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Original research

Long-term survival benefit of ramipril in patients with acute myocardial infarction complicated by heart failure

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ABSTRACT

Aims ACE inhibition reduces mortality and morbidity in patients with heart failure after acute myocardial infarction (AMI). However, there are limited randomised data about the long-term survival benefits of ACE inhibition in this population.

Methods In 1993, the Acute Infarction Ramipril Efficacy (AIRE) study randomly allocated patients with AMI and clinical heart failure to ramipril or placebo. The duration of masked trial therapy in the UK cohort (603 patients, mean age=64.7 years, 455 male patients) was 12.4 and 13.4 months for ramipril (n=302) and placebo (n=301), respectively. We estimated life expectancy and extensions of life (difference in median survival times) according to duration of follow-up (range 0–29.6 years).

Results By 9 April 2019, death from all causes occurred in 266 (88.4%) patients in placebo arm and 275 (91.1%) patients in ramipril arm. The extension of life between ramipril and placebo groups was 14.5 months (95% CI 13.2 to 15.8). Ramipril increased life expectancy more for patients with than without diabetes (life expectancy difference 32.1 vs 5.0 months), previous AMI (20.1 vs 4.9 months), previous heart failure (19.5 vs 4.9 months), hypertension (16.6 vs 8.3 months), angina (16.2 vs 5.0 months) and age >65 years (11.3 vs 5.7 months). Given potential treatment switching, the true absolute treatment effect could be underestimated by 28%.

Conclusion For patients with clinically defined heart failure following AMI, ramipril results in a sustained survival benefit, and is associated with an extension of life of up to 14.5 months for, on average, 13 months treatment duration.

INTRODUCTION

Survival following acute myocardial infarction (AMI) remains a key objective for healthcare professionals and governments throughout the globe. Despite major advances in the emergency treatment and secondary prevention of AMI, a substantial proportion of patients with AMI develop heart failure,¹ which in turn is associated with high morbidity, mortality and healthcare expenditure.^{2–4}

Moreover, therapeutic inhibition of ACE has become an established strategy in the care of patients with AMI and heart failure.⁵ ACE inhibition reduces death and disease in patients with left ventricular systolic dysfunction, and coronary artery disease.^{6–8} In the Acute Infarction Ramipril

Efficacy (AIRE) study, ACE inhibition at an average follow-up time of 15 months after randomisation significantly reduced all-cause mortality, and continued to confer a survival benefit at 3 years following study closure in the AIRE Extension (AIREX) study.^{9–12} Further placebo-controlled survival studies testing the efficacy of ACE inhibition in patients with AMI and clinical heart failure are unlikely because withholding a treatment of accepted clinical value would be unethical. Therefore, in the AIRE survival (AIRE-S) study, we aimed to assess the long-term (29.6 years after randomisation) magnitude of impact on length of survival observed after treatment with ramipril (target dose 5 mg twice daily) when compared with placebo.

METHODS

Patients and design

The rationale, design, organisation, outcome definitions and results of the AIRE study have been published previously.^{10–12} This was an independent, multinational, multicentre, double-blind, randomised, placebo-controlled, parallel-group investigation into the potential benefits from treatment with the lipophilic ACE inhibitor ramipril (target dose 10 mg/day) following AMI complicated by clinical evidence of heart failure. When the trial was conducted, in 1993, all patients had an ECG and cardiac enzyme confirmation of AMI, and also evidence of transient or persisting heart failure as judged by the attending physician (chest X-ray evidence of left ventricular failure, or auscultatory evidence of pulmonary oedema, or a third heart sound in the presence of persistent tachycardia), usually sufficient to justify diuretic or vasodilator therapy. Patients were excluded if they had evidence of clinical instability, contraindications to the use of ACE inhibitor therapy, heart failure of primary valvular or congenital origin or overt heart failure deemed to require treatment with an open-label ACE inhibitor. After applying these entry criteria to consecutive patients admitted to the UK centres (30 centres), 44% of patients approached declined to give their consent to take part in the study. All 603 of the remaining patients were then randomised by telephone to receive masked treatment with either ramipril (titrated over 4 days to 5 mg twice daily) or a matching placebo. Treatment was started between 2 and 9 days after the onset of AMI. Patients initially received either 1.25 or 2.5 mg ramipril or

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Long-Term Management of Patients After Myocardial Infarction C. Tissa Kappagoda, P. V. Greenwood, 1987-11-01

Coronary Care Manual Peter L. Thompson, 2010-12-03 The Coronary Care Manual has been designed as a practical manual for the management of the acute coronary patient and aims to achieve a balance between the large and rapidly changing evidence base and practical application in the CCU ICU ED and ambulance The book has been written by a group of Australian and international experts for medical nursing and paramedic staff working in these areas This edition covers a broad range of coronary care medicine in enough detail to eliminate the need to refer to a larger reference book providing a portable companion for a night on call Completely redesigned with new artwork the manual is organized to suit academics and practitioners alike covers a broad range of coronary care medicine provides specific advice on the management of common clinical problems eliminates the need to refer to a larger reference book features a consistent style and focus with standardised artwork for figures is now also available as an eBook A code inside the Coronary Care Manual enables a full text download allowing you to browse and search electronically make notes and bookmarks in the electronic files and highlight material

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Cumulated Index Medicus ,1994

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associated with ageing which are essential to our understanding and management of the elderly sick population The title has been extensively revised and updates to reflect the enormous changes in treatment options and medical conditions emerged since publication of the third edition Written by worldwide experts of international repute this is the most up to date and comprehensive single reference source currently available Principles Practice of Geriatric Medicine Fourth Edition incorporates More than 30 new chapters including Preventive geriatrics Anorexia of Aging Managements of Weight Loss Dehydration Vitamins and Minerals in the Elderly Cancer and Aging Mild Cognitive Impairment Treatment of Behavioral Disorders The Older Patient with Down s Syndrome Drug Abuse in Older People Breast Cancer Women s health A truly global perspective including new chapter on Care of the elderly in Israel old age in a young land Geriatric Medicine in China Geriatric medicine education in Europe Geriatrics from the European Union Perspective India Day Hospitals Perspectives from Latin America The title will be indispensable for all those involved in the treatment of older patients Gerontologists to keep up to date with the latest developments in the field General practitioners and specialists in health policy and community care who increasingly have to deal with a significant number of older people Academic researchers in geriatric medicine who are in need for an all encompassing reference work Medical registrars UK residents US in order to pass their exams Teachers of Geriatric Medicine Hospital libraries with increasing budgets to spend on much needed resource in this growing field Professionals within the pharmaceutical industry in order to monitor treatment options and new prescription developments

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Acute Coronary Syndromes: A Companion to Braunwald's Heart Disease E-Book Pierre Theroux,2010-09-24 Acute Coronary Syndromes a Companion to Braunwald s Heart Disease covers the state of the art scientific and clinical information you need to rapidly evaluate and

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