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# ERCAO Carotid Occlusion Recanalization Trial

#### Carotid stenosis

 Causing embolism, responsible for 20-30% of ischemic stroke

 Endarterectomy and stenting reduces future stroke rate, but contra-indicated only for total carotid occlusions (CAO)



# Why?

- With cessation of antegrade flow, the risk of embolic stroke is low
- \* Surgical bypass failed to show benefit
- It is rare, and endovascular recanalization is difficult and dangerous

#### No embolism does not mean safety

- Severe carotid stenosis or CAO leads to cerebral hypo-perfusion
- Annual stroke risk in patients with objective cerebral ischemia is as high as 20%



Klijn CJ, et al. Stroke 1997;28:2084 Grubb RL Jr, et al. JAMA 1998;280:1055 Derdeyn CP, et al. Neurology 1999;53:251

# Surgery provides no benefit

- \* Endarterectomy is not possible in CAO
- \* EC/IC bypass failed to yield benefit in CAO
  - High complication
  - \* Poor pt selection



EC/IC Bypass Study Group. N Engl J Med 1985;313:1191 Powers WJ, et al. JAMA 2011;306(18):1983

### An cardiologist's doubt

- Carotid intervention should also improve hemodynamics
- \* But can we demonstrate perfusion failure?
- \* How can we prove the benefit of rectifying perfusion, if any?

### **Objective cerebral ischemia**

baseline

post stenting





stress mean transit time

# Example case



# Progressive "dementia"



# Final angiograms



#### Our experience with CAO

- \* 235 CAO attempts in 1409 carotid procedure since 2002 (16%)
- \* Feasibility and safety has been reported, with overall 73% success and 4.3% 30d stroke/death
  - \* 83% success and 3.3% 30d stroke/death since 2016

JACC 2007;49:765-771 Circ Cardiovasc Intervent 2008;1:119-125 NTUH data on file





#### **Overall clinical outcome**

#### Composite endpoints: stroke/TIA + ICH + Death



Same endpoints, excluding peri-procedural complications



Logrank test: p=0.0326

Hazard ratio: 0.48, p= 0.040 after adjusting age, sex, DM, HTN, hyperlipidemia, smoking, CAD, CKD, prior ipsilateral ischemic event

Logrank test: p=0.0236

Hazard ratio: 0.44, p= 0.024 after adjusting age, sex, DM, HTN, hyperlipidemia, smoking, CAD, CKD, Prior ipsilateral ischemic event

#### **ERCAO trial**

- Prospective randomized control trial to evaluate the benefit of Endovascular Revascularization for CAO in addition to optimal medical therapy
- To include as many patients as possible and to observe real-world outcomes, 2 parallel registry arms are also included
- \* Funded by NTUH A1 Project grant 201702049RINA
- \* ClinicalTrials.gov ID NCT03179774



#### Inclusion criteria

- Classic NASCET carotid symptoms on optimal medical treatment after documentation of CAO, or
- \* Objective ischemia by CTP
- Collaborative research team including neurologist, neuro-psychiatrist, clinical psychiatrist, radiologist, and interventionist

# CT perfusion

- \* Performed off-line at a workstation using commercial software (CT Perfusion 3, Advantage 4.2; GE Healthcare)
- Cerebral blood volume (CBV), cerebral blood flow (CBF), time to peak (TTP), and mean transit time (MTT) will be calculated
- \* By independent radiologist

# Neurocognitive function (NCF)

#### \* Questionnaire and tasking

- \* Mini-Mental State Examination (MMSE)
- Alzheimer Disease Assessment Scale–Cognitive subscale (ADAS-Cog),
- Verbal fluency (category naming: fruits, vegetables, and fishes)
- \* Color trailing test parts 1 and 2
- \* Conducted by independent neuro-psychiatrist

# Battery of CANTAB

Stage	Tests	Estimated Minutes
Practice and attention	Motor Screening Task (MOT)	2
	Reaction Time (RTI)	5
	Rapid Visual Information Processing (RVP)	7
Memory	Paired Associates Learning (PAL)	10
	Delayed Matching to Sample (DMS)	12
	Pattern Recognition Memory (PRM)	5
Decision making and executive function	Spatial Working Memory (SWM)	8
	Spatial Span (SSP)	8
	Intra-Extra Dimensional Set Shift (IED)	7
	One Touch Stockings of Cambridge (OTS)	10

#### **Optimal medical therapy**

- \* Dual antiplatelet therapy
- \* Aggressive treatment of dyslipidemia
- Treatment of hypertension and diabetes according to national guidelines
- \* Smoking cessation
- \* Lifestyle modification, including diet and exercise

#### Endovascular Intervention

- \* A standard endovascular procedure will be performed
- Technical success: if the occlusion segment was recanalized with final residual diameter stenosis of <20%, establishing grade 3 antegrade TICI flow</li>
- \* Procedure success: technical success without 30d stroke/death

#### **Current enrollment status**



#### Conclusions

- Hemodynamic insufficiency may be as important as embolism in carotid artery disease
- Benefit of carotid intervention may not be limited to stroke prevention, but also includes cerebral perfusion and neurocognitive improvement
- \* ERCAO trial may prove the long-term neurological and survival benefit of carotid revascularization in CAO patients

# Thanks for your attention