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TROMBOEMBOLISMO

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FORUM

MULTIDISCIPLINARIA  
DE LA



21-22 OCTUBRE 2010

HOTEL ABADES NEVADA PALACE - GRANADA

# ETV tras hemorragia intacraneal

Dr. Lobo. *H Txagorritxu*

Dr. Nieto. *H Virgen de la Luz*

Dr. Monreal. *H Trias i Pujol*

- **Hemiplejía por HIC (15 d)**
- **Disnea y datos de TVP**



# Venous thromboembolism and treatment in patients with acute stroke and traumatic brain injury

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Current Opinion in Critical Care 2008, 14:149–155

## Purpose of review

Patients with acute stroke and traumatic brain injury are at risk to develop venous thromboembolism. This review analyzes the available literature to propose guidelines for the prevention and treatment of venous thromboembolism in these groups of patients.

## Recent findings

In acute ischemic stroke, low-dose low-molecular-weight heparin has the best benefit–risk ratio to prevent venous thromboembolism. Patients with primary intracerebral hemorrhage and traumatic brain injury should receive intermittent pneumatic compression, followed by low-dose low-molecular-weight heparin or unfractionated heparin 3–4 days after stroke onset or 24 h after injury or surgery, respectively, and after cessation of bleeding. Concerning treatment, in patients with deep-vein thrombosis lower doses of heparin are indicated to prevent pulmonary embolism, and a vena cava filter should be considered. In patients with pulmonary embolism, treatment could be more aggressive, because of a high mortality risk.

## Summary

Adequate prevention of venous thromboembolism with intermittent pneumatic compression or pharmacological prophylaxis is important. The best treatment of venous thromboembolism remains unclear. In case of pulmonary embolism, more aggressive treatment is warranted.

## Keywords

acute stroke, anticoagulant treatment, traumatic brain injury, venous thromboembolism

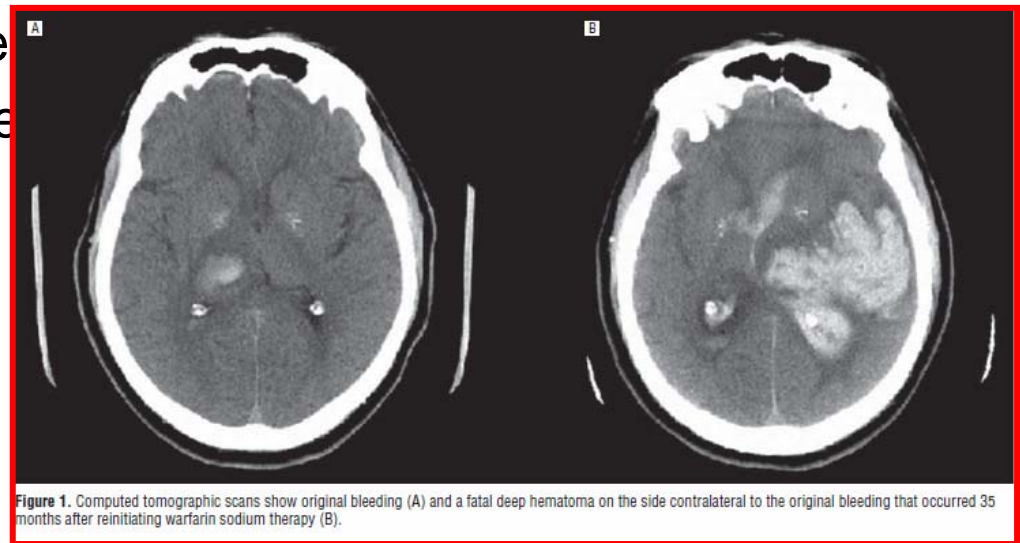
Curr Opin Crit Care 14:149–155  
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1070-5295

..”Data on treatment of VTE in patients with intracranial hemorrhage are extremely scarce”.



# Hemorragia Intracerebral

- El 10-15% de los ACV
- Mortalidad a 30 días del 35-52%
  - la mitad en los primeros dos días.
- Tendencia a recidiva espontánea
- Solo 20% autónomos a los 6 meses





ELSEVIER

## Deep venous thrombosis after acute intracerebral hemorrhage <sup>☆</sup>

Toshiyasu Ogata <sup>a,\*</sup>, Masahiro Yasaka <sup>a</sup>, Yoshiyuki Wakugawa <sup>a</sup>, Tooru Inoue <sup>b</sup>,  
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Available online 13 J

### Abstract

**Background:** We evaluated the incidence of deep venous thrombosis (DVT) DVT.

**Methods:** We enrolled 52 patients with acute ICH between June 2005 and Se deficit, hemorrhage size and laboratory data, and performed ultrasonography  
**Results:** DVT was detected a total of 21 patients (40.4%) after two weeks. F severe disturbance of consciousness ( $p=0.020$ ) and paralysis ( $p=0.035$ ) or Health Stroke Scale (NIHSS) score was significantly higher in patients w diameter of ICH were more likely to develop DVT ( $p=0.021$ ). D-dimer va than those without ( $p=0.002$ ). Logistic regression analysis indicated that bo for the occurrence of DVT.

**Conclusions:** We need be aware that acute ICH patients with severe neur developing DVT.

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**Keywords:** Deep venous thrombosis; Intracerebral hemorrhage; Neurological finding

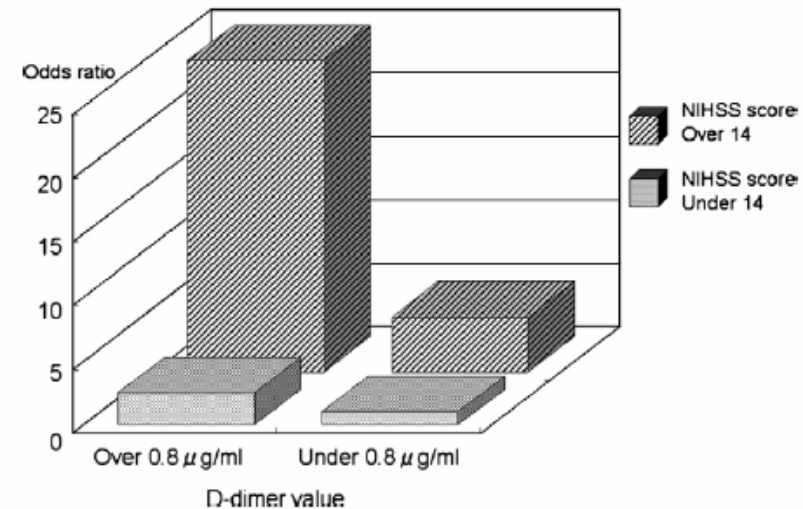


Fig. 1. Odds ratio for the occurrence of DVT in patients with NIHSS score under 14 and D-dimer value under 0.8 μg/ml. DVT denotes deep vein thrombosis; NIHSS, National Institutes of Health and Stroke Scale.







## Antithrombotic Therapy for Venous Thromboembolic Disease\*

American College of Chest Physicians Evidence-Based Clinical Practice Guidelines (8th Edition)

1.13.2. For patients with acute proximal DVT, if anticoagulant therapy is not possible because of the risk of bleeding, we recommend

placement of an IVC filter (Grade 1C).

4.6.2. In patients with acute PE, if anticoagulant therapy is not possible because of risk of bleeding, we recommend placement of an IVC filter (Grade 1C).



**Table 1** Clinical characteristics and 3 month outcome of the 12 294 patients with venous thromboembolism (VTE), according to the site of recent bleeding

	Gastrointestinal (GI), n = 116	Intracranial, n = 94	Other, n = 96	No recent bleeding, n = 11 988
<b>Clinical characteristics</b>				
Gender (males)	56 (48%)	49 (52%)	44 (46%)	5990 (50%)
Age [years (mean ± SD)]	71 ± 15 <sup>†</sup>	66 ± 13	63 ± 19	66 ± 17
<b>Underlying diseases</b>				
Creatinine levels > 1.2 mg dL <sup>-1</sup>	22 (19%)	5 (5.3%)*	13 (14%)	1675 (14%)
Chronic lung disease	19 (18%)	6 (6.4%)	9 (9.4%)	1345 (13%)
Chronic heart failure	11 (10%)	3 (3.2%)	8 (8.3%)	719 (6.8%)
<b>Risk factors for VTE</b>				
Cancer	38 (33%) <sup>†</sup>	7 (7.4%) <sup>†</sup>	37 (39%) <sup>†</sup>	2415 (20%)
Surgery < 2 months	30 (26%) <sup>†</sup>	29 (31%) <sup>†</sup>	32 (33%) <sup>†</sup>	1541 (13%)
Prior VTE	12 (10%)	6 (6.4%) <sup>†</sup>	11 (11%)	1947 (16%)
<b>Clinical presentation</b>				
Symptomatic PE	52 (45%)	43 (46%)	47 (49%)	5209 (43%)
<b>Time elapsed since bleeding</b>				
Mean days ± SD	17 ± 10	20 ± 9	13 ± 10	
Median (days)	15	20	11	
<b>Initial therapy</b>				
UFH	17 (15%)*	6 (6.4%)	15 (16%)*	984 (8.2%)
Mean UFH dose (IU kg <sup>-1</sup> d <sup>-1</sup> )	332 ± 72	262 ± 100	345 ± 93	340 ± 109
LMWH	98 (84%)*	83 (88%)	78 (81%) <sup>†</sup>	10 853 (91%)
Mean LMWH dose (IU kg <sup>-1</sup> d <sup>-1</sup> )	162 ± 52 <sup>†</sup>	136 ± 66 <sup>†</sup>	166 ± 52 <sup>†</sup>	182 ± 38
IVC filter	16 (14%) <sup>†</sup>	28 (30%) <sup>†</sup>	10 (10%) <sup>†</sup>	188 (1.6%)
<b>Time to initial therapy</b>				
Mean ± SD (days)	0.1 ± 0.5	0.4 ± 3.3	0.3 ± 3.6	0.04 ± 0.6
Median (days)	0	0	0	0
<b>Long-term therapy</b>				
AVK drugs	50 (43%) <sup>†</sup>	23 (24%) <sup>†</sup>	43 (45%) <sup>†</sup>	8481 (71%)
LMWH	49 (42%) <sup>†</sup>	61 (65%) <sup>†</sup>	41 (43%) <sup>†</sup>	2953 (25%)
<b>Three-month outcome</b>				
Major bleeding	12 (10%) <sup>†</sup>	0	7 (7.3%) <sup>†</sup>	276 (2.3%)
Fatal bleeding	7 (6.0%) <sup>†</sup>	0	1 (1.0%)	61 (0.5%)
Recurrent VTE	3 (2.6%)	5 (5.3%)	3 (3.1%)	321 (2.7%)
Overall mortality	20 (17%) <sup>†</sup>	4 (4.3%)	19 (20%) <sup>†</sup>	964 (8.0%)

Comparisons with patients with no recent bleeding: \* $P < 0.05$ ; <sup>†</sup> $P < 0.01$ ; <sup>‡</sup> $P < 0.001$ . PE, pulmonary embolism; UFH, unfractionated heparin; LMWH, low-molecular-weight heparin; SD, standard deviation; IVC, inferior vena cava; AVK, anti-vitamin K drugs.

# Restarting Anticoagulation Therapy After Warfarin-Associated Intracerebral Hemorrhage

Daniel O. Claassen, MD; Noojan Kazemi, MBBS; Alexander Y. Zubkov, MD, PhD; Eelco F. M. Wijdicks, MD; Alejandro A. Rabinstein, MD

**Table 3. Follow-up Data in 48 Patients With Warfarin-Associated ICH**

Patients restarted warfarin therapy after a median of 10 (range, 7-28) days.

**Table 1. Clinical Characteristics<sup>a</sup>**

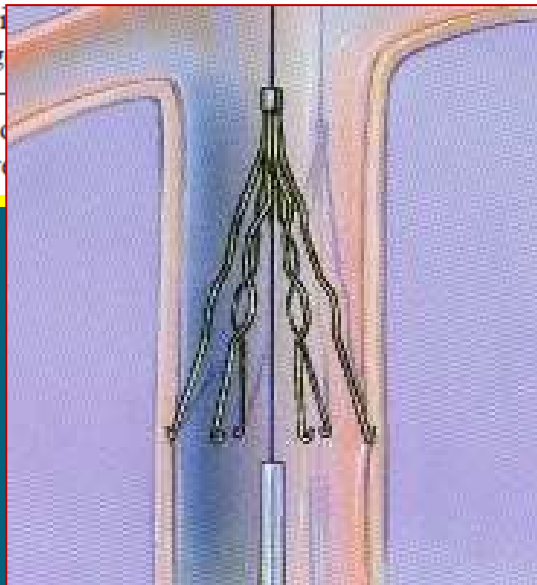
Variable	Restarted Group (n = 23)
Age, mean (range), y	70.8 (45-)
Sex, No. M/F	13/10
Reason for anticoagulation	
Atrial fibrillation	9 (39)
Valve replacement	10 (44)
Thrombus/DVT	3 (13)
Other <sup>c</sup>	1 (4)
Hypertension treated with medication	19 (83)
Diabetes mellitus	5 (22)
Coronary artery disease/CHF	5 (22)
Previous stroke	7 (30)
Concomitant neoplasm	1 (4)

main outcome measures thromboembolic events.

	Restarted Group	Nonrestarted Group
Mean follow-up, mo	49.8	36.1
Mean mRS score		
At discharge	3.1	2.6
At latest follow-up	4.6	3.7
Mean time to death, mo	55.6	21.8
End point events, No. of patients		
Thromboembolic stroke	0	3
Thromboembolism, nonstroke	0	2
Nonembolic ischemic stroke	2	2
Nontraumatic ICH	1	0
Traumatic ICH	2	0
GI hemorrhage	2	2
Myocardial infarction	4	6

# Anticoagulation or Inferior Vena Cava Filter Placement for Patients With Primary Intracerebral Hemorrhage

No hay datos experimentales para sustentar ningún tipo de recomendación



## Definition of Inadequate Cardiopulmonary Reserve in Patients With Suspected PE in the Study of Hull et al<sup>31</sup>

Inadequate cardiopulmonary reserve defined by any of the following:

Hypotension (systolic blood pressure <90 mm Hg)

Syncope

Right ventricular failure

Pulmonary edema

Acute tachyarrhythmias

Respiratory failure (any of the following:  $P_{O_2}$  <50 mm Hg,  $P_{CO_2}$  >45 mm Hg,  $FEV_1$  <1.0 L, vital capacity <1.5 L)

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# No hay experiencia clínica acumulada

...la recomendación más habitual de los expertos es la colocación de un Filtro de Cava

Of an inferior vena caval filter...  
data are sparse and not suitable for  
is open for randomized trials in  
patients.  
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oli are common preventable causes of mor-  
cerebral hemorrhage (ICH). The frequency of  
h acute ICH ranges from 0.5 to 13% in scant  
reatment of these complications is to reduce  
of intracranial rebleeding. There is a paucity  
ability of the recommendations for patients

estima que la probabilidad de EP fatal si  
a un EP o TVP proximal sintomáticos es  
de alrededor del 25%, y sin embargo la  
probabilidad de resangrado intracraneal con  
tratamiento anticoagulante es (3-5 veces el  
espontáneo) de 3-5%, y “solo” la mitad de ellos  
fallecerá.

# AHA/ASA Guideline

## Guidelines for the Management of Spontaneous Intracerebral Hemorrhage

### A Guideline for Healthcare Professionals From the American Heart Association/American Stroke Association

*The American Academy of Neurology affirms the value of this guideline as an educational tool for neurologists.*

*The American Association of Neurological Surgeons and the Congress of Neurological Surgeons have reviewed this document and affirm its educational content.*

Lewis B. Morgenstern, MD, FAHA, FAAN, Chair;  
J. Claude Hemphill III, MD, MAS, FAAN, Vice-Chair; Craig Anderson, MBBS, PhD, FRACP;  
Kyra Becker, MD; Joseph P. Broderick, MD, FAHA; E. Sander Connolly, Jr, MD, FAHA;  
Steven M. Greenberg, MD, PhD, FAHA, FAAN; James N. Huang, MD; R. Loch Macdonald, MD, PhD;  
Steven R. Messé, MD, FAHA; Pamela H. Mitchell, RN, PhD, FAHA, FAAN;  
Magdy Selim, MD, PhD, FAHA; Rafael J. Tamargo, MD; on behalf of the American Heart Association Stroke Council and Council on Cardiovascular Nursing

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*Results—*

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*Purpose*—The aim of this guideline is to present current and comprehensive recommendations for the diagnosis and treatment of acute spontaneous intracerebral hemorrhage.

*Methods*—A formal literature search of MEDLINE was performed. Data were synthesized with the use of evidence tables.

Writing committee members met by teleconference to discuss data-derived recommendations. The American Heart Association Stroke Council's Levels of Evidence grading algorithm was used to grade each recommendation. Prerelease review of the draft guideline was performed by 6 expert peer reviewers and by the members of the Stroke Council Scientific Statements Oversight Committee and Stroke Council Leadership Committee. It is intended that this guideline be fully updated in 3 years' time.

*Results*—Evidence-based guidelines are presented for the care of patients presenting with intracerebral hemorrhage. The focus was subdivided into diagnosis, hemostasis, blood pressure management, inpatient and nursing management, preventing medical comorbidities, surgical treatment, outcome prediction, rehabilitation, prevention of recurrence, and future considerations.

*Conclusions*—Intracerebral hemorrhage is a serious medical condition for which outcome can be impacted by early, aggressive care. The guidelines offer a framework for goal-directed treatment of the patient with intracerebral hemorrhage. (*Stroke*. 2010;41:2108-2129.)



# R.



- En Agosto 2009 había en RIETE 27.029 pacientes ; 141 (0.5%) de ellos habían padecido una HIC reciente

**Table I. Clinical characteristics, treatment strategies and 3-month outcome of 27,029 VTE patients with or without recent cerebral hemorrhage.**

	<b>Cerebral hemorrhage</b>	<b>No cerebral hemorrhage</b>	<b>p value</b>
<b>Patients, N</b>	<b>141</b>	<b>26888</b>	
<b>Clinical characteristics,</b>			
Gender (males)	77 (55%)	13168 (49%)	0.206
Age (mean, IQR)	71 (58-77)	70 (56-79)	0.871
Age >70 years	76 (54%)	13377 (50%)	0.399
Body weight (mean, IQR)	70 (64-80)	73 (64-82)	0.054
<b>Additional risk factors for VTE,</b>			
Cancer	12 (8.5%)	5644 (21%)	<0.001
Prior VTE	6 (4.3%)	4190 (16%)	<0.001
<b>Underlying diseases,</b>			
Chronic lung disease	9 (6.4%)	2665 (9.9%)	0.201
Chronic heart disease	6 (4.3%)	1504 (5.6%)	0.711
CrCl levels <30 mL/min	9 (6.4%)	1706 (6.4%)	0.947
<b>Initial VTE presentation,</b>			
Pulmonary embolism	70 (50%)	12625 (47%)	0.612
<b>Initial therapy,</b>			
LMWH	116 (82%)	24270 (90%)	0.004
LMWH, IU/kg/day (mean, IQR)	133 (71-185)	187 (163-200)	<0.001
UFH	14 (9.9%)	2029 (7.6%)	0.264
UFH, IU/kg/day (mean, IQR)	267 (235-393)	360 (300-426)	0.043
Inferior vena cava filter	44 (31%)	560 (2.1%)	<0.001
No drug therapy	3 (2.1%)	171 (0.6%)	0.063
<b>Long term therapy,</b>			
Vitamin K antagonists	30 (21%)	18895 (70%)	<0.001
LMWH	95 (67%)	6532 (24%)	<0.001
LMWH, IU/Kg/day (mean, IQR)	110 (68-163)	150 (112-180)	<0.001
<b>90-day outcome,</b>			
Major bleeding	3 (2.1%)	588 (2.2%)	0.970
Fatal bleeding	0	156 (0.6%)	0.441
Recurrent DVT	5 (3.5%)	278 (1.0%)	0.017
Recurrent PE	4 (2.8%)	286 (1.1%)	0.067
Fatal PE	7 (5.0%)	426 (1.6%)	0.008
Fatal PE initial	4 (2.8%)	322 (1.2%)	0.093
Fatal PE recurrent	3 (2.1%)	104 (0.4%)	0.019
Overall death	14 (9.9%)	2103 (7.9%)	0.346

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<b>Pacientes, N</b>	<b>52</b>	<b>81</b>	
<b>Initial VTE presentation,</b>			
With PE	27 (52%)	38 (47%)	0.59
TAS (mmHg)	127 ± 32	122 ± 25	0.35
pO2	64 ± 14	69 ± 18	0.23
<b>Initial therapy,</b>			
LMWH, IU/kg/day (X ± SD)	<b>120 ± 86</b>	139 ± 58	0.15
LMWH dosis <100 IU/kg/day	20 (39%)	17 (21%)	<b>0.03</b>
Inferior vena cava filter	20 (38%)	20 (25%)	0.12
No drug therapy	1 (1.9%)	2 (2.5%)	1.00
<b>Eventos</b>			
Hemorragia grave	2 (3.8%)	1 (1.2%)	0.56
Recurrencia TVP	4 (7.7%)	1 (1.2%)	0.08
Recurrencia EP	1 (1.9%)	3 (3.7%)	1.00
EP mortal	2 (3.8%)	5 (6.2%)	0.70
Mortalidad global	6 (12%)	8 (9.9%)	0.78

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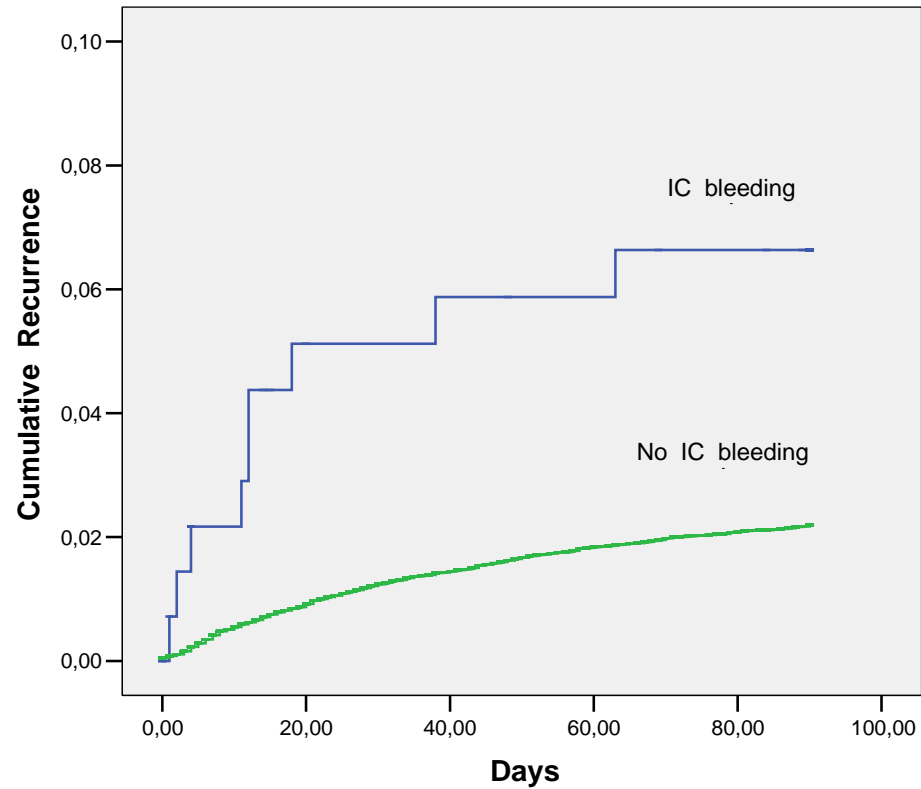
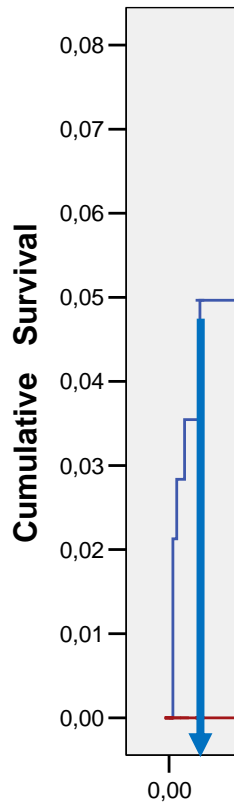
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<b>Initial therapy,</b>			
LMWH	116 (82%)	24270 (90%)	0.004
LMWH, IU/kg/day (mean, IQR)	133 (71-185)	187 (163-200)	<0.001
UFH	14 (9.9%)	2029 (7.6%)	0.264
UFH, IU/kg/day (mean, IQR)	267 (235-393)	360 (300-426)	0.043
Inferior vena cava filter	44 (31%)	560 (2.1%)	<0.001
No drug therapy	3 (2.1%)	171 (0.6%)	0.063
<b>Long term therapy,</b>			
Vitamin K antagonists	30 (21%)	18895 (70%)	<0.001
LMWH	95 (67%)	6532 (24%)	<0.001
LMWH, IU/Kg/day (mean, IQR)	110 (68-163)	150 (112-180)	<0.001
<b>90-day outcome,</b>			
Major bleeding	3 (2.1%)	588 (2.2%)	0.970
Fatal bleeding	0	156 (0.6%)	0.441
Recurrent DVT	5 (3.5%)	278 (1.0%)	0.017
Recurrent PE	4 (2.8%)	286 (1.1%)	0.067
Fatal PE	7 (5.0%)	426 (1.6%)	0.008
Fatal PE initial	4 (2.8%)	322 (1.2%)	0.093
Fatal PE recurrent	3 (2.1%)	104 (0.4%)	0.019
Overall death	14 (9.9%)	2103 (7.9%)	0.346

...the incidence of fatal PE was higher in patients with HIC (5.0% vs. 1.6%; OR: 3.2; 95% CI: 1.5-6.9).



Las recurrencias fueron 3 veces más frecuentes en los pacientes con HIC que en los pacientes normales

Todos los EP fatales se produjeron en la primera semana

Table IV. Univariate analysis for fatal PE or major bleeding.

	Fatal PE	No fatal PE	Major bleeding	No major bleeding
<b>Patients, N</b>	<b>7</b>	<b>134</b>	<b>3</b>	<b>138</b>
<b>Clinical characteristics,</b>				
Gender (males)	3 (43%)	74 (55%)	1 (33%)	76 (55%)
Age >70 years	4 (57%)	72 (54%)	0	76 (55%)
Body weight <	2 (29%)	27 (20%)	1 (33%)	28 (20%)
Inpatients	5 (71%)	102 (76%)	2 (67%)	105 (76%)
<b>Additional risk factors for VTE,</b>				
Cancer	1 (14%)	11 (8.2%)	0	12 (8.7%)
Prior VTE	0	6 (4.5%)	1 (33%)	5 (3.6%)
<b>Underlying diseases,</b>				
Chronic lung disease	0	9 (6.7%)	0	9 (6.5%)
Chronic heart failure	0	6 (4.5%)	0	6 (4.4%)
CrCl levels <30 mL/min	2 (29%)	77 (5.2%)	1 (33%)	8 (5.8%)
<b>VTE presentation,</b>				
Symptomatic PE	4 (51%)	66 (49%)	2 (67%)	68 (49%)
<b>Initial therapy,</b>				
Low-molecular-weight heparin	3 (43%)*	113 (84%)	3 (100%)	113 (82%)
LMWH dose IU/kg/day (mean, IQR)	81 (67-198)	133 (71-185)	100 (80-171)	133 (69-186)
Unfractionated heparin	2 (29%)	12 (9.0%)	0	14 (10%)
Inferior vena cava filter	0	44 (33%)	3 (100%)*	41 (30%)
<b>Long term therapy,</b>				
Vitamin K antagonists	0	30 (22%)	0	30 (22%)
Low-molecular-weight heparin	1 (14%)†	94 (70%)	3 (100%)	92 (67%)

# Conclusiones I

- La ETV es frecuente tras un episodio de HIC grave
- La HBMP no esta contraindicada en la profilaxis
- No hay base experimental para realizar recomendaciones terapéuticas

# Conclusiones II

- Las dosis “intermedias” de HBPM parecen relativamente seguras
- Con las pautas habituales la mortalidad por EP es muy superior a la morbimortalidad por hemorragia.
- En la práctica habitual se utilizan menos filtros de los que sería deseable.