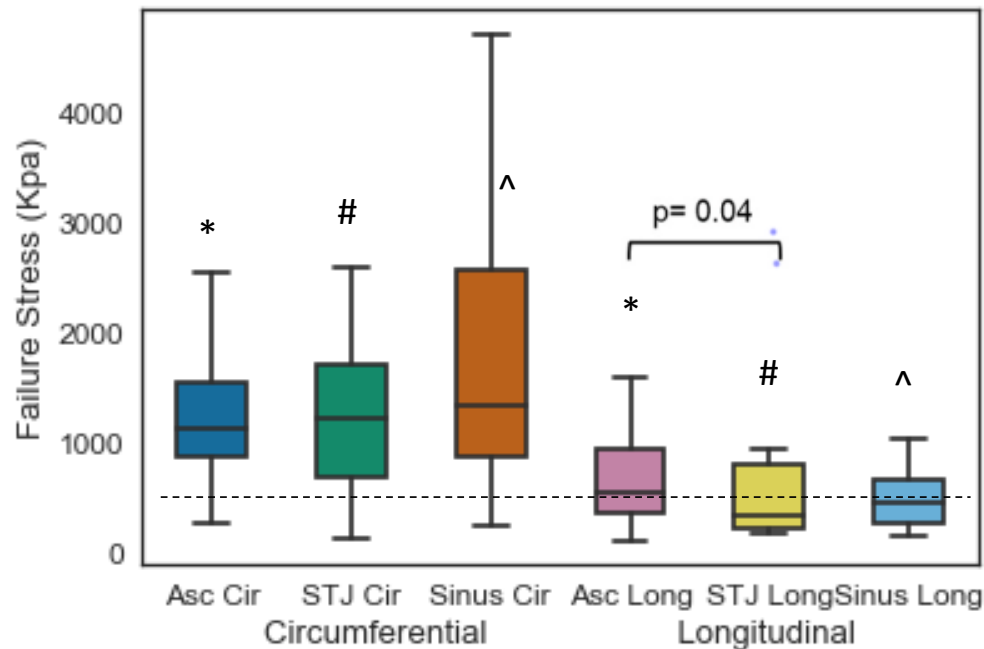
Peak Longitudinal but not Circumferential Stress Increased with Type A Dissection



- N=7 Dissection
 - Dissection CTA
 - ≥ 2 predissection CTA ≥ 1 yr before dissection
 - 1 predissection TTE
- N=7 controls
 - No BAV, CTD, aTAA
 - Matched Age, BSA

Why Longitudinal Stress and Mortality?

Failure Strength of the Aorta Lower in Longitudinal than Circumferential Direction

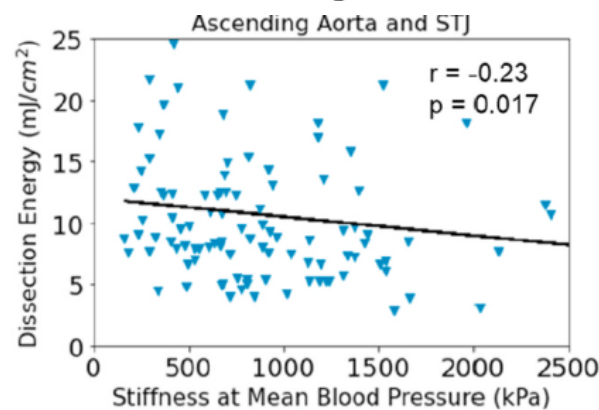
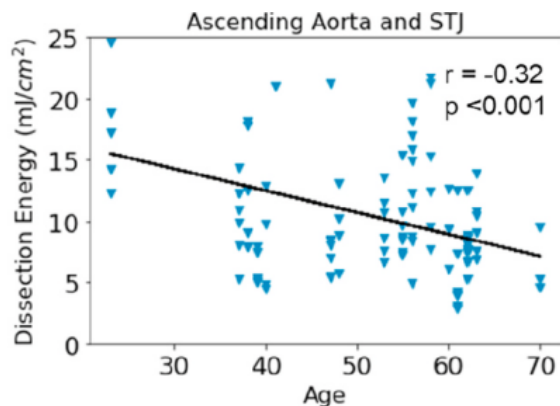
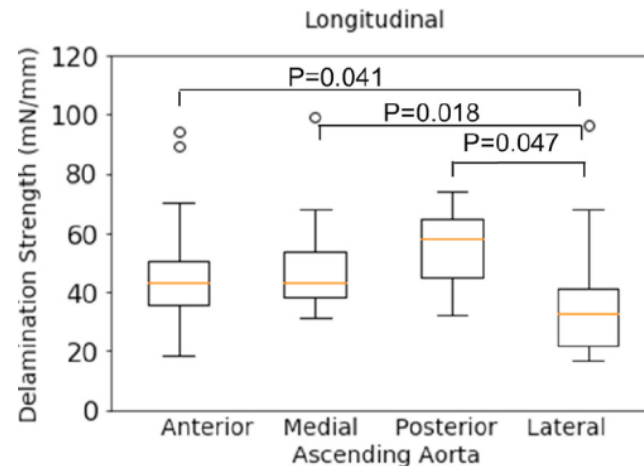
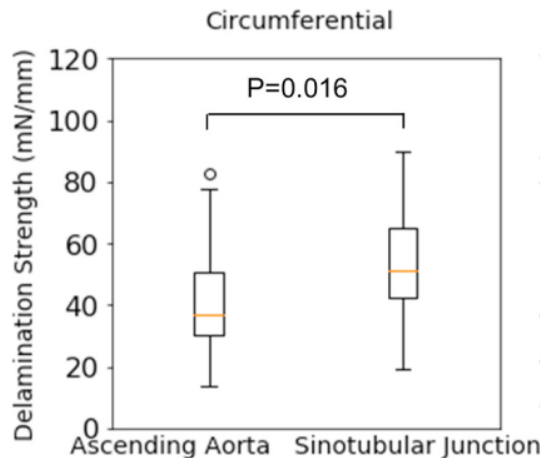


*, #, ^ p<0.01 circumferential vs longitudinal

- N=23 normal aortic roots
- 47% women
- Aorta is **weaker longitudinally** than circumferentially
- Longitudinal stresses result in transverse intimal tear
- Weakest region: **sinotubular junction (STJ)**

Dissection Peeling Properties of Aorta

Normal Aorta with Lower Delamination Strength in Ascending Aorta than STJ

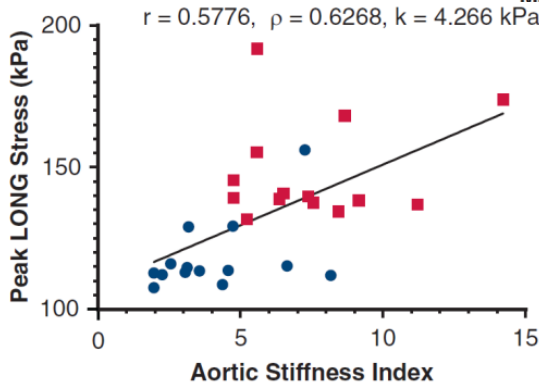
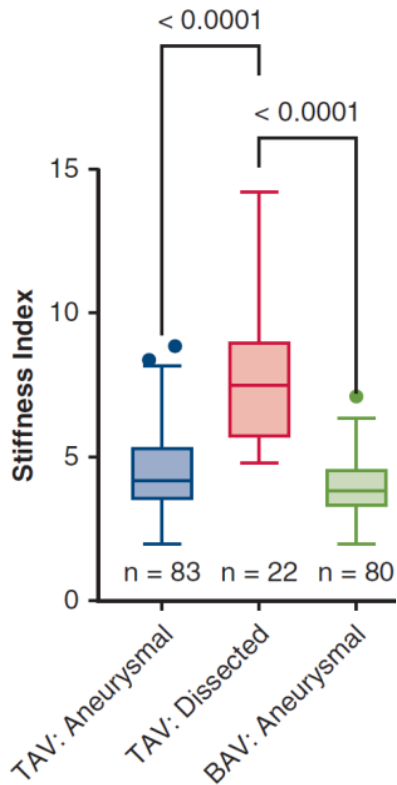


- N=19 normal aortas
 - 53% men
 - Mean age 51 yo
- Delamination strength circumferentially Ascending Ao < STJ
- Delamination strength longitudinally lowest in the greater curve Ao

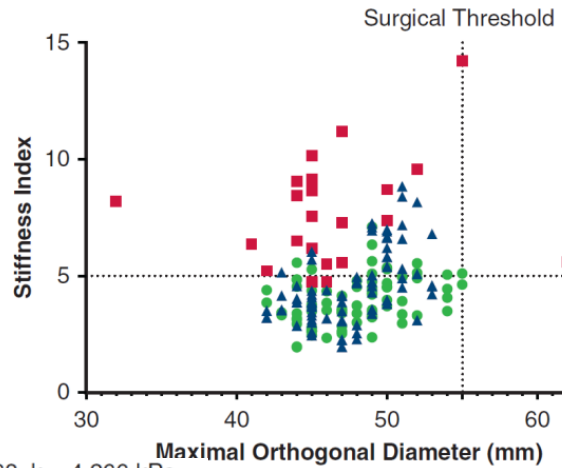


Aortic Stiffness Predicts Dissection in TAA

$$\text{Stiffness Index} = \frac{\ln(\text{SBP}/\text{DBP})}{\text{AoS} - \text{AoD}} \text{AoD}$$



● Non-dissected (n = 14) ■ Dissected (n = 14)



▲ TAV: Aneurysmal (n = 83)
 ■ TAV: Dissected (n = 22)
 ● BAV: Aneurysmal (n = 80)

N=22 Dissection, 73% men

- ≥1 TTE and ≥1 CTA predissection ≥1yr before dissection

• 46mm, Stiffness Ind 7.73

N=83 TAV pts, 83% men

- 48mm, Stiffness Ind 4.51

N=80 BAV pts, 65% men

- 48mm, Stiffness Ind 3.92

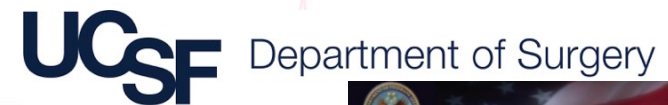
- Stiffness Index >5 predicted Aortic Dissection

- Stiffness Index Correlates w Peak Longitudinal Stress



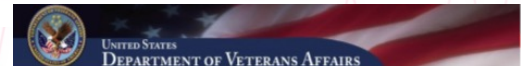
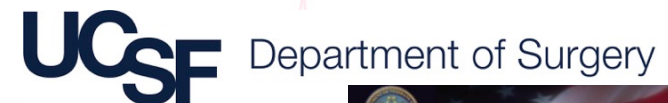
Conclusions

- Ascending thoracic aortic aneurysm dissection rates are lower than previous literature:
 - ~1%/yr for 5.5cm and <1%/yr for <5.5cm
 - Hinge point at 6cm
- Dissection Paradox: rate of dissection low at smaller sizes but most dissections occur at smaller sizes
 - Presentation of Dissection: ~60% are <5.5cm and 40% are < 5.0cm
- Given the dissection size paradox, better predictors of dissection are required than the present diameter based guidelines



Conclusions

- Peak longitudinal stress has been shown to correlate with type A dissection (n=7) and all-cause mortality (n=273)
- Aortic stiffness index > 5 may be predictive of type A dissection (n=22) and correlates with peak longitudinal stress.
- Circumferential delamination during dissection is easier in ascending aorta than STJ
- Longitudinal delamination is easier in greater curvature of ascending aorta
- Dissection energy is lower at older age and increasing aortic stiffness



Acknowledgements

- Funding:

- Marfan Foundation



- American Heart Association



- National Institutes of Health



- University of California San Francisco Summer Explore



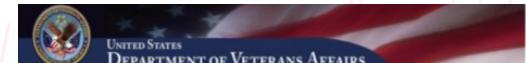
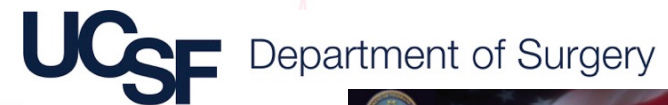
- Lab:

- Co-PI: Liang Ge, PhD

- Postdoc: Melody Xuan, PhD, Axel Gomez, MD

- Students: Siavash Zamirpour, Arushi Gulati, Shiv Verma, Preston Calloway

- Collaborators: Dimitrios Mitsouras PhD, Joseph Leach MD PhD, David Saloner PhD, Julius Guccione PhD





VA



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Thank you



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