

Clinical Decision Making In Asymptomatic Patients MedCAC 1/25/12

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Why treat Asymptomatic Carotid Stenosis?

- Carotid Stenosis is responsible for 20% of ischemic strokes (Abbott AL, et al Int J Stroke, 2007;2:27-39)
- Only 1/3 of strokes are preceded by warning TIA*
- Estimated 13 million Americans have silent cerebral infarctions*
- Stroke mortality is 10% @ 30D, 30% permanent disability, 20% Institutionalized at 3 months*

* AHA Heart Disease and Stroke Statistics -2011 Update, Circulation 2011;123:e11-209

Selecting Asymptomatic Patients to Treat

- Identify patients most likely to benefit from intervention
- Identify patients least likely to suffer harm from intervention
- Identify intervention (CEA, CAS, BMT) that has the lowest overall stroke and death rate

Asx CS and Stroke

- Stenosis: >60% - 2.3% / yr
> 75% - 4-5% / yr
- Plaque progression: 48% @ 1 yr
- Ulceration: 12.5% @ 1 yr
- Silent infarctions: 4.8% @ 1 yr
- Risk factor stratification : stenosis, age, BP, Contralateral symptoms, plaque character:
<5% to >20% / year

Survival important in estimating cumulative stroke risk

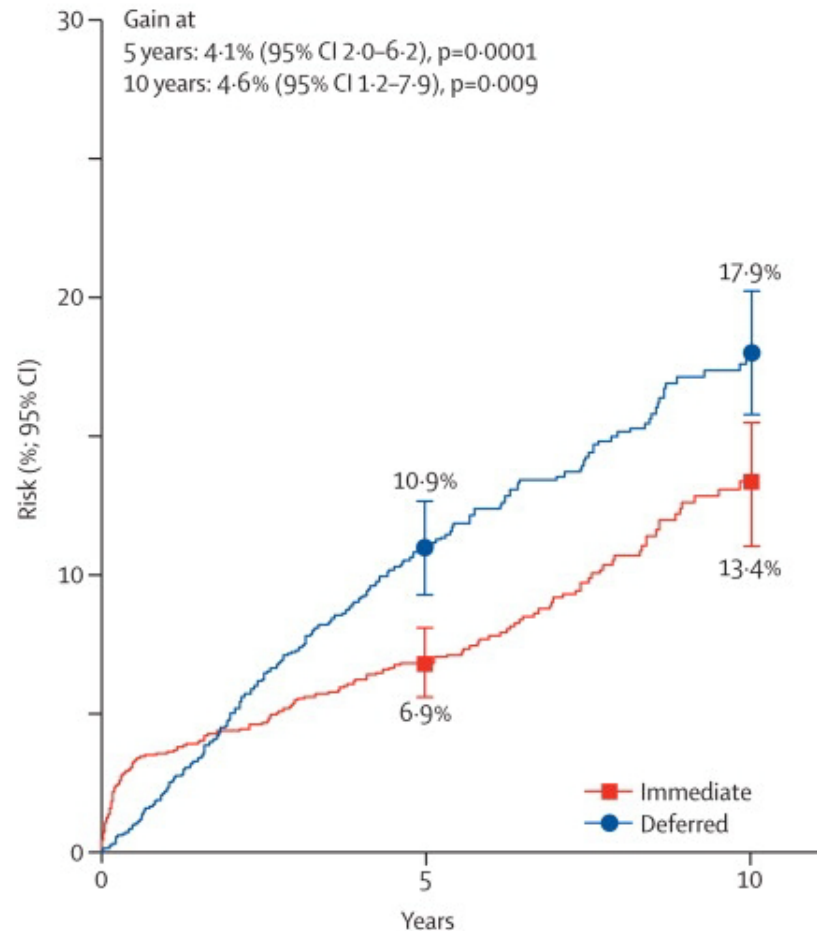
Long Term Results in Patients with Asymptomatic Carotid Stenosis: ACST 10 year Data

Halliday A, et al

Lancet 2010;376 1074-1084

ACST 10 Year Results

A Any stroke or perioperative death



Perioperative events/CEAs (%) + other events

Years 0-5
44/1509 (2.9%) + 56
14/360 (3.9%) + 140

Years 5-10
0/23 (0.0%) + 43
2/87 (2.3%) + 48

Immediate
Deferred

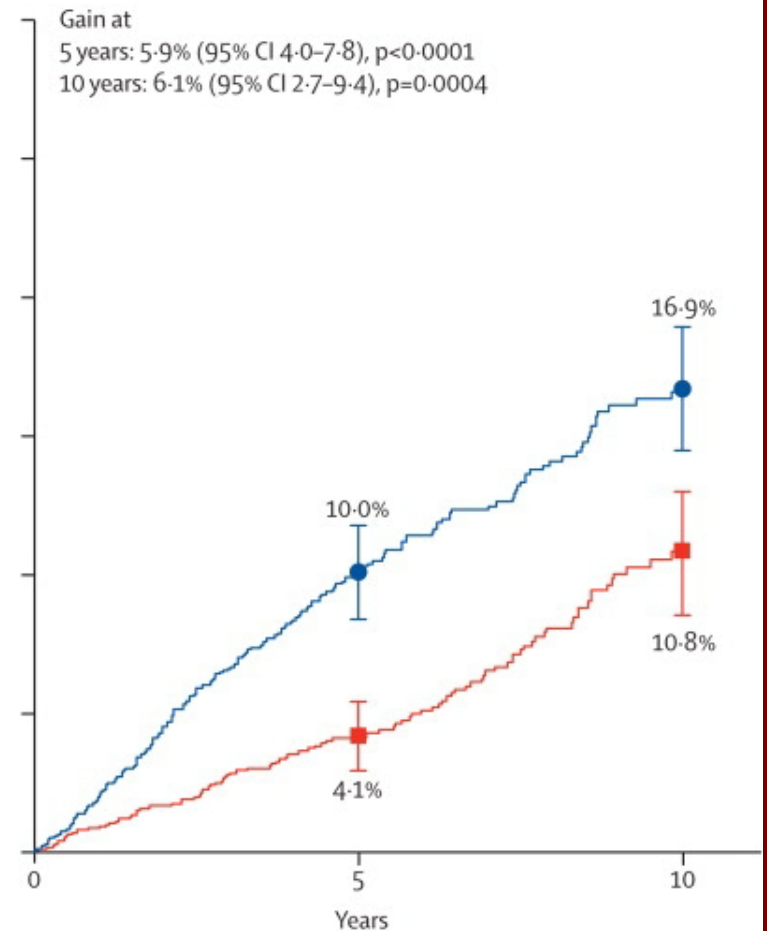
Number at risk

Immediate 1560
Deferred 1560

1003
981

293
281

B Any non-perioperative stroke



Events/person-years

Years 0-5
56/6540 (0.9% py)
140/6553 (2.1% py)

Years 5-10
43/3042 (1.4% py)
48/3003 (1.6% py)

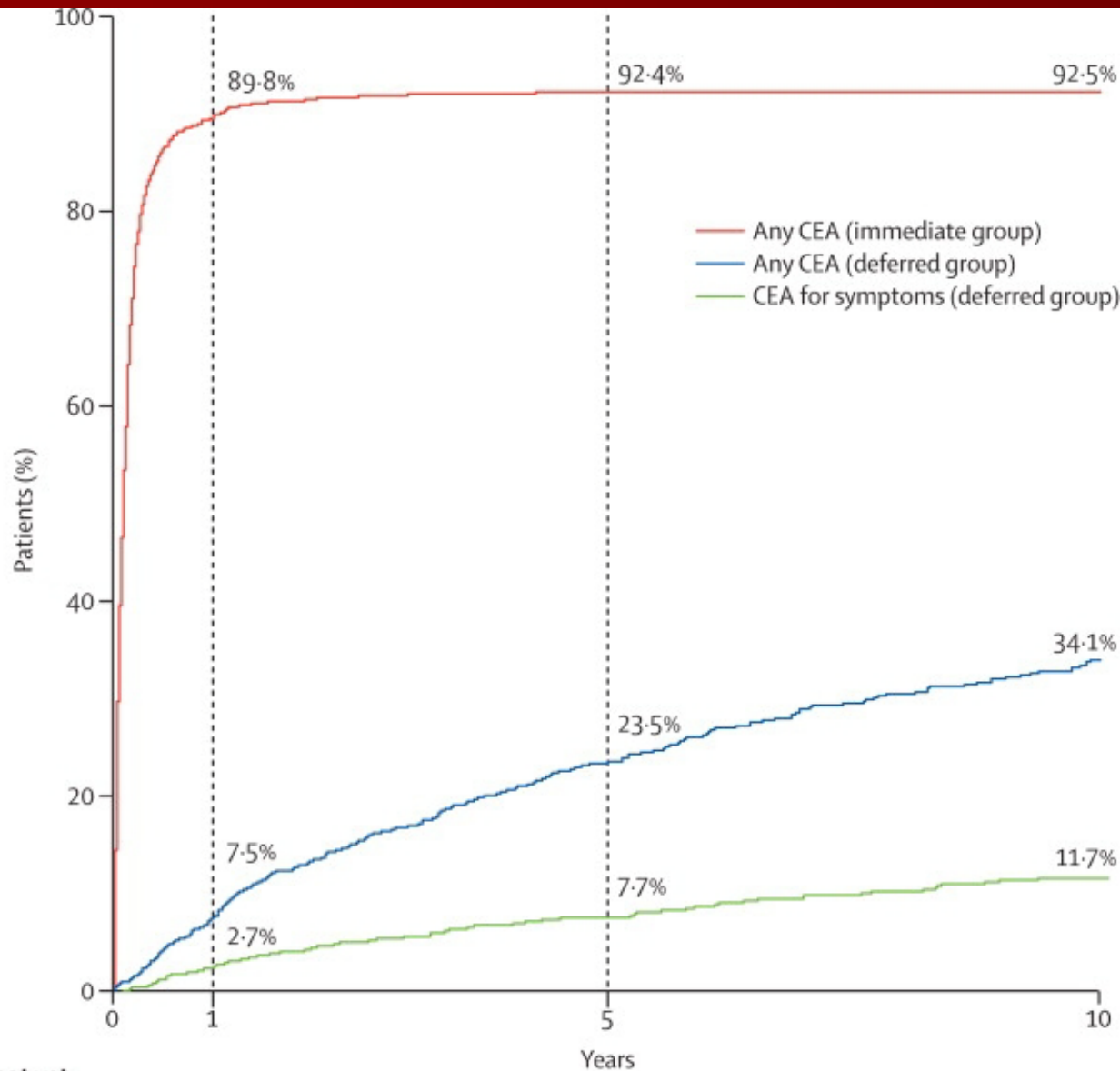
Immediate
Deferred

1560
1560

1003
981

293
281

ACST Incidence of Late CEA in Medical Arm

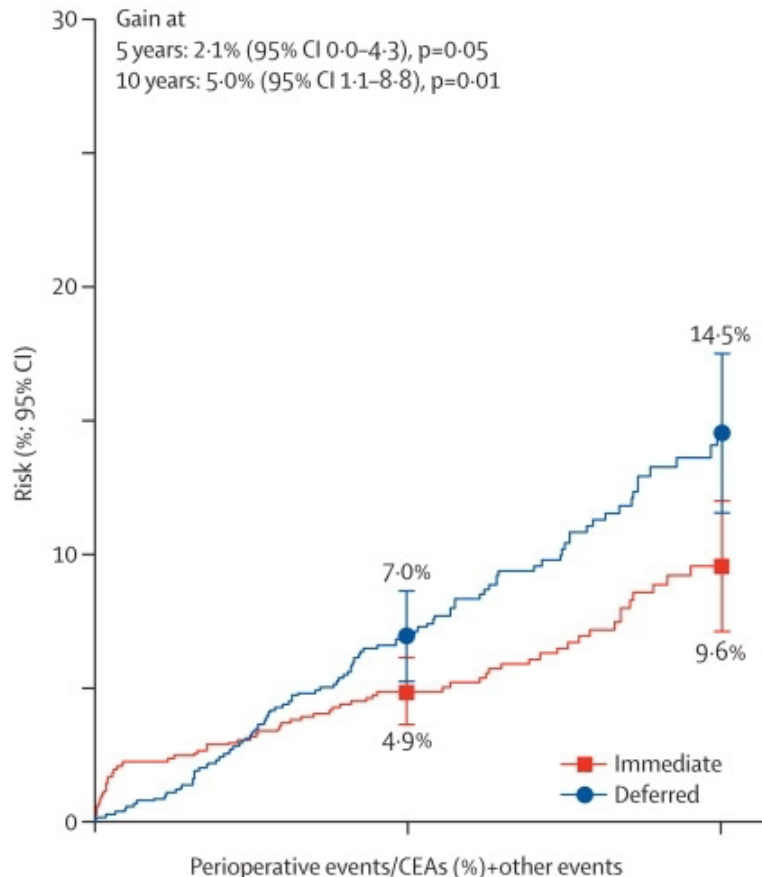


Number not yet
operated on

Immediate	1560	149	123	24
Deferred	1560	1386	818	205

ACST Risk Reduction - Lipids

A On lipid-lowering therapy before stroke:
stroke or perioperative death (mean age 68.0 years)

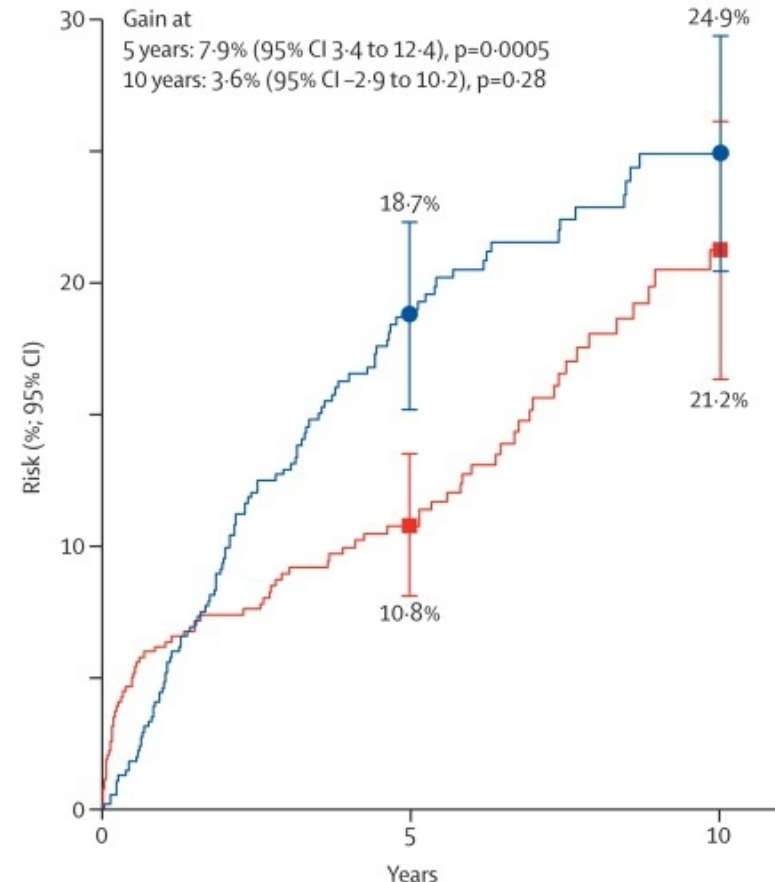


	Years 0-5	Years 5-10	
Immediate	22/993 (2.2%) + 25	0/15 (0.0%) + 20	Immediate
Deferred	9/259 (3.5%) + 56	2/69 (2.9%) + 32	Deferred

Number at risk

Immediate	1013	702	197
Deferred	999	697	176

C Not on lipid-lowering therapy before stroke:
stroke or perioperative death (mean age 69.6 years)



	Years 0-5	Years 5-10	
Immediate	22/516 (4.3%) + 31	0/8 (0.0%) + 23	Immediate
Deferred	5/101 (5.0%) + 84	0/18 (0.0%) + 16	Deferred

Number at risk

Immediate	547	301	96
Deferred	561	284	105

Selecting Intervention for Asymptomatic Stenosis

- Major morbidity after CEA is cardiac
- Major morbidity after CAS is stroke
- CREST suggested age affected outcomes
 - <70 favored CAS, >70 favored CEA
 - ? plaque character, arch, CAD
- Reducing Cardiac Morbidity should improve results of CEA
- Reducing Stroke Morbidity should improve results of CAS

Periprocedural risks: CEA vs CAS

(Murad et al JVS 2011;53:792-7)

Table II. Absolute risk difference per 1000 patients

<i>Outcome</i>	<i>RD (95% CI)</i>	<i>Quality of evidence</i>	<i>Interpretation</i>
Death	3.44 (-1.29, 11.44)	Moderate ^a	CAS is associated with 3 more deaths (from 1 fewer to 11 more)
MI	-10.15 (-13.17, -5.16)	High	CAS is associated with 10 fewer MI's (from 13 fewer to 5 fewer)
Stroke	18.77 (1.96, 42.23)	High	CAS is associated with 19 more strokes (from 2 more to 42 more)

CAS, Carotid artery stenting; CEA, carotid artery endarterectomy; CI, confidence interval; MI, myocardial infarction; RD, risk difference presented as a percentage with endarterectomy as a reference.

Analysis assumed the median control event rate from patients undergoing carotid endarterectomy in the included trials and relative risks from random effects meta-analyses.

^aQuality of evidence downgraded due to imprecision of meta-analytic estimate.

Factors Associated with Stroke risk in CAS

- Arch Configuration
- Access issues – angulation, stenosis
- Lesion Character – “echolucent” plaque, calcification, length >15mm, preocclusive stenosis
- Age

*A combination of these factors may increase
OR of stroke with CAS by 2.5-5.6*

(Settaci et al. Siena CA Stenting Score. Stroke 2010;41:1259-65}

Reducing Morbidity in Asymptomatic Patients

- **Cardiac Screening** – patients with active or occult coronary ischemia should be identified and treated before consideration for carotid intervention
- **Stabilize all medical conditions** – CHF, COPD, HTN, DM
- **Institute BMT** in all patients prior to intervention – antiplatelet, beta blockers, statins
- **Avoid intervention** in pts with severe comorbidities or limited life expectancy

Intervention in Asymptomatic Stenosis: CREST results

- 1182 Asx pts periprocedural S/D
CAS 2.5% CEA 1.4%
- Patients were good candidates for either intervention
- Intervention carried out by experienced CEA and CAS practitioners
- Results represent the best currently attainable

Selecting Intervention in Asymptomatic Patients

- Level 1 data supports CEA in Asx patients based on degree of stenosis even with relatively low event rate in the Medical Arm – based on low morbidity of CEA
- While “Best Medical Therapy” has improved, so have the results of Intervention – CREST results
- Improved patient selection should result in application of intervention to those most likely to benefit and least likely to suffer complications

Clinical Decision Making in Asymptomatic Patients

- Level 1 data supports CEA in pts with low periprocedural event rates
- Medical Comorbidities must be treated and stabilized for optimal results
- Only pts with significant life expectancy (3-5 years) should be considered for intervention
- Stratifying stroke risk among pts with $>60\%$ stenosis is an important goal
- Trials comparing BMT, CEA and CAS are needed