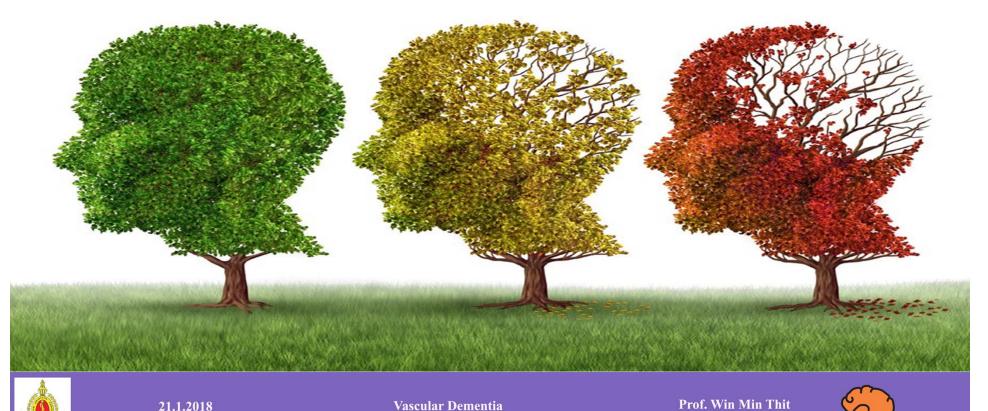
#### **Professor Win Min Thit**

Professor/Head Department of Neurology Yangon General Hospital 21.1.2018 (Sunday)



### Vascular Dementia

# "A forgetful person, in no real distress who can no longer do their job, can no longer be independent and who cannot really sustain any ordinary sensible conversation."



MMA, Yangon

- 1. What is dementia?
- Burden of dementia
- Anatomy and function
- Historical Background
- Aetiology of Vascular dementia
- 6. Criteria for vascular dementia
- 7. Classification of dementia
- Types and proposed mechanisms of vascular dementia
- Clinical features of vascular dementia
- 10. Clinical tools for dementia
- 11. Management of vascular dementia





#### 1. What is Dementia?

#### Dementia ....

- memory impairment
- impairment in other ≥2 cognitive domains
  - Orientation
  - Attention
  - Language
  - Visuo-spatial functions
  - Motor control
  - Praxis
  - Executive functions
- Progressive impairment in functional status





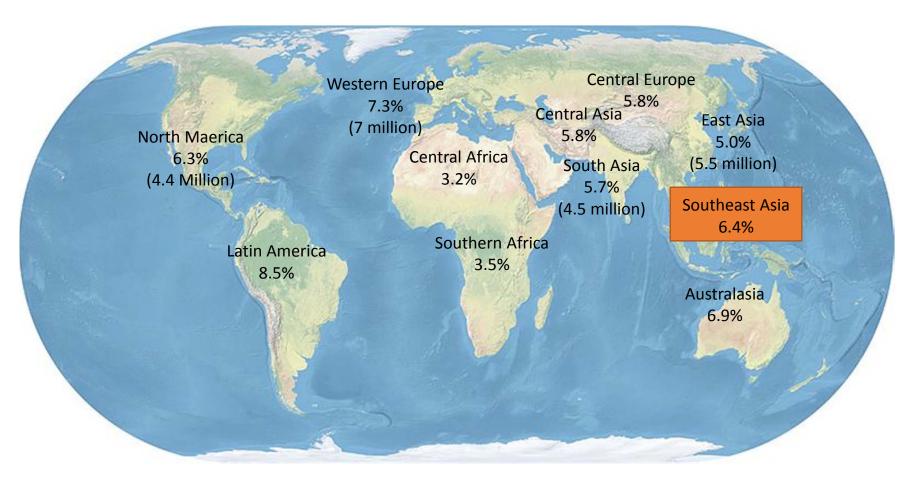
#### 2. Burden of Dementia

- Globally nearly 9.9 million people dementia each year; this figure translates into 1 new case every 3 seconds.
- In 2015, dementia affected 47 million people worldwide, a figure that is predicted to increase to 75 million in 2030 and 132 million by 2050.
- In 2015, dementia costs were estimated at US\$ 818 billion
- By 2030, it is estimated that the cost of caring for dementia people worldwide will have risen to US\$ 2 trillion.





#### Global prevalence of dementia



World health Organization (WHO) Dementia: a public health priority. Geneva WHO, 2012.



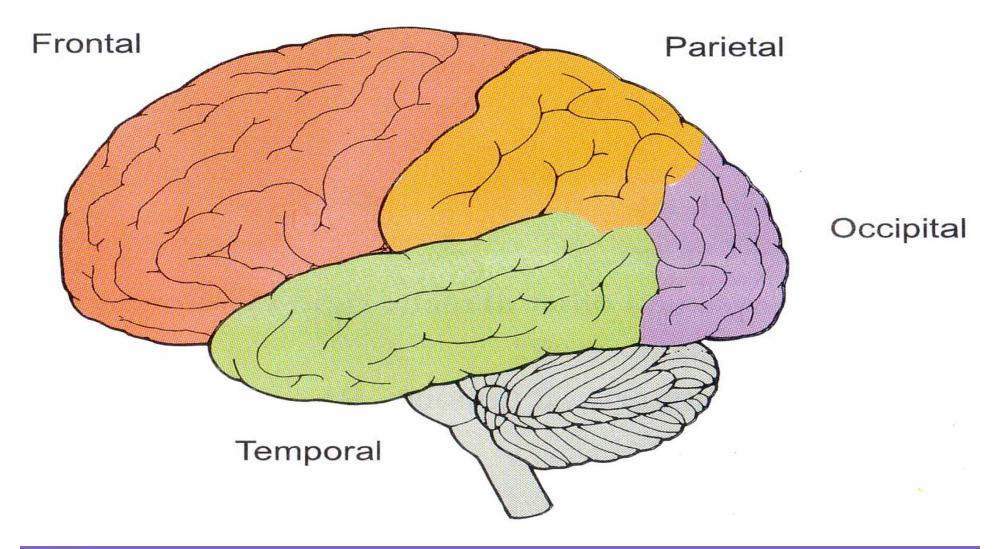


#### 3. Anatomy & Physiology



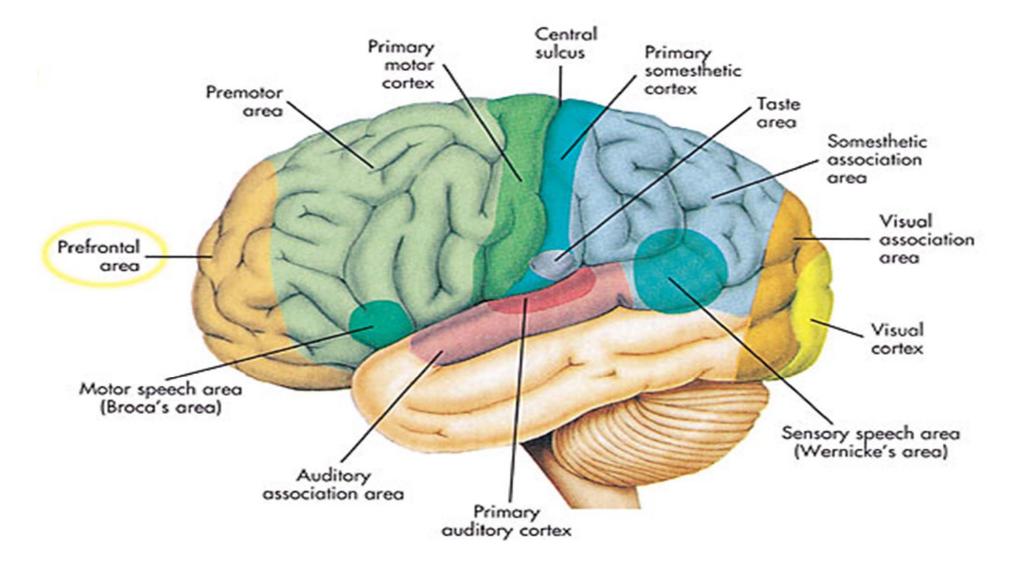


#### Lobes of the brain





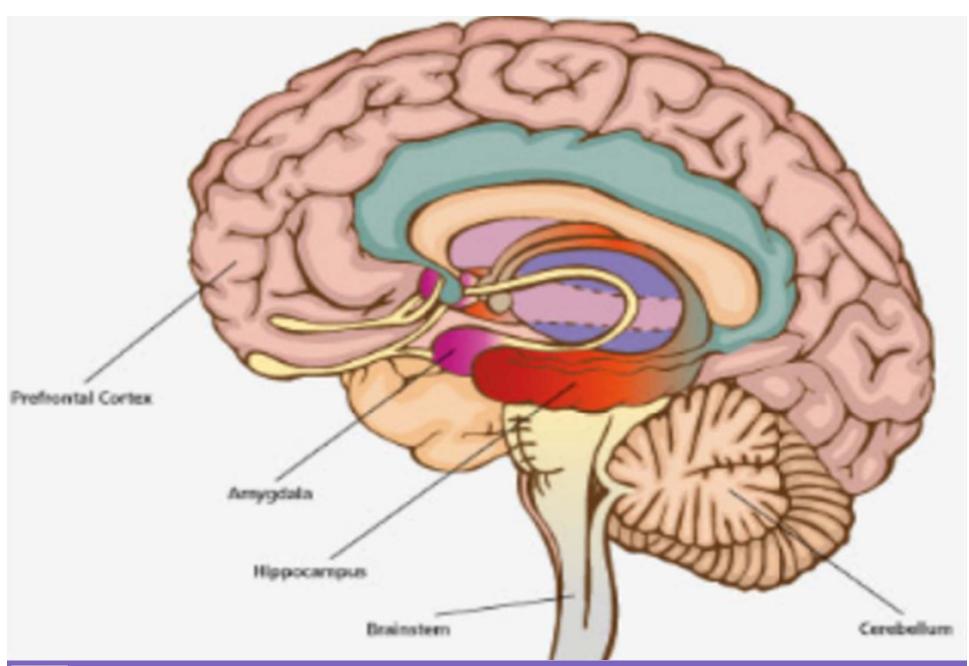




#### **Anatomy and Cortical Lobe Functions**











#### **Cortical lobar functions**

Lobe	Function	Effects of damage
		Cognitive / Bahavioural
Frontal	Personality	Disinhibition
	Emotional control	Lack of initiation
	Social behavior	Anti-social behavior
	Language	Impaired memory
		Expressive dysphasia
	Micturition	Incontinence
	Contralateral motor control	Contralateral Hemiplegia
Parietal:	Language	Dysphasia
dominant	Reading	Dyslexia
	Writing	Dysgraphia
	Calculation	Dyscalculia
		Apraxia, Agnosia

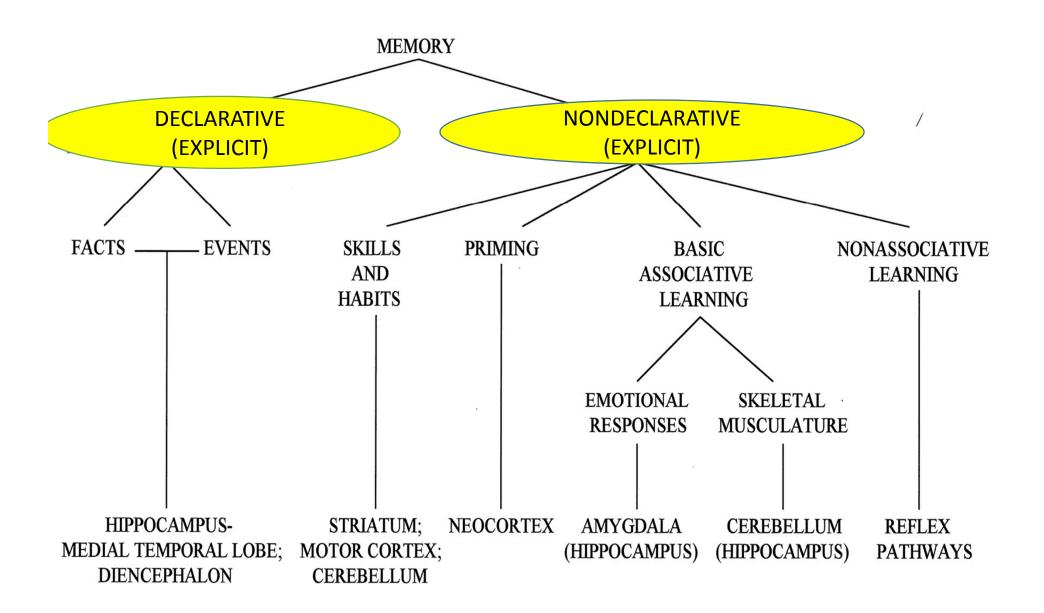




Parietal: non-	Spatial orientation	Spatial disorientation	
dominant		Neglect of non-dominant side	
	<b>Constructional skills</b>	Constructional apraxia	
		Dressing apraxia	
Temporal:	<b>Auditory perception</b>	Receptive aphasia	
dominant	Language	Dyslexia	
	Verbal memory	Impaired memory	
	Smell	Complex hallucinations (smells, sound,	
	Balance	vision, memory)	
Temporal: non-	Auditory perception	Impaired musical skills	
dominant	Melody / pitch perception	(tone perception)	
	Non-verbal memory	Impaired non-verbal memory	
	Smell	Complex hallucinations (smells, sound,	
	Balance	vision, memory)	
Occipital	Visual processing	Visual inattention	
		Visual loss	
		Visual agnosia	











#### 4. Historical Background

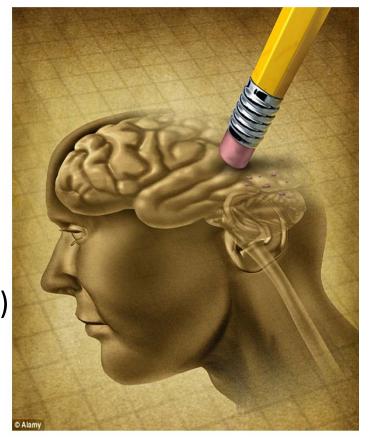
- In 1672, Thomas Willis first reported cases of vascular dementia post apoplexy (later known as post stroke dementia). He separated mental retardation from dementia.
- In 1894, Otto Binswanger and Alois Alzheimer separated vascular dementia from dementia paralytica caused by neurosyphilis.
   They described dementia postapoplexy, arteriosclerotic brain degeneration, vascular cortical atrophy and Binswanger's disease.
- Hachinski in 1974 used the term "multi-infarct dementia".
- In 1995, the name "vascular cognitive impairment" was proposed by Bowler & Hachinski to emphasize the need for prevention and early diagnosis.





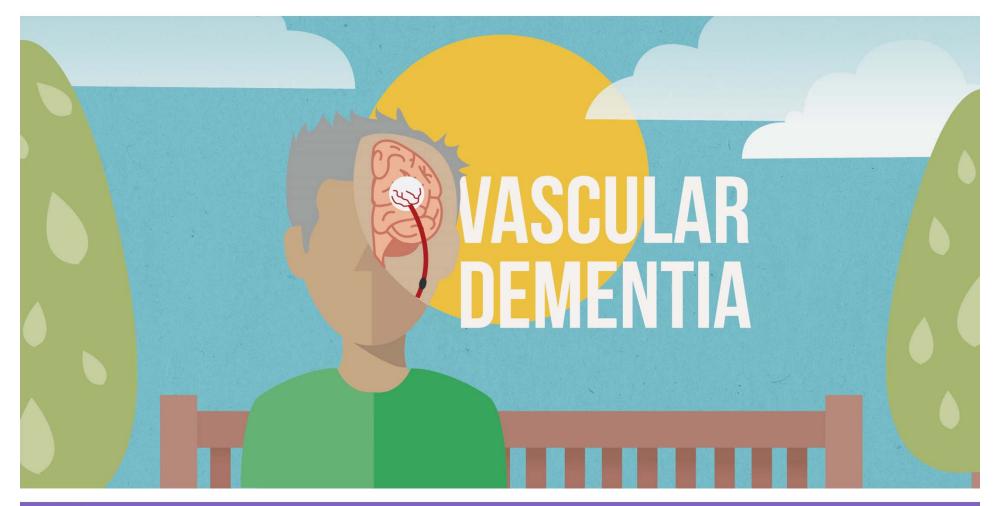
#### 5. Aetiology of VaD

- Stroke (but no clear link with location)
- Hypertension (in mid-life)
- Hypotension (in late-life)
- Hyperlipidaemia (in some studies)
- Diabetes (& 'metabolic syndrome')
- Smoking (and probably other risk factors)
- Genetic causes (rare)
- But age is strongest 'risk factor'





#### 6. Criteria for Vascular Dementia







#### NINDS-AIREN Criteria (Roman et al, 1993)

#### A) Dementia

- B) Cerebrovascular disease (confirmation by brain imaging -must)
  - ✓ Infarcts-single strategic, multiple cotical, multiple lacunar
  - ✓ Small vessel (Leukoaraiosis/Binswanger)
  - ✓ICH, Hypoperfusion
  - √ Genetic CADASIL
  - √ Vasculitis

#### C) A and B must be reasonably related

Based on Román GC et al: Vascular dementia: Diagnostic criteria for research studies. Report of the NINDS-AIREN International Workshop held at the National Institutes of Health, Bethesda, Md, April 19-21, 1991. *Neurology* 1993;43:250-260.





#### DSM-IV Criteria for VaD

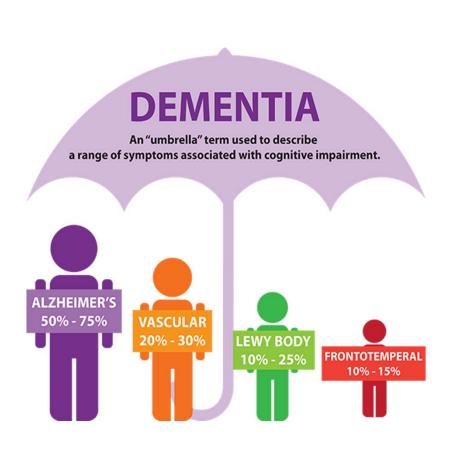
- 1. Multiple Cognitive Deficits including amnesia
- 2. Significant impairment in social or occupational functioning which is a change
- 3. Presence of focal neurological signs and symptoms or laboratory evidence (=neuroimaging) of cerebrovascular disease judged to be aetiologically related to dementia (stepwise decline dropped)
- 4. Deficits not only during a delirium

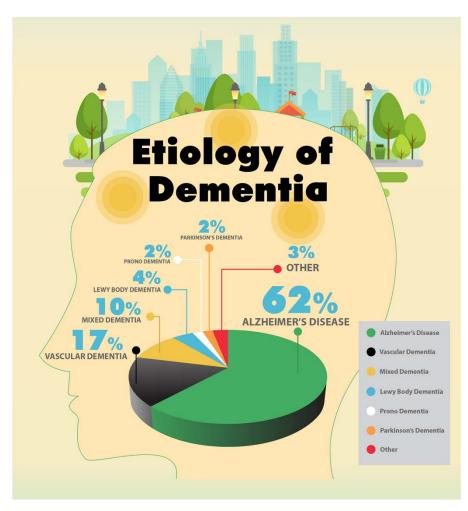
Diagnosis and Statistical Manual of Mental Disorders, Fourth Edition, American Psychiatric Association, 2000.





#### 7. Classification of dementia









#### Classification of dementia (depends on Anatomy)

	Cortical dementia	Subcortical dementia	
Neuropathology	Cortical association areas	Striatum, Thalamus	
Severity	More severe	Mild to moderate	
Speed of cognition	normal but frequent errors	slow	
Neuropsychology	More severe memory impairment unaided by cues, Dysphasia, dyspraxia, agnosia	memory impairment, Recall aided by cues	
Mood	Depression less common	Apathy, Depression	
Motor abnormalities	common, Gegenhalten	Extrapyramidal, dysarthria	





## Classification of dementia (depends on Pathophysiology)



Alzheimer's dementia

(60%)



Vascular dementia

(20%)



**Others** 

- Lewy Body dementia -Frontal lobe dementia -Parkinson disease dementia -Corticobasal degeneration -Normal pressure hydrocephalus

Mixed dementia (combination of AD and vascular dementia)



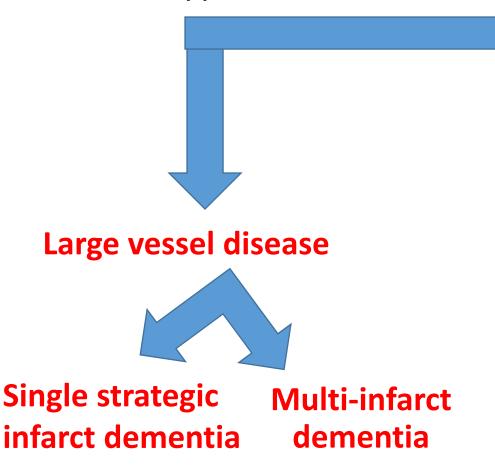


### 8. Types & Proposed mechanism of vascular dementia





#### Types of Ischaemic Vascular dementia



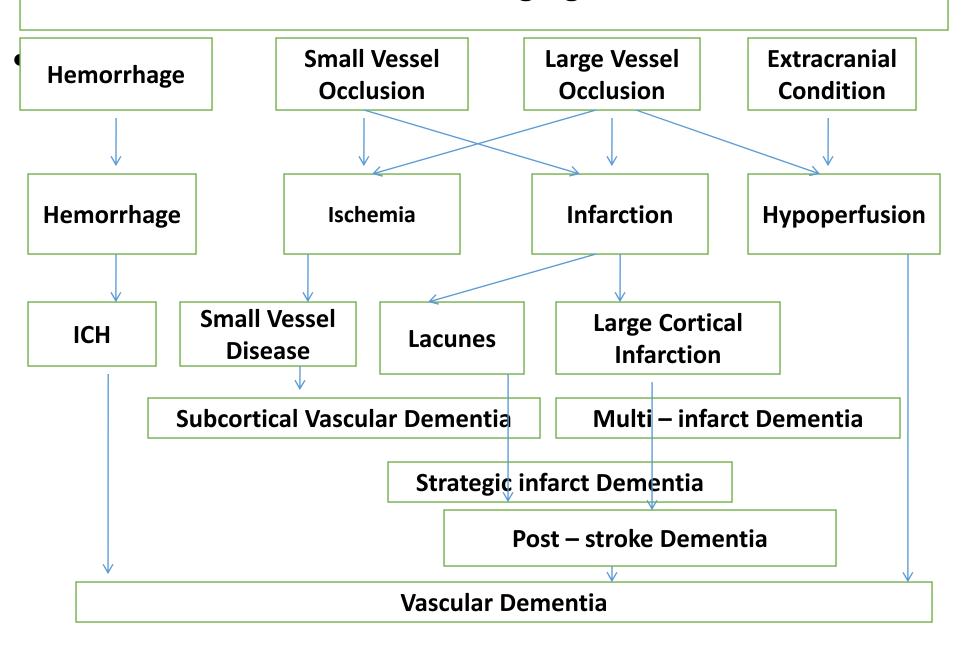
Small vessel disease (Subcortical vascular dementia)

- 1. Multiple small deep infarcts (Lacunes)
- Diffuse ischaemic lesions of deep white matter (Binswanger / Leukoaraiosis)





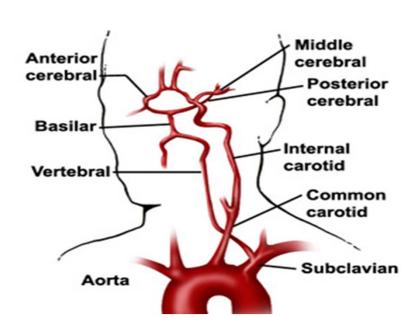
#### Vascular Risk Factors, Genetic, Aging, Environmental Factors

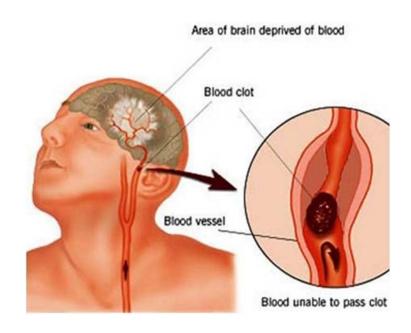


#### Dementia due to large vessel disease

Vascular Dementia

- 1. Single strategic infarct dementia
- 2. Multi-infarct dementia









#### 1. Single strategic infarct

**Anterior Cerebral Artery (ACA) (Inferomedial frontal infarct)** 

- Abulia, memory impairment, language impairment

Lt Middle Cerebral Artery (MCA) infarct (Dominant parieto-temporal, temporo-occipital association, angular gyrus)

- Aphasia with cognitive impairment

Rt MCA (Non- Dominant pareito-temporal, temporo-occipital association)

- visuo-spatial functions

Posterior cerebral artery (PCA) (Bilateral inferomedial temporal or thalamic infarct)

- Amnesia

Watershed infarcts (Superior frontal or parietal)
Lacunar infarcts (Bilateral thalamic)

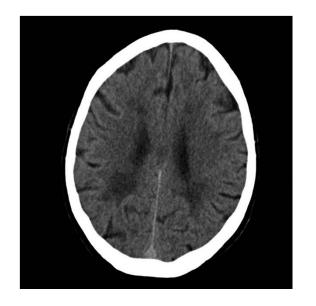
- Amnesia

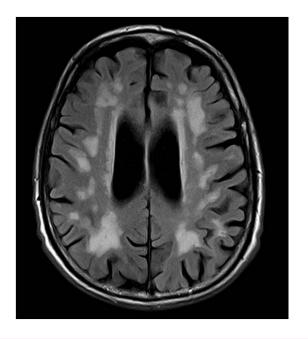




#### 2. Multi-infarct dementia (MID)

- Multiple large infarct
- Destroying critical brain volume
- Not necessarily at strategic sites
- Dementia occurs 'stroke by stroke', with progressive focal loss of function
- Diagnosis is obtained from the history and confirmed by CT or MRI scan (presence of multiple areas of infarction)





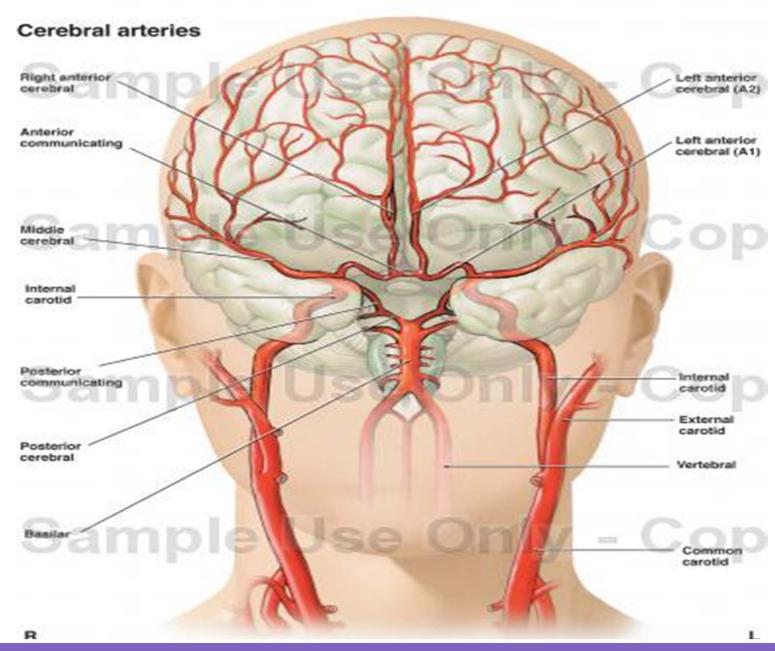




#### Dementia due to small vessel disease











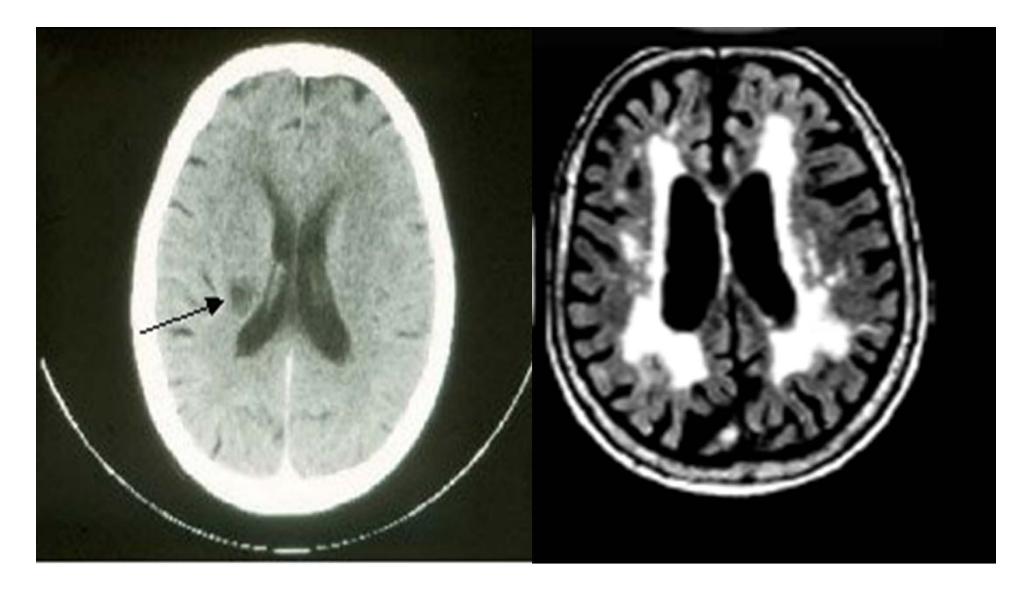
#### 3. Small vessel disease

- 1. Multiple small deep infarcts (Lacunes)
- Widespread patchy or diffuse ischaemic lesions of deep cerebral white matter (Binswanger disease)
  - ✓ Symptoms develop more gradually
  - ✓ Abulia
  - ✓ Abnormal behaviour with up and down emotion
  - ✓ Pseudobulbar palsy
  - ✓ Pyramidal signs
  - ✓ Disturbed gait
  - ✓ Loss of bladder control and Urinary incontinence
  - ✓ CT/MRI periventricular or subcortical lucencies (Leukoaraiosis)

SVD can progress silently for many years before becoming clinically evident







Lacunes (CT)

Leukoaraiosis (MRI)





#### **CADASIL**

- Cerebral Autosomal Dominant Arteriopathy with Subcortical Infarcts and Leucoencephalopathy.
- mutations in NOTCH3 gene (which had a role in arterial development and is expressed on vascular smooth-muscle cells)
- Clinical phenotype:
  - Migraine
  - Recurrent strokes & TIAs
  - Dementia
  - Psychiatric disturbance
  - Onset usually in the third to sixth decade
  - About a 1/3 of patients develop migraine with aura-early sign







#### 9. Clinical Features







#### Features Associated with Dementia



- Agitation
- Aggression
- Sleep disturbances
- Apathy (can be misdiagnosed as depression)
- Depression or anxiety
- Personality changes

- Behavioral disinhibition
- Impaired insight
- Hallucinations (visual > auditory)
- Delusions (often paranoid or persecutory)





#### Clinical features consistent with Vascular Dementia:

- Early gait disturbances, frequent falls
- Parkinsonian features
- Early urinary symptoms
- Personality change, mood disorders (vascular depression), psychomotor retardation
- Predominant abnormal executive function (Affects subcortical & frontal lobes)
- Memory and language deficits less obvious & occur late
- higher risk for institutionalization than AD due to more Behavioural &Psychiatric symptoms





DO I HAVE VASCULAR DEMENTIA?

#### 10. Clinical tools for Dementia







#### MINI MENTAL STATE EXAMINATION (MMSE)

#### MINI MENTAL STATE EXAM

Please name the:

Year? Season? Date? Day of Week? Month?

Orientation to time /5

Where are we?

State? City? Suburb? Hospital? Floor/Ward?

Orientation to place /5

"I am now going to test your memory" Name 3 objects. Ask them to repeat all 3. 1 Point for each object remembered. Repeat

until learnt all 3 so that recall can be tested.

Registration /3 # of trials

Serial 7s

'please count backwards from 100 in sevens" 93, 86, 79, 72, 65

alternatively Spell WORLD backwards DLROW

Attention and Calculation /5

"Please repeat the 3 objects I asked you to remember"

Recall /3

"Please name these objects" Point to a wristwatch and a pencil

Naming /2

"Please repeat the following phrase"
"No ifs, ands or buts"

Repetition /1

"Please follow this command" "Take this paper in your right hand, fold it in half and place it in your lap"

Complex command /3

Please read and obey the following command

#### CLOSE YOUR EYES

"Please write a sentence" Must have a noun, verb and make sense

"Please copy the following drawing"



1 point each for the last 3 commands /3

24-30-normal range

18-23-moderate cognitive impairment

TOTAL /30

0-17 -marked

congnitive impairment





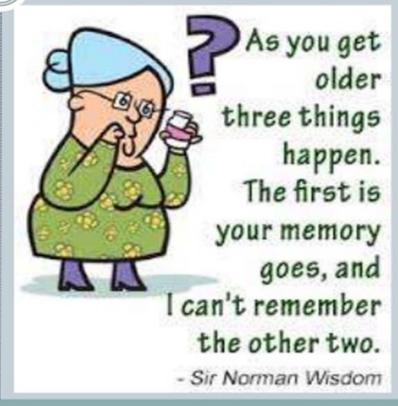
MONTREAL C	OGNITIVE ASSESS	SMENT (MOCA	Educa	AME: tion: Sex:	Date of birth : DATE :	
S End Begin	(B) (2) (4) (3)		Copy	Draw CLOCK (3 points)	(Ten past eleven)	POINTS
©	1.1		1.1	[ ] Contour Nu	l [] mbers Hands	_/s
NAMING		A B		Y		/3
MEMORY	Read list of words, subject must repeat them. Do 2 tri: Do a recall after 5 minutes.	als.	ACE VELVET	CHURCH	DAISY RED	No points
ATTENTION	Read list of digits (s digit/ s		repeat them in the repeat them in the		[]21854	_/2
Read list of letters. Th	e subject must tap with his l				AAJAMOFAAB	_/1
Serial 7 subtraction s	tarting at 100 [ ]		[] 79 ractions: 3 pts, 2 or 3	[ ] 72 correct: 2 pts, 1 cor	[ ] 65 rect: 1 pt, 0 correct: 0 pt	_/3
LANGUAGE	Repeat: I only know that Jo The cat always his	ohn is the one to help I under the couch who		100m. [ ]		_/2
Fluency / Name	maximum number of words	in one minute that be	gin with the letter	[ ]_	(N≥ 11 words)	_/1
ABSTRACTION	Similarity between e.g. ban	ana - orango = fruit	] train - bicycl	e [ ] watch -	ruler	_/2
Optional	WITH NO CUE  Category cue	FACE VELVET		AISY RED	Points for UNCUED recall only	_/5
	Multiple choice cue					₩.
ORIENTATION		lonth [ ] Year	[ ]Day	[ ]Place	[ ]City	_/6
© Z.Nosreddine MD V WWW.mocatest	ersion November 7, 2004		Normal	26/30 TOTA	AL Add 1 point if 5 12 yr ei	/30





#### 11. Management of Vascular Dementia

## TREATMENT OF DEMENTIA







#### Primary Prevention of VaD

#### **Target**

Brain at risk of CVD

#### **Action (treatment of risk factors)**

- Life style modification
- Arterial hypertension
- Cardiac abnormality
- Lipid abnormality: DIET, statins
- Diabetes mellitus
- Homocysteine







#### Secondary Prevention of VaD

#### **Target**

CVD brain at risk of VCI/VaD

#### **Action**

- Treatment of acute stroke (rtPA)
- Prevention of stroke recurrence
- Slow progression of VaD related changes
- Treatment of vascular risk factors
- Neuroprotection ? Cerebrolysin, ?Citicoline
   VCI = vascular congnitive impairment



#### Once Vascular Dementia is present,

- Acetyl cholinesterase inhibitors (AChEI) may have mild moderate benefit
- Memantine may be useful adjunctive to AChEI in moderate severe dementia
- Anti depressants (specifically SSRIs)
- Atypical antipsychotics
- No adequately controlled trials demonstrating pharmacologic efficacy for any agent in ischemic vascular (multi-infarct) dementia.





#### 12. Take Home Messages

- Association between stroke and dementia is frequent and post stroke dementia adversely influences the outcome in stroke patients.
- Vascular dementia can be caused by both large and small vessel diseases.
- Small vessel disease can progress silently for many years before becoming clinically evident.
- Look for vascular risk factors and focal neurological signs
- Post stroke dementia is very important because appropriate control of vascular risk factors could lead to prevention
- Primary prevention depend on early identification and control of vascular risk factors
- Secondary prevention must include energetic therapy to prevent stroke recurrence.





## SEE THE PERSON NOT THE DEMENTIA

Q & A & Discussion



#### Thank You



