

# Prévention de la mort subite dans la maladie coronaire

## **Place du traitement médical**

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# DÉCLARATION DE LIENS D'INTÉRÊT AVEC LA PRÉSENTATION

**Intervenant : Cynthia Barbraud, Liège**

Je n'ai pas de lien d'intérêt à déclarer

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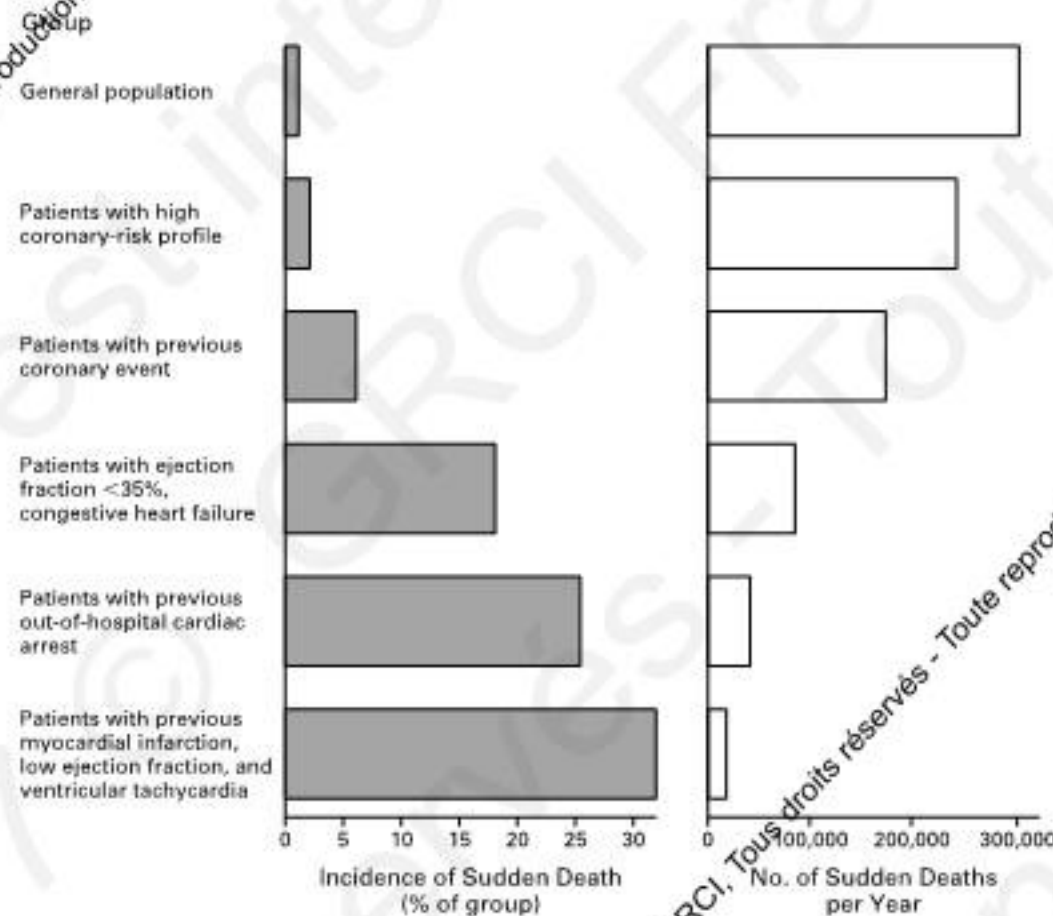
# Mort subite : 1ere manifestation de la coronaropathie

- >30% des MSC sont la manifestation initiale de la maladie coronaire
- Coronaropathie identifiée dans 80% des MSC

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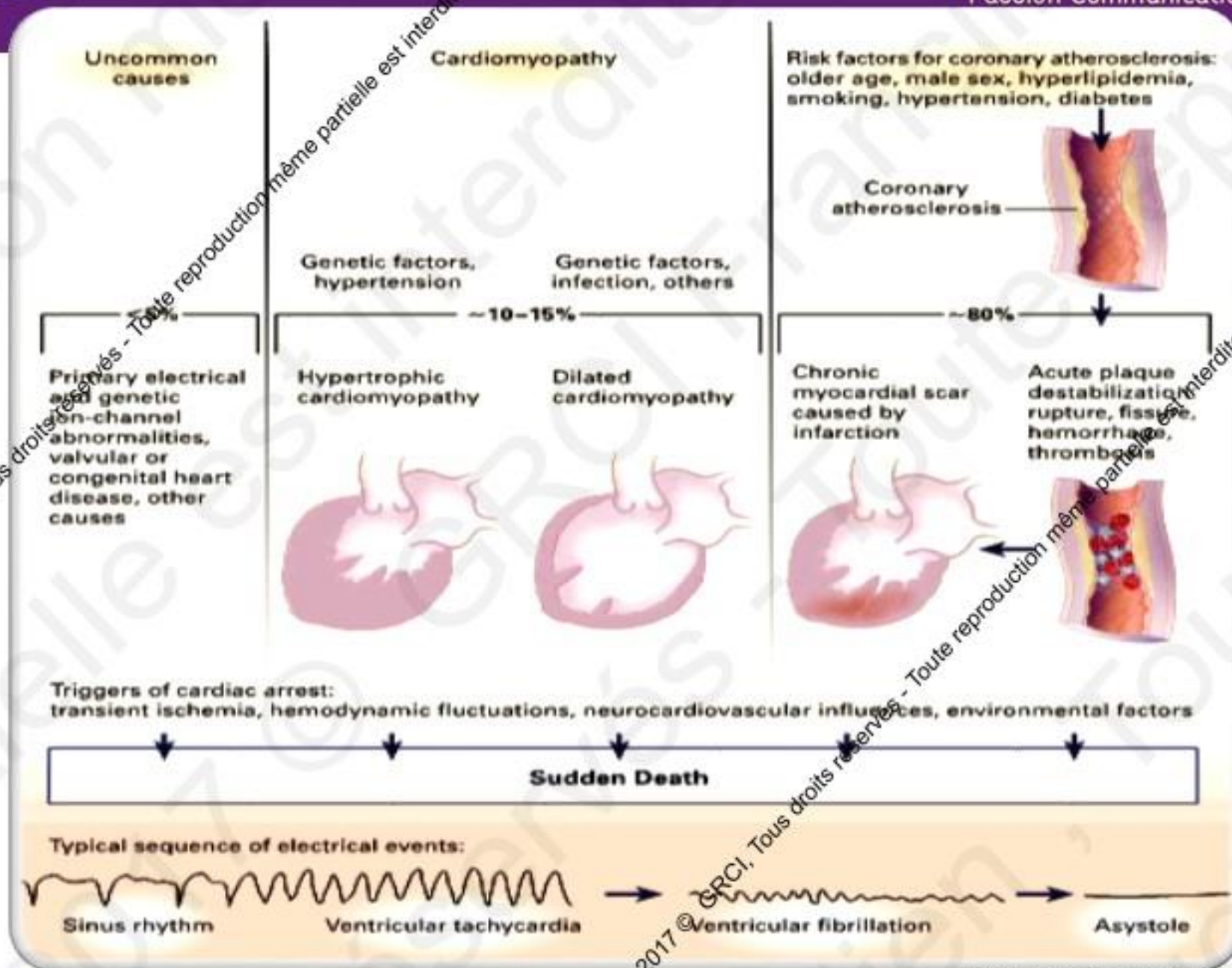
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# Incidence de la mort subite



**Figure 1.** The Incidence of Sudden Death in Specific Populations and the Annual Numbers of Sudden Deaths in Those Populations. Most of the deaths occur in the larger, lower-risk subgroups. Modified from Myerburg et al.<sup>10</sup> with the permission of the publisher.

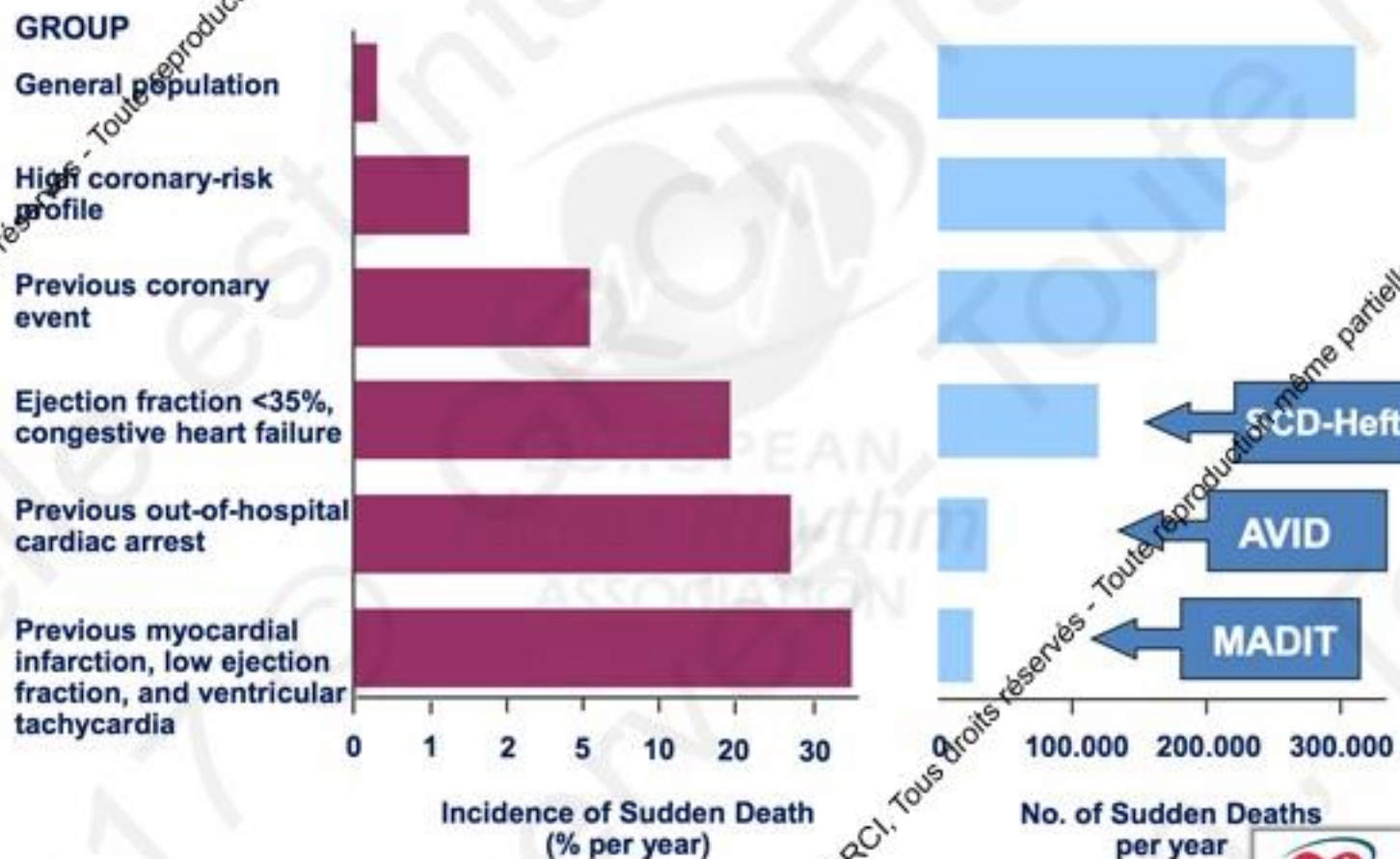




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## Incidence of SCD in Specific Populations



Myerburg RJ. *Circulation*. 1998;97:1514-1521.



# Facteurs de risque de mort subite

- Prévention primaire population générale, facteurs de risque cardiovasculaire
- Prévention secondaire post SCA :
  - Ischémie résiduelle
  - Dysfonction VG
  - Présence d'ESV
  - Âge

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TABLE IIB Risk factors associated with short-term mortality from day of index myocardial infarction up to 45 days. Patients from TRACE and DIAMOND only were analyzed

Risk factors	All-cause mortality		Arrhythmic mortality		Nonarrhythmic cardiac mortality	
	OR (95% CI)	p Value	OR (95% CI)	p Value	OR (95% CI)	p Value
Age ( $\uparrow$ 10 years)	1.52 (1.24–1.87)	<0.001	1.29 (1.00–1.66)	0.05	1.52 (1.12–2.06)	0.007
Smoker	1.58 (1.00–2.49)	0.05	1.59 (0.85–2.98)	0.1	1.07 (0.58–1.98)	0.8
Previous MI	1.44 (0.98–2.11)	0.06	1.84 (1.09–3.09)	0.02	1.18 (0.67–2.07)	0.6
Previous hypertension	1.72 (1.13–2.60)	0.01	2.28 (1.34–3.89)	0.003	1.16 (0.64–2.12)	0.1
SBP ( $\uparrow$ by 10%)	0.80 (0.71–0.90)	<0.001	0.73 (0.61–0.86)	0.001	0.87 (0.72–1.04)	0.1
HR ( $\uparrow$ by 10%)	1.23 (1.11–1.37)	<0.001	1.28 (1.11–1.47)	0.001	1.26 (1.09–1.47)	0.002
NYHA (compared with level I)		<0.001		0.007		<0.001
II	1.36 (0.78–2.35)		1.10 (0.53–2.25)		2.53 (0.85–7.56)	
III	2.16 (1.17–4.00)		1.47 (0.65–3.32)		5.39 (1.74–16.70)	
IV	8.35 (4.23–16.5)		4.50 (1.8–11.22)		28.19 (8.99–88.42)	
DIAMOND-MI vs. TRACE	1.39 (0.91–2.11)	0.1	2.06 (1.17–3.63)	0.01	1.33 (0.73–2.43)	0.4

Abbreviations: OR = odds ratio, CI = confidence interval. Other abbreviations as in Table 1.



# Mort subite et ischémie aiguë 24-48h

- Ischémie abaisse le seuil de dépolarisation,  $\nearrow$  tonus sympathique
- Reperfusion allonge la repolarisation : long QT local dans la zone ischémique
- Changement de l'électrophysiologie des myocytes déclenche réentrée

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# Beta Bloquants

- Effet anti ischémique et anti arythmique
- CAST 1994
- Registre VALIANT 2008 BB phase aiguë [Am J Cardiol. 2008 Dec 1;102\(11\):1427-32](#)

En cas de TV soutenue ou FV

5,8% arythmie en phase aiguë, risque x3 mortalité hospitalière

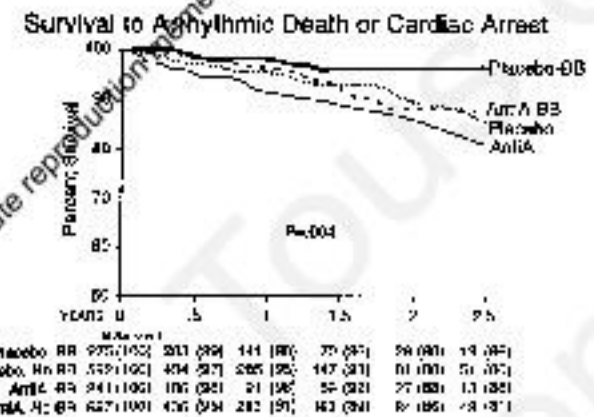
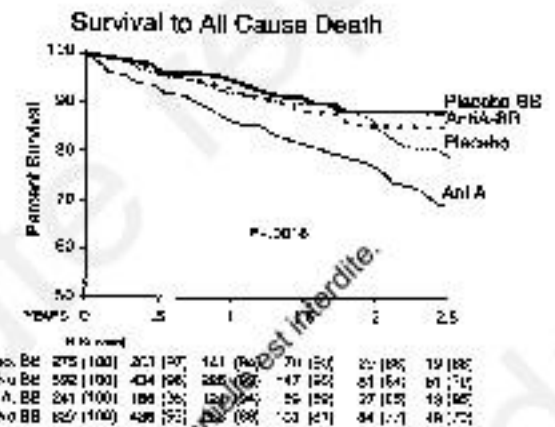
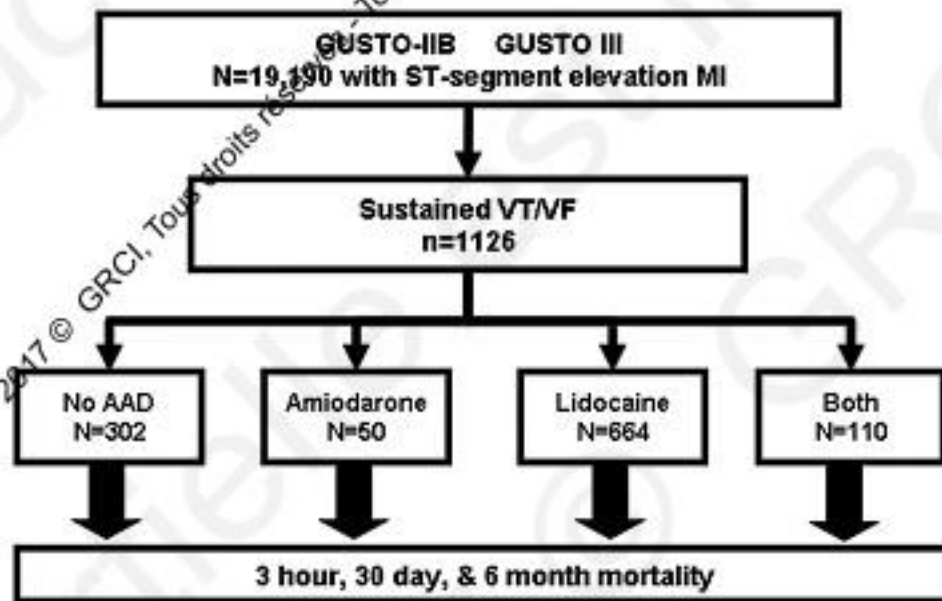


FIGURE 1. Kaplan-Meier survival of 1,725 CAST patients with an ejection fraction > 40% stratified for treatment with antiarrhythmic (AntiA) therapy or placebo and beta-blockers (BB) during 2.5 years of follow-up. BB, the end point was all-cause death; BB/BB the end point was arrhythmic death or nonfatal cardiac arrest. (Reproduced with permission from Kennedy et al.)

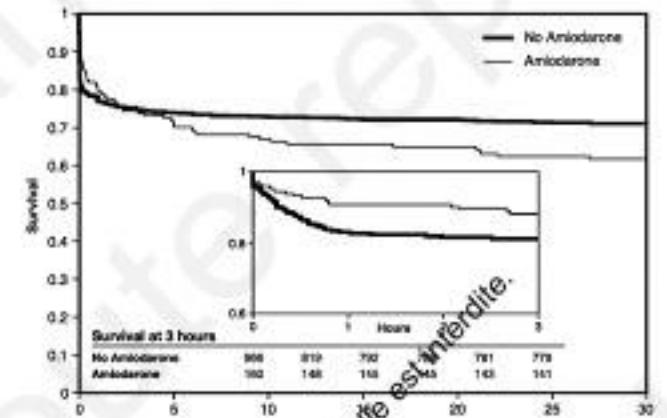
5. Kennedy HL, Brooks MM, Barker AH, Bergstrand R, Huther ML, Beanlands DS, Biggers JT, Goldstein S, for the CAST Investigators. Beta-blocker therapy in the Cardiac Arrhythmia Suppression Trial. *Am J Cardiol* 1994; 74: 674 – 680.

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# Lidocaïne

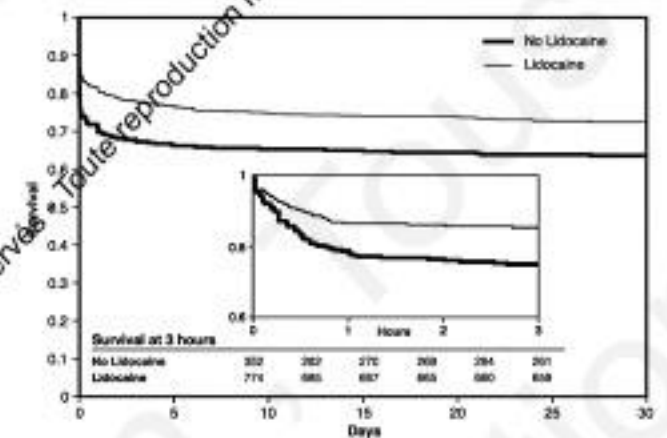


263. Piccini JP, Schulte PJ, Pieper KS, Mehta RH, White HD, Van de Werf F, Ardissino D, Califf RM, Granger CB, Ohman EM, Alexander JH. Antiarrhythmic drug therapy for sustained ventricular arrhythmias complicating acute myocardial infarction. Crit Care Med 2011;39:78-83.



Survival at 30 days

No Amiodarone	905	722	693	687	619	60
Amiodarone	180	112	138	132	131	85



Survival at 30 days

No Lidocaine	302	230	230	226	223	220	200
Lidocaine	774	691	676	668	665	660	637

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# Prévention associée aux SCA : phase hospitalière

Beta-blocker treatment is recommended for recurrent polymorphic VT.

**I**

**B**

Intravenous amiodarone is recommended for the treatment of polymorphic VT.

**I**

**C**

Intravenous lidocaine may be considered for the treatment of recurrent sustained VT or VF not responding to beta-blockers or amiodarone or in the presence of contraindications to amiodarone.

**IIb**

**C**

Prophylactic treatment with anti-arrhythmic drugs (other than beta-blockers) is not recommended.

**III**

**B**

Oral treatment with beta-blockers should be considered during the hospital stay and continued thereafter in all ACS patients without contraindications.

**IIa**

**B**

Radiofrequency catheter ablation at a specialized ablation centre followed by the implantation of an ICD should be considered in patients with recurrent VT, VF or electrical storms despite complete revascularization and optimal

**IIa**

**C**

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## Mort subite phase **précoce post SCA** 48h-3 mois

- 35% de risque de mort subite si FEVG basse
- Pas de bénéfice significatif d'implanter un défibrillateur, excès de mortalité totale (DINAMIT MADIT II)

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# Flécaïnide

- CAST study 6 jours-2 ans post SCA 1991
- ESV et FE < 55%

SUPPRESSION TRIAL — ECHT ET AL. 783

Death and Cardiac Arrest (with Resuscitation) in the CAST, According to Treatment Group.

	ENCAINIDE GROUP		FLÉCAÏNIDE GROUP		BOTH GROUPS		TOTAL
	ACTIVE DRUG	PLA-CEBO	ACTIVE DRUG	PLA-CEBO	ACTIVE DRUG	PLA-CEBO	
	432	425	323	318	755	743	1498
arrêts	44	19	19	7	63	261	89
arrest	42	15	18	6	60	211	81
ation	5	1	2	0	7	1	8
	29	12	14	4	43	161	59
ation	3	1	2	0	5	1	6
ic to	13	3	4	2	17	59	76
ation	2	0	0	0	2	0	2
	2	4	1	1	3	51	54

Units of categories.

or Flecainide or Corresponding Placebo.  
The number at risk is shown along the bottom.

these groups in mortality due to arrhythmia. There was a trend for more of the patients receiving active drug to have ventricular tachycardia or ventricular tachycardia degenerating into ventricular fibrillation on monitoring. More deaths due to arrhythmia in which asystole was the documented rhythm occurred in the active-treatment groups. There were also more patients receiving active drug in whom no monitoring was performed or for whom the monitored rhythm was unknown. Overall, the mean time from the onset of an event to monitoring was similar —  $12.4 \pm 11.7$  minutes in the active-drug groups and  $12.1 \pm 8.3$  minutes in the pla-

Table 3. First h

Ravvise

Ventricular tachycardia  
Ventricular tachycardia  
ventricular fibrillation  
Ventricular fibrillation  
Asystole

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# Amiodarone

## CAMIAT 1997:

- ESV post SCA, amiodarone 2ans vs placebo EMIAT
- Diminution 48% MSC et de 21 % de décès toutes causes NS
- 60% sous BB

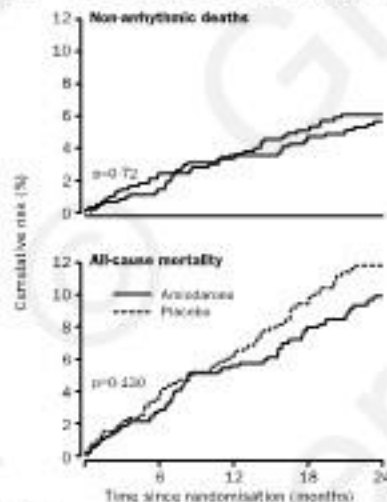


Figure 3: Rates of all-cause mortality and non-arhythmic death by intention-to-treat analysis

## EMIAT 1997 :

- FE<40% post SCA
- Diminution MSC mais pas de diminution mortalité toutes causes
- 44% sous BB

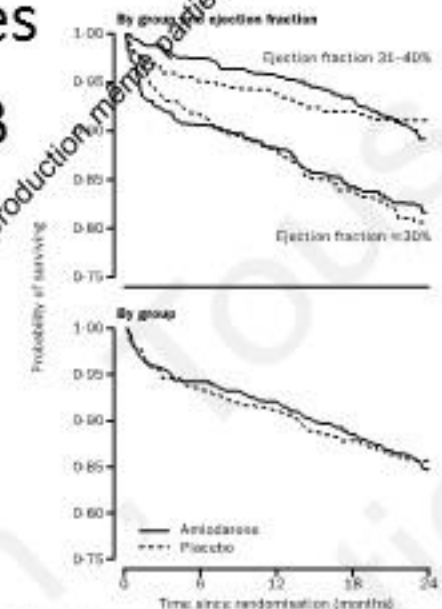


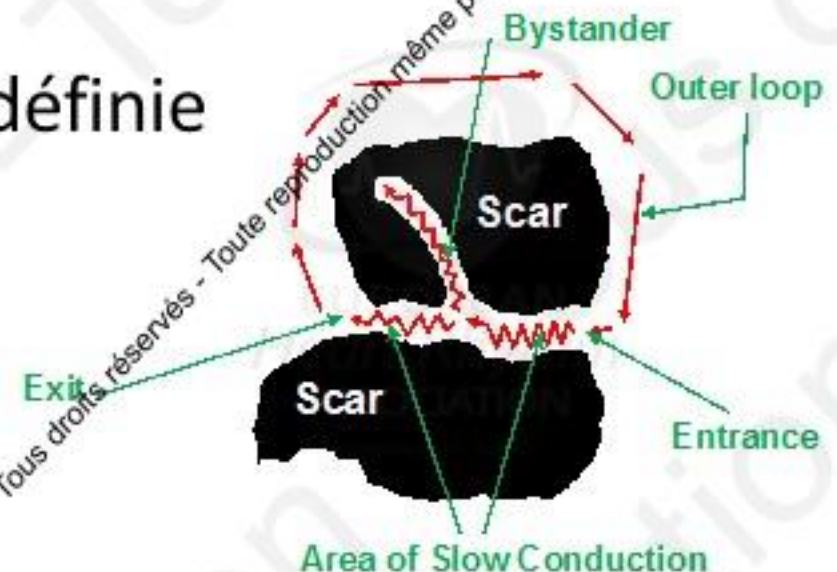
Figure 2: Kaplan-Meier estimates of all-cause mortality by group and ejection fraction

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# Mort subite et maladie coronarienne stable

- Remodeling ventriculaire et insuffisance cardiaque
- Cicatrice favorise la réentrée
- Limite temporelle du risque indéfinie



# Beta Bloquants

- Effet anti ischémique et anti arrythmique
- CAST 1994

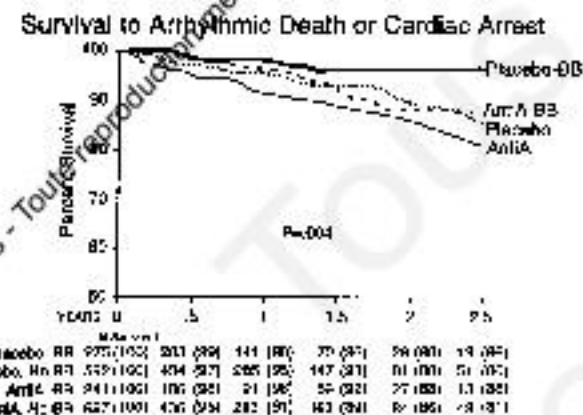


FIGURE 1. Kaplan-Meier survival of 1,125 CAST patients with an ejection fraction <math>\le 40\%</math> stratified for treatment with antiarrhythmic (AntiA) therapy or placebo and  $\beta</math>-blockers (BB) during 2.5 years of follow-up.  $\square$  The end point was all-cause death,  $\square$  both the end point was arrhythmic death or nonfatal cardiac arrest. Reproduced with permission$

5. Kennedy HL, Brooks AH, Barker AH, Bergstrand R, Huther ML, Beanlands DS, Bigler JT, Goldstein S, for the CAST Investigators. Beta-blocker therapy in the Cardiac Arrhythmia Suppression Trial. *Am J Cardiol* 1994; 74: 674 – 680.

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# Traitement de l'insuffisance cardiaque

## Recommendations to prevent or delay the development of overt heart failure or prevent death before the onset of symptoms

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref <sup>c</sup>
Treatment of hypertension is recommended to prevent or delay the onset of HF and prolong life.	I	A	126, 129, 150, 151
Treatment with statin is recommended in patients with or at high-risk of CAD whether or not they have LV systolic dysfunction, in order to prevent or delay the onset of HF and prolong life.	I	A	137–140, 152
Counselling and treatment for smoking cessation and alcohol intake reduction is recommended for people who smoke or who consume excess alcohol in order to prevent or delay the onset of HF.	I	C	131–133
Treating other risk factors of HF (e.g. obesity, dysglycaemia) should be considered in order to prevent or delay the onset of HF.	IIa	C	134, 141, 153–155
Empagliflozin should be considered in patients with type 2 diabetes in order to prevent or delay the onset of HF and prolong life.	IIa	C	130
ACE-I is recommended in patients with asymptomatic LV systolic dysfunction and a history of myocardial infarction in order to prevent or delay the onset of HF and prolong life.	I	A	5, 144, 145
ACE-I is recommended in patients with asymptomatic LV systolic dysfunction without a history of myocardial infarction, in order to prevent or delay the onset of HF.	I	B	5
ACE-I should be considered in patients with stable CAD even if they do not have LV systolic dysfunction, in order to prevent or delay the onset of HF.	IIa	A	142
Seta-blocker is recommended in patients with asymptomatic LV systolic dysfunction and a history of myocardial infarction, in order to prevent or delay the onset of HF or prolong life.	I	B	146
ICD is recommended in patients: a) with asymptomatic LV systolic dysfunction (LVEF ≤30%) of ischaemic origin, who are at least 40 days after acute myocardial infarction. b) with asymptomatic non-ischaemic dilated cardiomyopathy (LVEF ≤30%), who receive OM therapy, in order to prevent sudden death and prolong life.	I	B	149, 156–158

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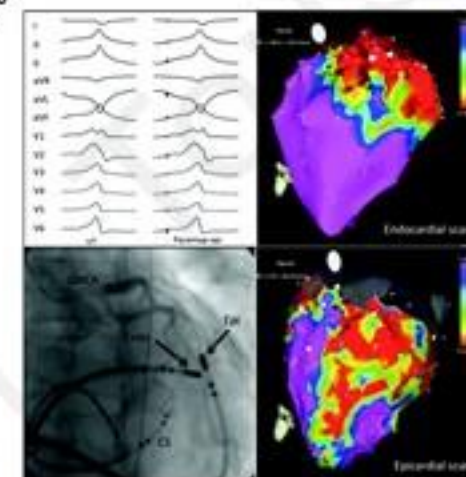
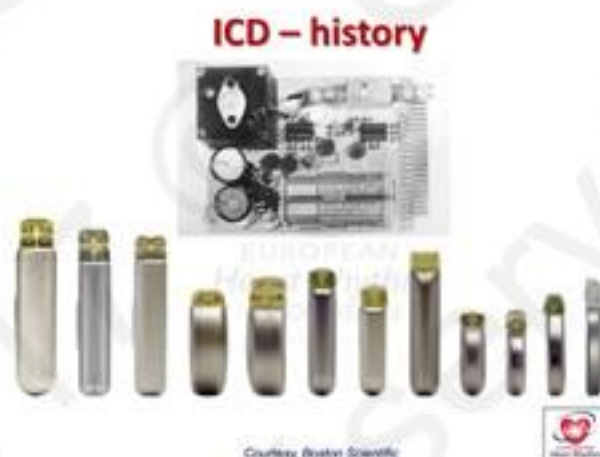
# Angiotensin–Neprilysin Inhibition vs. Enalapril

**Table 2. Primary and Secondary Outcomes.\***

Outcome	LCZ696 (N = 4187)	Enalapril (N = 4212)	Hazard Ratio or Difference (95% CI)	P Value
Primary composite outcome — no. (%)				
Death from cardiovascular causes or first hospitalization for worsening heart failure	914 (21.8)	1117 (26.5)	0.80 (0.73–0.87)	<0.001
Death from cardiovascular causes	558 (13.3)	693 (16.5)	0.80 (0.71–0.89)	<0.001
First hospitalization for worsening heart failure	537 (12.8)	658 (15.6)	0.79 (0.71–0.89)	<0.001
Secondary outcomes — no. (%)				
Death from any cause	711 (17.0)	835 (19.8)	0.84 (0.76–0.93)	<0.001
Change in KCCQ clinical summary score at 8 mo <sup>†</sup>	-2.99±0.36	-4.63±0.28	1.64 (0.63–2.65)	0.001
New-onset atrial fibrillation <sup>‡</sup>	84 (3.1)	83 (3.1)	0.97 (0.72–1.31)	0.83
Decline in renal function <sup>§</sup>	94 (2.2)	88 (2.6)	0.86 (0.65–1.13)	0.28



- Traitement médical au long cours indispensable
- Mais insuffisant si facteurs de risque en raison de l'incidence élevée de mort subite dans ce groupe à risque







### HOW THE POLYPILL WILL HELP YOU LIVE LONGER

Heart attack survivors' pill will include:

**Aspirin 75mg**  
(anti blood clotting)

**Simvastatin 40mg**  
(anti cholesterol)

**Atenolol 50mg**  
(blood pressure lowering)

**Lisinopril 10mg**  
(blood pressure lowering)

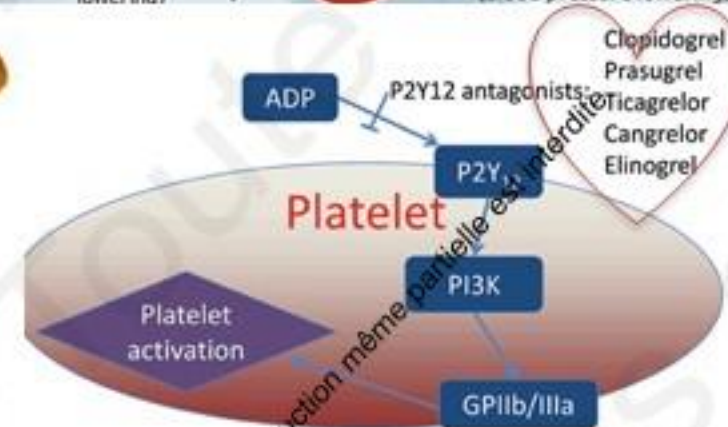
Stroke survivors' pill will include:

**Aspirin 75mg**  
(anti blood clotting)

**Simvastatin 40mg**  
(anti cholesterol)

**Lisinopril 10mg**  
(blood pressure lowering)

**Hydrochlorothiazide 12.5mg**  
(blood pressure lowering)



**VALSARTAN**

Valsartan blocks the effects of a body chemical. This relaxes blood vessel tightening and the buildup of sodium and fluid.

**SACUBITRIL**

Sacubitril blocks the activity of an enzyme called neprilysin. When neprilysin is active, it breaks down helpful peptides. Sacubitril prevents neprilysin so peptide levels can go up. These peptides help relax blood vessels and decrease sodium and fluid in the body.

ENTRESTO helps make it easier for your heart to do its job.



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