

# *μsesiones*

*Anticoagulación precoz en el ictus  
en pacientes con FA no valvular*

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17.01.2025

*If untreated, the risk of early recurrence of ischemic stroke in patients with AF is between 0.5% and 1.3% per day.*

# Guía ESO 2019

## Recomendaciones

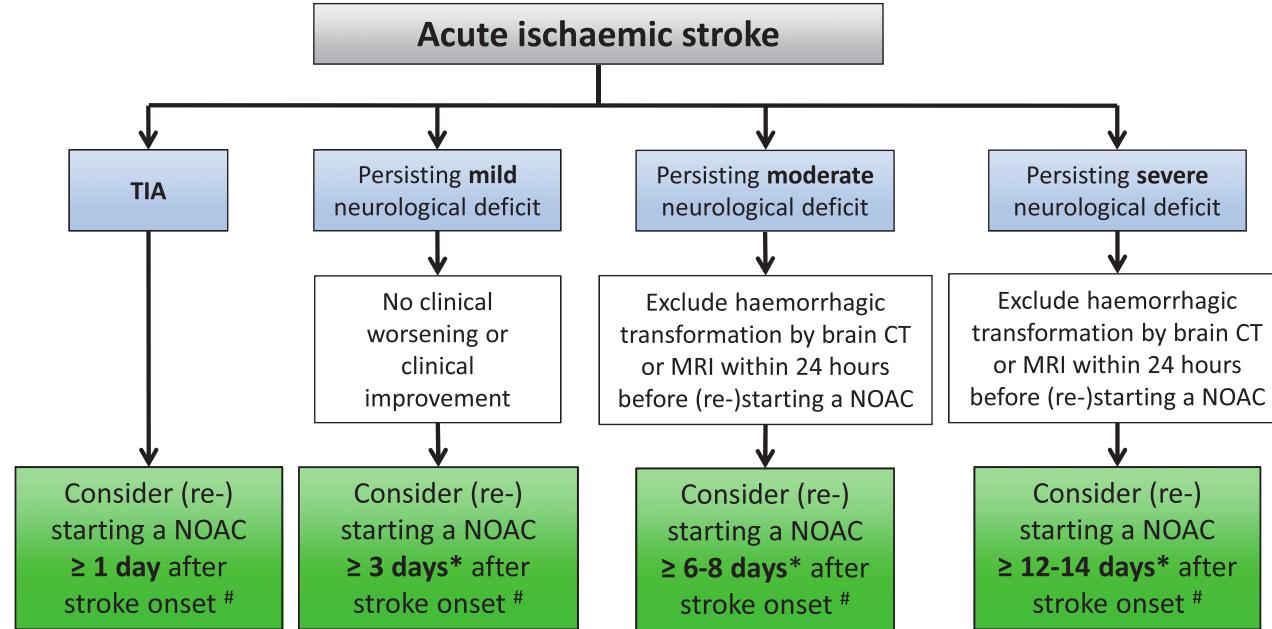
- Antiagregación durante las primeras 48 horas
- Iniciar anticoagulación **DOAC** (o AVK; evitando heparina)

| NIHSS <8 <sup>1</sup> | NIHSS 8-16 | NIHSS >16 |
|-----------------------|------------|-----------|
| 3-4 días              | 7 días     | 14 días   |

1. Volumen de infarto <1.5 cm<sup>3</sup>

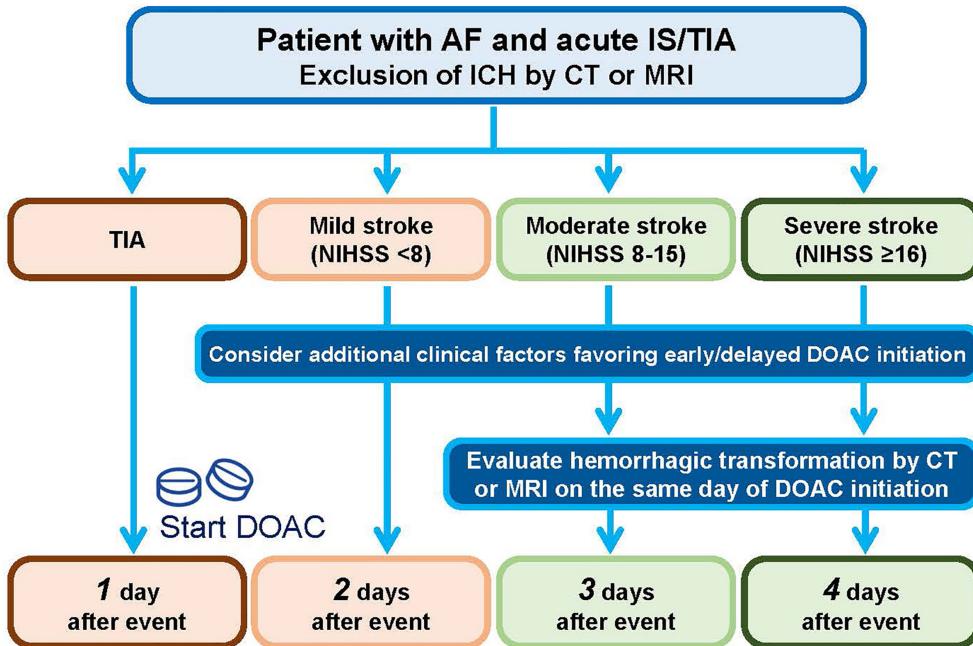
|   |          | Quality of evidence | Strength of recommendation |
|---|----------|---------------------|----------------------------|
| Medical treatment   |          |                     |                            |
| We do not recommend antiplatelet agents (single or dual), over no antiplatelet therapy.   | Moderate | Weak                |                            |
| We recommend VKA therapy over no antithrombotic medication  | Moderate | Strong              |                            |
| We recommend VKAs (INR 2-3) over antiplatelet therapy.  | Moderate | Strong              |                            |
| We recommend NOACs over VKAs  | High     | Strong              |                            |
| We suggest NOACs over aspirin in patients who have failed or are unsuitable for VKA therapy.  | Moderate | Weak                |                            |
| Timing and bridging of medical treatment  |          |                     |                            |
| We cannot make recommendations about the optimal time for initiating anticoagulation treatment in patients with acute ischemic stroke.  | Low      | Weak                |                            |
| Expert opinion: We suggest antiplatelet therapy in the first 48 h after ischemic stroke associated with AF. We consider it reasonable to start anticoagulant therapy at day 3 or 4 from the index stroke in patients with mild stroke and small infarcts (<1.5 cm) and at day 7 for moderate infarcts. For large infarcts, OACs might be best delayed for 14 days after the index stroke. |          |                     |                            |
| We suggest that bridging therapy should be avoided prior to anticoagulation with VKAs or NOACs.   | Low      | Weak                |                            |

# Guía ESC 2018 (1-3-6-12)



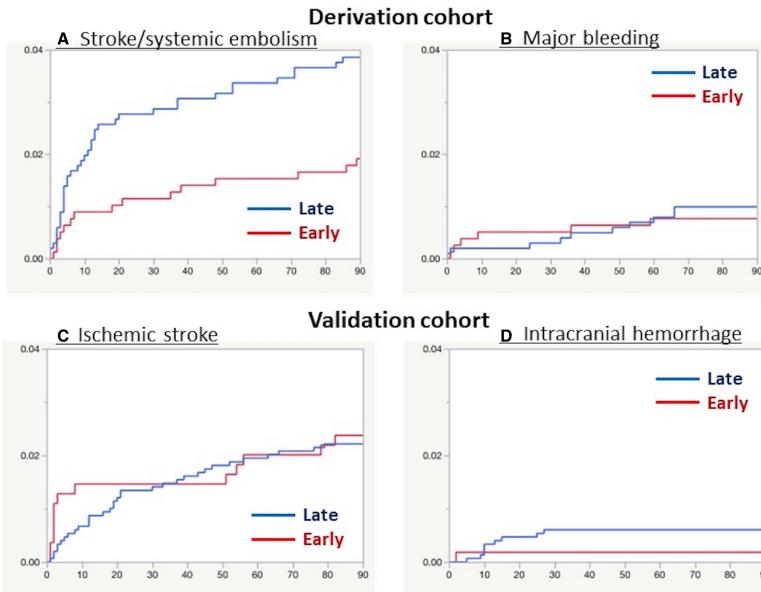
**Figure 14** (Re-) initiation of anticoagulation after transient ischaemic attack/stroke. (Re-) start only in the absence of contraindications and if stroke size is not expected to substantially increase the risk of secondary haemorrhagic transformation. \*Consider shorter delays to (re-) start a non-vitamin K antagonist oral anticoagulant if there is a very high risk of stroke recurrence (e.g. left atrial appendage thrombus) and no haemorrhagic transformation on follow-up brain imaging (using computed tomography or magnetic resonance imaging). Consider longer delays to (re-)start a non-vitamin K antagonist oral anticoagulant according to the recommendations made in the European Society of Cardiology Atrial Fibrillation Guidelines 2016. <sup>#</sup>Without proven evidence; consider inclusion of patient in an ongoing trial.

# 1-2-3-4 😬



Development and validation of the practical “1-2-3-4-day” rule for starting direct oral anticoagulants after ischemic stroke/TIA with atrial fibrillation

- Japón (2 observacionales) + validación con registro Europeo
- **Mediana** de inicio de DOAC para los cuatro grupos de ictus; definen *inicio precoz* como **un día menos**:
  - Japón: 2-3-4-5
  - *Inicio precoz*: 1-2-3-4
  - Europa: 2-5-8-8



¡Ojo!

- Estudio no aleatorizado
- Más fibrinolisis / trombectomía en el grupo precoz
  - neuroimagen en 24 horas del ictus
- Ictus graves excluidos
- Mayor efecto del **rivaroxaban** en japoneses

## Todo el mundo pasa de las guías

| AIT     | NIH <8   | NIH 8-16 | NIH >16  |
|---------|----------|----------|----------|
| 2 días  | 3-4 días | 7 días   | 14 días  |
| 2 (1-3) | 5 (2-10) | 8 (4-14) | 8 (4-14) |

*Mediana y rango intercuartílico*

### (i) Menor riesgo de ictus... en Japón

- HR 0,5 [95% CI 0,27 - 0,89]
- mismo riesgo de hemorragias graves

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## Early versus Later Anticoagulation for Stroke with Atrial Fibrillation

U. Fischer, M. Koga, D. Strbian, M. Branca, S. Abend, S. Trelle, M. Paciaroni, G. Thomalla, P. Michel,

| Tipo de ictus | Menor     | Moderado  | Mayor      |
|---------------|-----------|-----------|------------|
| Precoz        | <48 horas | <48 horas | 6-7 días   |
| Tardío        | 3-4 días  | 6-7 días  | 12-14 días |

¿Un ictus moderado?

## Menor

- $\leq 1,5$  cm

## Moderado

- Una rama de ACM o ACA

## Mayor

- ACM, ACA o ACP completa
- $\geq 2$  ramas de ACM
- $\geq 2$  territorios
- $\geq 1,5$  cm en cerebelo o tronco

# Early vs Late Anticoagulation in Minor, Moderate, and Major Ischemic Stroke With Atrial Fibrillation

## Post Hoc Analysis of the ELAN Randomized Clinical Trial

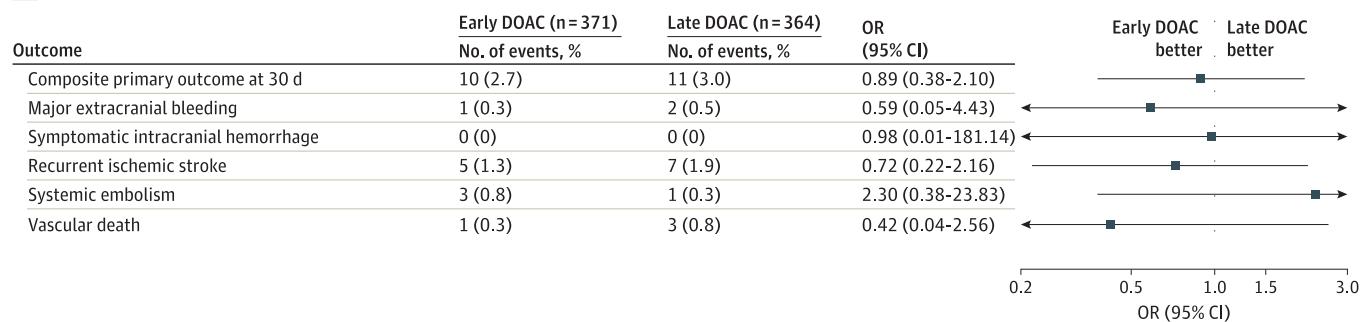
**RESULTS** A total of 1962 of the original 2013 participants (909 [46.3%] female; median [IQR] age, 77 [70-84] years) were included. The primary outcome occurred in 10 of 371 participants (2.7%) with early DOAC initiation vs 11 of 364 (3.0%) with late DOAC initiation among those with minor stroke (odds ratio [OR], 0.89; 95% CI, 0.38-2.10); in 11 of 388 (2.8%) with early DOAC initiation vs 14 of 392 (3.6%) with late DOAC initiation among those with moderate stroke (OR, 0.80; 95% CI, 0.35-1.74); and in 8 of 219 (3.7%) with early DOAC initiation vs 16 of 228 (7.0%) with late DOAC initiation among those with major stroke (OR, 0.52; 95% CI, 0.21-1.18). The 95% CI for the estimated risk difference of the primary outcome in early anticoagulation was -2.78% to 2.12% for minor stroke, -3.23% to 1.76% for moderate stroke, and -7.49% to 0.81% for major stroke. There was no significant treatment interaction for the primary outcome. For infarct size, interrater reliability was moderate ( $\kappa = 0.675$ ; 95% CI, 0.647-0.702) for local vs core laboratory raters and strong ( $\kappa = 0.875$ ; 95% CI, 0.855-0.894) between core laboratory raters.

**CONCLUSIONS AND RELEVANCE** The treatment effect of early DOAC initiation did not differ in people with minor, moderate, or major stroke assessed by brain imaging. Early treatment was not associated with a higher rate of adverse events, especially symptomatic intracranial hemorrhage, for any infarct size, including major stroke.

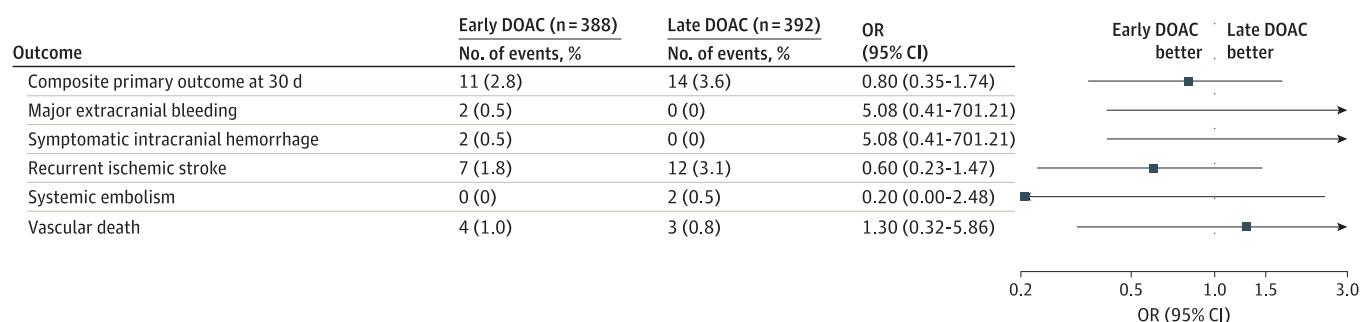
**TRIAL REGISTRATION** ClinicalTrials.gov Identifier: NCT03148457

**Figure 2. Unadjusted Odds Ratios (ORs) of the Primary Composite Outcome and Secondary Outcomes at 30 Days**

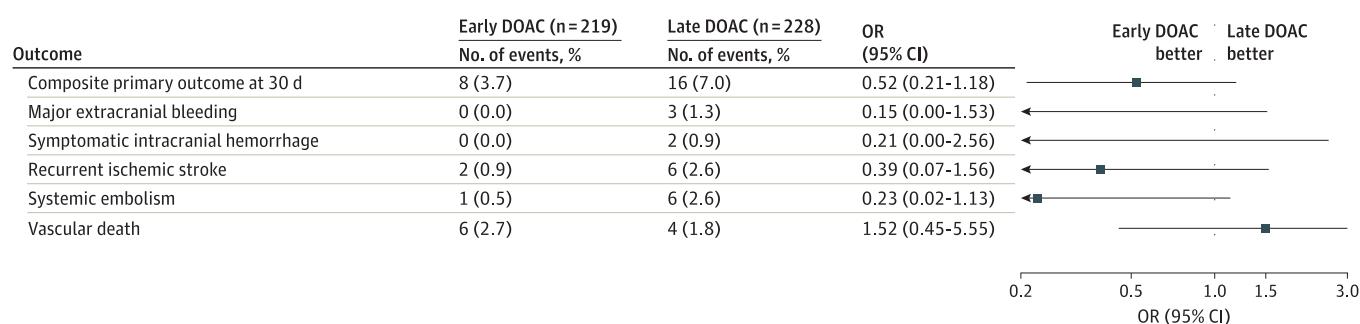
**A** Minor stroke



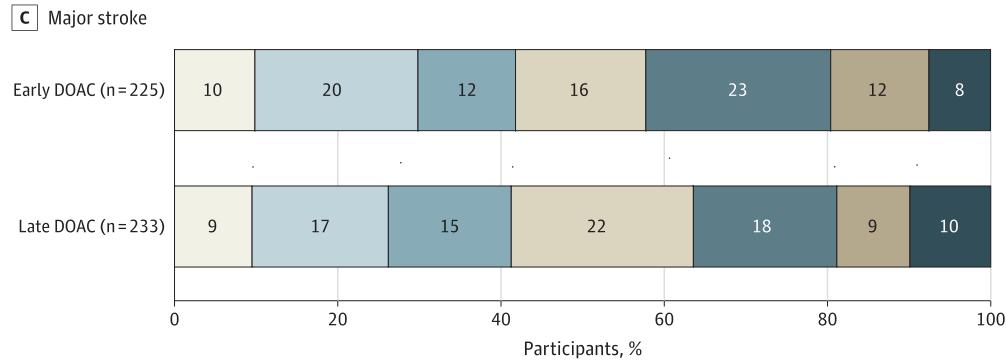
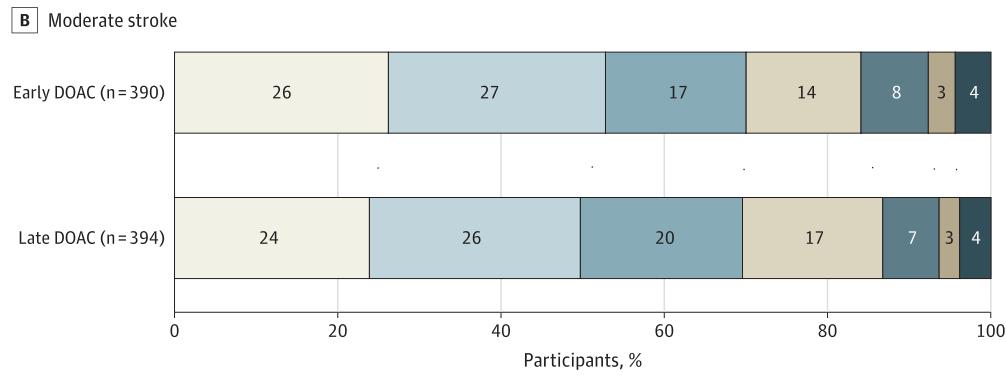
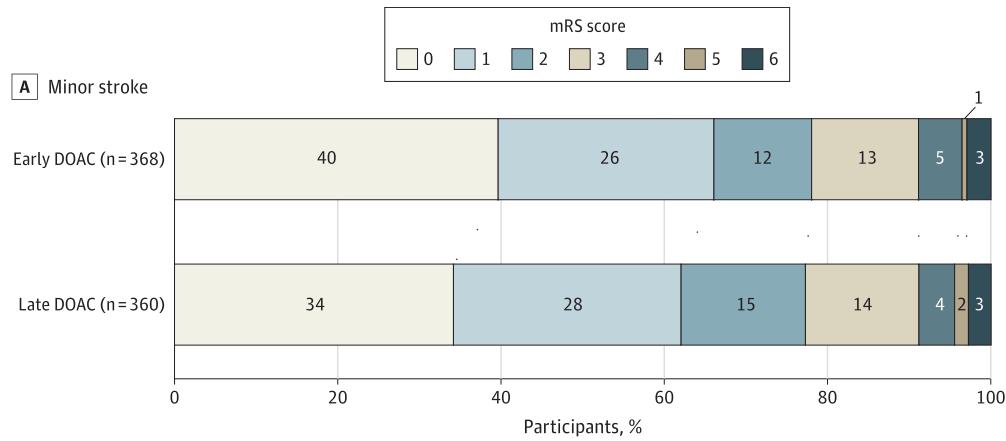
**B** Moderate stroke



**C** Major stroke



The incidences of the primary and secondary outcomes at 30 days were comparable for all infarct size groups, including minor (A), moderate (B), and major (C) stroke. DOAC indicates direct oral anticoagulant.



In all infarct size groups, including minor (A), moderate (B), and major (C) stroke, the mRS score distribution at 90 days was similar between early and late direct oral anticoagulant (DOAC) initiation. For mRS score shifts between early and late DOAC initiation, the odds ratio was 0.86 (95% CI, 0.65-1.13) for minor stroke, 0.92 (95% CI, 0.69-1.22) for moderate stroke, and 1.04 (95% CI, 0.69-1.56) for major stroke.

# Recomendaciones

Ictus o AIT asociado a FA no valvular tras comprobar que no hay hemorragia...

## Las guías del 2019

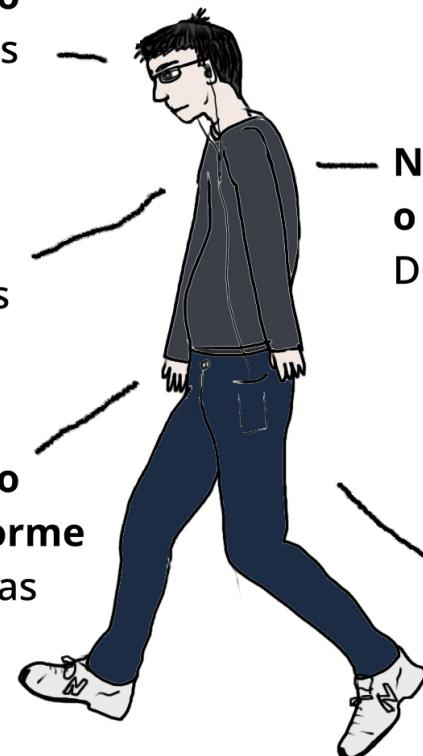
AIT o infarto

AAS 48 horas

NIHSS 8-16

DOAC 7 días

NIHSS >16 o  
infarto enorme  
DOAC 14 días



NIHSS <8

o <1,5 cm

DOAC 3-4 días

Infarto grande  
DOAC 7 días

## Me acabo de leer un artículo

AIT

DOAC <24 horas

OUCH!



Infarto pequeño o grande  
DOAC <48 horas

Infarto enorme  
DOAC 7 días

# Recomendaciones

- En AIT se podría anticoagular en 24 horas con imagen
- En infartos pequeños o medianos parece seguro  $\leq 48$  horas
- En infartos mayores es posible a la semana
- Las guías contemplan dos semanas en infartos grandes
- La evidencia actualmente no apoya ninguna actitud<sup>1</sup>

1. pero parece favorecer tiempos más cortos

# Muchas gracias

Podéis encontrar estas diapositivas en

[cerebro.neocities.org](http://cerebro.neocities.org)

## Referencias

1. Goeldlin, M. B. et al. Early vs Late Anticoagulation in Minor, Moderate, and Major Ischemic Stroke With Atrial Fibrillation: Post Hoc Analysis of the ELAN Randomized Clinical Trial. *JAMA Neurology* **81**, 693 (2024).
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5. Fischer, U. et al. Early versus Later Anticoagulation for Stroke with Atrial Fibrillation. *New England Journal of Medicine* **388**, 2411-2421 (2023).