

Cerebral blood flow: possible hemodynamic links between atrial fibrillation and cognitive decline

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Background & State of the Art

● Atrial fibrillation (AF)

- Common arrhythmia: irregular and faster beat;
- 33.5 million people worldwide in 2010 (to be doubled in 40 years);
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- Neurological degeneration: loss of memory, socio-cognitive alterations;
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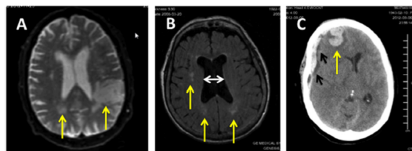
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- 81 million people worldwide in 2040;
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- Recent independent **association** between **AF** and **dementia**
 ⇒ Potential **hemodynamic mechanisms**: microembolisms, altered cerebral blood flow, hypoperfusion and microbleeds.

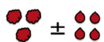
Background & State of the Art



Spectrum of macro- to micro-cerebral injuries from AF (*Jacobs et al., 2015*)



Macro Emboli

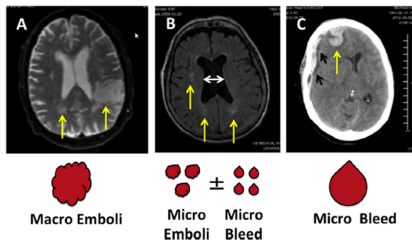


Micro Emboli ± Micro Bleed



Micro Bleed

Background & State of the Art

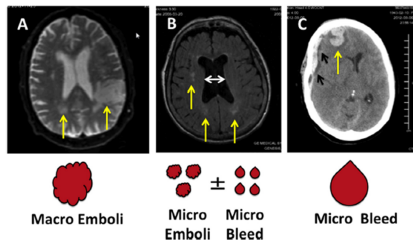


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● Open Questions

- Linking mechanisms between AF and dementia;
- Consequences of AF rate/rhythm control on cognitive decline;
- Lacking clinical measures in the cerebral microcirculation.

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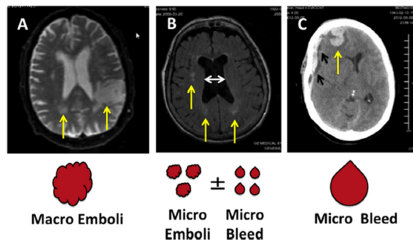
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● Modeling approach for the cerebral hemodynamics during AF

Anselmino et al., *Sci. Rep.*, 2016; Scarsoglio et al., *J. R. Soc. Interface*, 2017; Scarsoglio et al., *Chaos*, 2017

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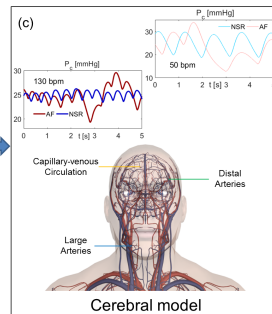
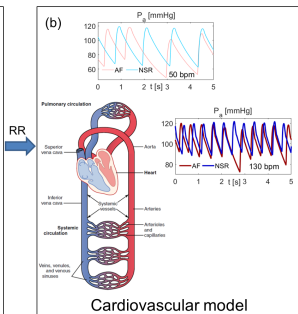
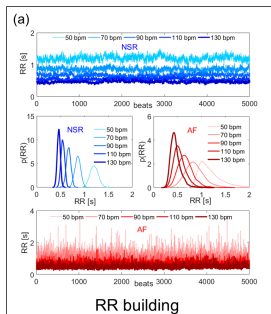
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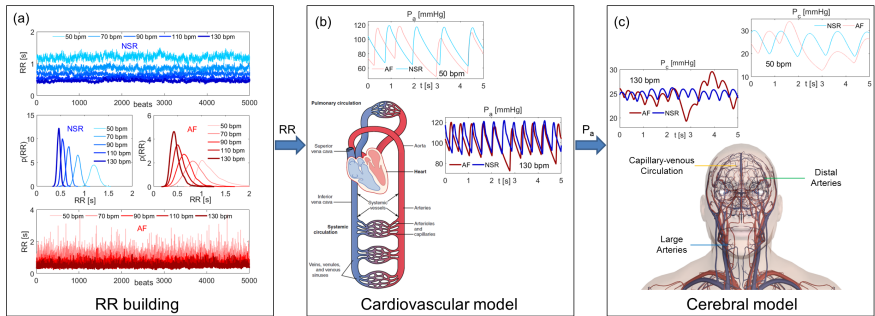
⇒ Impact of heart rate (HR) during AF on cerebral hemodynamics

Methods: Computational algorithm



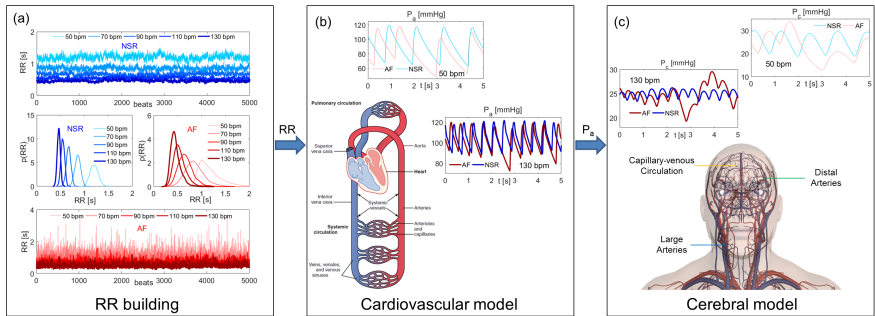
- Stochastic RR beating extraction & validated lumped modeling;

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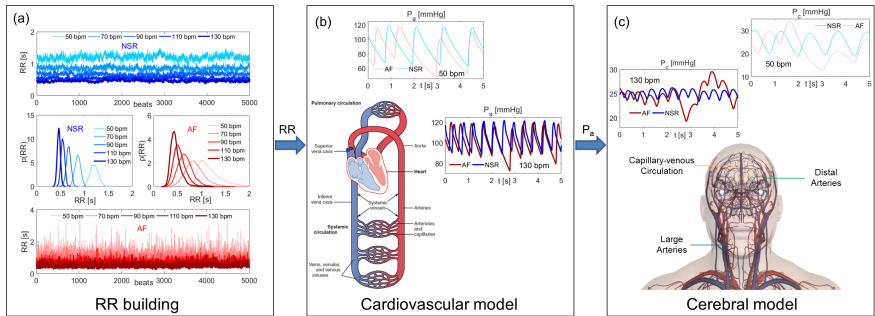
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- NSR (normal sinus rhythm, blue) and AF (red);

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- Stochastic RR beating extraction & validated lumped modeling;
- Mean HR = 50, 70, 90, 110, 130 bpm;
- NSR (normal sinus rhythm, blue) and AF (red);
- 5000 cardiac cycles (RR beats) simulated for each configuration.

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 - Governing equations for each region:
 - ⇒ Continuity equation
 - ⇒ Momentum equation
 - ⇒ Constitutive relation between P and V

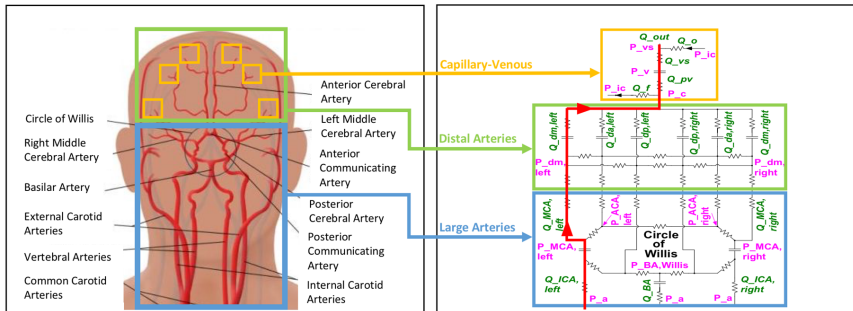
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 - Baroreceptor, autoregulation, and CO_2 reactivity mechanisms;
- Focus on the **proximal-to-distal pathway** (left side):
 - Large arteries ($P_a, P_{MCA,left}, Q_{ICA,left}, Q_{MCA,left}$);
 - Distal arteries ($P_{dm,left}, Q_{dm,left}$);
 - Capillary-venous circulation (P_c, Q_{pv}).

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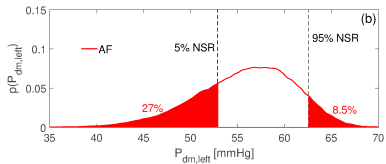
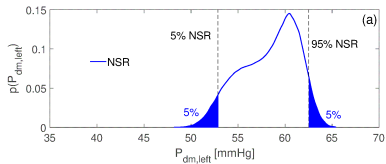


- Focus on the proximal-to-distal pathway (left side):

- Large arteries (P_a , $P_{MCA, left}$, $Q_{ICA, left}$, $Q_{MCA, left}$);
- Distal arteries ($P_{dm, left}$, $Q_{dm, left}$);
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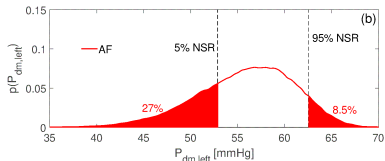
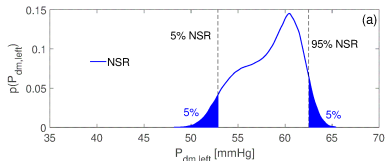
Methods: Data analysis

● Percentile evaluation

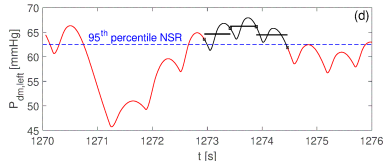
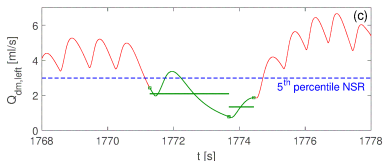


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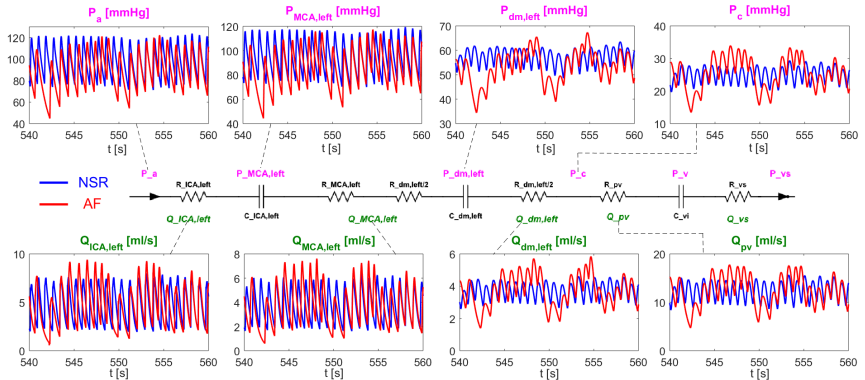
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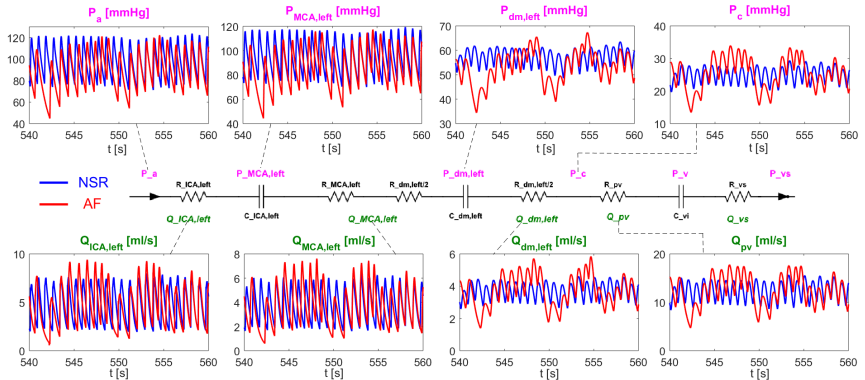
● Recurrence of extreme events (hypoperfusions and hypertensive events) in AF



Results: proximal-to-distal pathway

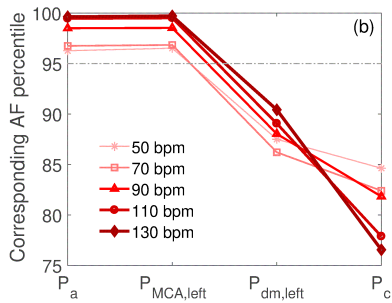
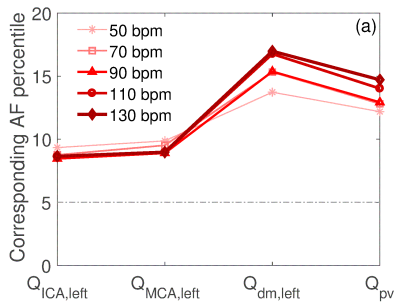


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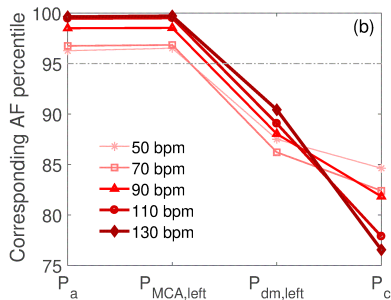
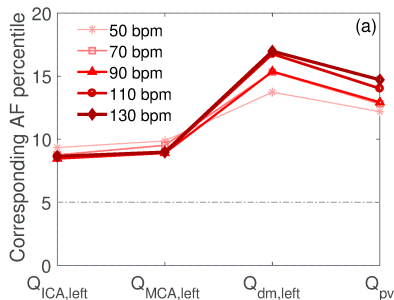


- HR=70 bpm. Compared to NSR, AF triggers a **higher variability** of the cerebral hemodynamic variables, increasingly proceeding towards the distal circulation.

Results: percentile analysis

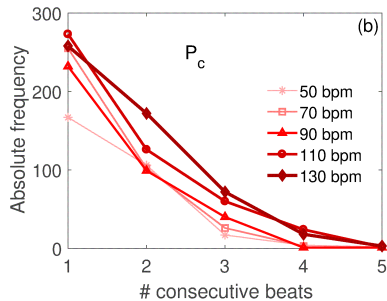
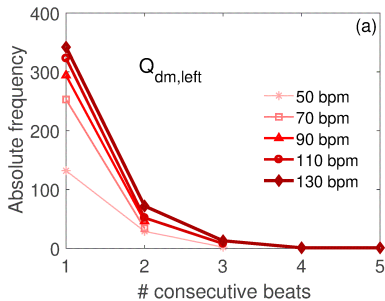


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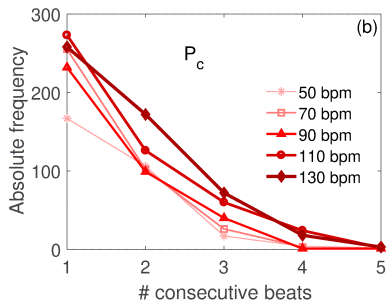
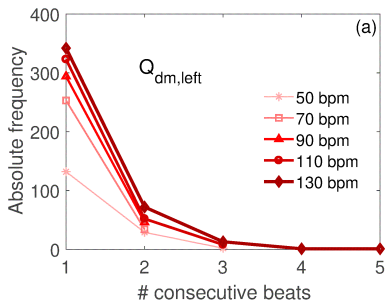


- The increased variability during AF leads to critical hemodynamic events of **reduced blood flow or excessive pressure** in the deepest cerebral circulation (arterioles and capillaries).

Results: hypoperfusions and hypertensive events



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- Absolute frequency over 5000 beats of (a) hypoperfusions ($Q_{dm,left}$) and (b) hypertensive events (P_c) during AF.

Results: total number of one-beat extreme events

Hypoperfusions				
	$Q_{ICA,left}$	$Q_{MCA,left}$	$Q_{dm,left}$	Q_{pv}
50 bpm	1	2	196	124
70 bpm	0	0	321	136
90 bpm	0	0	386	216
110 bpm	0	0	451	352
130 bpm	0	0	534	415
Hypertensive events				
	P_a	$P_{MCA,left}$	$P_{dm,left}$	P_c
50 bpm	0	0	231	456
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- Critical events (over 5000 beats) mainly occur in the distal region (rare episodes in the proximal region) and increase with HR.

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Compagnia di San Paolo is acknowledged for funding the present work within the Project CSTO160444 "*Cerebral hemodynamics during atrial fibrillation*".