

NOVARTIS AG
Form 6-K
September 02, 2014

SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 20549

FORM 6-K

**REPORT OF FOREIGN PRIVATE ISSUER
PURSUANT TO RULE 13a-16 or 15d-16 OF
THE SECURITIES EXCHANGE ACT OF 1934**

Report on Form 6-K dated August 30, 2014

(Commission File No. 1-15024)

Novartis AG

(Name of Registrant)

Lichtstrasse 35

4056 Basel

Switzerland

(Address of Principal Executive Offices)

Indicate by check mark whether the registrant files or will file annual reports under cover of Form 20-F or Form 40-F:

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Novartis International AG

Novartis Global Communications

CH-4002 Basel

Switzerland

<http://www.novartis.com>

MEDIA RELEASE • COMMUNIQUE AUX MEDIAS • MEDIENMITTEILUNG

Novartis new heart failure medicine LCZ696 cut cardiovascular deaths by 20% vs. ACE-inhibitor in landmark PARADIGM-HF trial

- *Study showed significantly more HF-REF patients on LCZ696 regimen were alive, had fewer hospitalizations than those given enalapril regimen(1)*
- *On all-cause mortality, LCZ696 doubled the effect that enalapril, an ACE-inhibitor, previously showed vs placebo when added to current best treatment for HF-REF(1),(2)*
- *26 million people across US and Europe live with heart failure(3), facing high risk of death and poor quality of life(4),(5)*

Basel, August 30, 2014 Today at the European Society of Cardiology congress and published simultaneously in the New England Journal of Medicine, Novartis revealed that its investigational heart failure medicine, LCZ696, was superior to ACE-inhibitor enalapril on key endpoints in the largest heart failure study ever done(1),(6). In PARADIGM-HF patients with heart failure with reduced ejection fraction (HF-REF) who were given LCZ696 were more likely to be alive and less likely to have been hospitalized for sudden deterioration of their heart failure than those given ACE-inhibitor enalapril(1). Patients received LCZ696 or enalapril on top of current best treatment.

The magnitude of benefit with LCZ696 against enalapril in HF-REF patients was highly statistically significant and clinically important. In the study, the benefit of LCZ696 was seen early, was sustained and was consistent across subgroups. LCZ696: (1)

- reduced the risk of death from cardiovascular causes by 20% (p=0.00004)
- reduced heart failure hospitalizations by 21% (p=0.00004)
- reduced the risk of all-cause mortality by 16% (p=0.0005)

Overall there was a 20% risk reduction on the primary endpoint, a composite measure of CV death or heart failure hospitalization ($p=0.0000002$) (1).

By demonstrating a very significant reduction in cardiovascular deaths while improving Quality of Life, Novartis' new heart failure medicine, LCZ696, represents one of the most important cardiology advances of the last decade," said David Epstein, Division Head, Novartis Pharmaceuticals. "We want to thank leading cardiologists from around the world for their collaboration with us and their determination in advancing this important new life saving therapy for heart failure patients."

LCZ696, a twice a day tablet being investigated for heart failure, has a unique mode of action which is thought to reduce the strain on the failing heart(6),(7),(8). It acts to enhance the protective neurohormonal systems of the heart (NP system) while simultaneously suppressing the harmful system (the RAAS)(6),(8). Currently available medicines for HF-REF work only to block the detrimental effects(6),(8). Despite existing therapies, the mortality rate remains very high with up to 50% of patients dying within 5 years of a diagnosis of heart failure(9),(10),(11). Approximately half of patients with heart failure have HF-REF(12).

Analysis of the safety data from PARADIGM-HF showed side effects were manageable in the study(1). Fewer patients on LCZ696 discontinued study medication for any adverse event compared to those on enalapril (10.7% vs 12.3%, respectively, $p=0.03$). The LCZ696 group had more hypotension and non-serious angioedema but less renal impairment, hyperkalemia and cough than the enalapril group.

Novartis plans to file the application for marketing authorization with the US FDA by the end of 2014 and in the EU in early 2015.

About the PARADIGM-HF study

PARADIGM-HF is a randomized, double-blind, phase III study evaluating the efficacy and safety profile of LCZ696 versus enalapril (a widely studied ACE inhibitor) in 8,442 patients with HF-REF(6),(13). The baseline characteristics showed the patients enrolled were typical HF-REF patients with NYHA Class II-IV heart failure. PARADIGM-HF was specifically designed to see if LCZ696 could decrease CV mortality by at least 15% vs. enalapril(6). Patients received LCZ696 or enalapril in addition to current best treatment regimen. The primary endpoint is a composite of time to first occurrence of either cardiovascular death or heart failure hospitalization, and is the largest heart failure study ever done(6).

Secondary endpoints are change in the clinical summary score for heart failure symptoms and physical limitations (as assessed by Kansas City Cardiomyopathy Questionnaire) at 8 months; time to all-cause mortality; time to new onset atrial fibrillation; and time to occurrence of renal dysfunction(6). It was initiated in December 2009 and in March 2014 the Data Monitoring Committee confirmed that patients given LCZ696 were significantly less likely to die from CV causes, leading to the trial being stopped early(14). The DMC also confirmed the primary endpoint had been met.

About LCZ696 in heart failure

LCZ696 is an ARNI (Angiotensin Receptor Neprilysin Inhibitor) and has a unique mode of action which is thought to reduce the strain on the failing heart(6),(8). It harnesses the body's natural defences against heart failure simultaneously acting to enhance the levels of natriuretic and other endogenous vasoactive peptides, while also inhibiting the renin-angiotensin-aldosterone system (RAAS).

Heart failure is a debilitating and potentially life-threatening disease in which the heart cannot pump enough blood around the body. Symptoms such as breathlessness, fatigue and fluid retention can appear slowly and worsen over time, significantly impacting quality of life(4),(15).

It is a significant and growing public health concern with a high unmet need for new treatments. Every year, HF costs the world economy \$108 billion(16), and hospitalizations comprise 60-70% of treatment costs(17),(18).

Disclaimer

The foregoing release contains forward-looking statements that can be identified by words such as being investigated, thought, plans, growing, or similar terms, or by express or implied discussions regarding potential marketing approvals for LCZ696, or regarding potential future revenues from LCZ696. You should not place undue reliance on these statements. Such forward-looking statements are based on the current beliefs and expectations of management regarding future events, and are subject to significant known and unknown risks and uncertainties. Should one or more of these risks or uncertainties materialize, or should underlying assumptions prove incorrect, actual results may vary materially from those set forth in the forward-looking statements. There can be no guarantee that LCZ696 will be approved for sale in any market, or submitted for approval in any additional markets, or at any particular time. Neither can there be any guarantee that LCZ696 will be submitted or approved for any additional indications or labeling in any market, or at any particular time. Nor can there be any guarantee that

LCZ696 will be commercially successful in the future. In particular, management's expectations regarding LCZ696 could be affected by, among other things, the uncertainties inherent in research and development, including unexpected clinical trial results and additional analysis of existing clinical data; unexpected regulatory actions or delays or government regulation generally; the company's ability to obtain or maintain proprietary intellectual property protection; general economic and industry conditions; global trends toward health care cost containment, including ongoing pricing pressures; unexpected manufacturing issues, and other risks and factors referred to in Novartis AG's current Form 20-F on file with the US Securities and Exchange Commission. Novartis is providing the information in this press release as of this date and does not undertake any obligation to update any forward-looking statements contained in this press release as a result of new information, future events or otherwise.

About Novartis

Novartis provides innovative healthcare solutions that address the evolving needs of patients and societies. Headquartered in Basel, Switzerland, Novartis offers a diversified portfolio to best meet these needs: innovative medicines, eye care, cost-saving generic pharmaceuticals, preventive vaccines, over-the-counter and animal health products. Novartis is the only global company with leading positions in these areas. In 2013, the Group achieved net sales of USD 57.9 billion, while R&D throughout the Group amounted to approximately USD 9.9 billion (USD 9.6 billion excluding impairment and amortization charges). Novartis Group companies employ approximately 135,000 full-time-equivalent associates and sell products in more than 150 countries around the world. For more information, please visit <http://www.novartis.com>.

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References:

- (1) McMurray JJV et al. Angiotensin-Neprilysin Inhibition versus Enalapril in Heart Failure, *N Engl J Med* 2014; DOI: 10.1056/NEJMoa1409077
- (2) Effect of ACE inhibitor vs placebo derived from SOLVD-Treatment trial, that showed enalapril reduced risk of death by 16% vs placebo; Effect of LCZ696 vs ACE inhibitor derived from PARADIGM-HF trial.
- (3) Ambrosy AP et al. *J Am Coll Cardiol* 2014;63:1123-33. doi: 10.1016/j.jacc.2013.11.053
- (4) Gheorghiade M, Pang P, Acute heart failure syndromes, *Journal of the American College of Cardiology* 2009; 53 (7):557-73
- (5) Harrison's Principles of Internal Medicine, Seventeenth Edition pages 1442 - 1455
- (6) McMurray JJ, Packer M, Desai AS, et al. Dual angiotensin receptor and neprilysin inhibition as an alternative to angiotensin-converting enzyme inhibition in patients with chronic systolic heart failure: rationale for and design of the Prospective comparison of ARNI with ACEI to Determine Impact on Global Mortality and morbidity in Heart Failure trial (PARADIGM-HF). *Eur J Heart Fail* 2013;15:1062-1073 (doi:10.1093/eurjhf/hft052)
- (7) Langenickel TH et al. Angiotensin receptor-neprilysin inhibition with LCZ696: a novel approach for the treatment of heart failure. *Drug Discovery Today: Therapeutic Strategies*.2012, Vol 9. No.4
- (8) Solomon SD et al. The angiotensin receptor neprilysin inhibitor LCZ696 in heart failure with preserved ejection fraction: a phase 2 double-blind randomised controlled trial. *Lancet*. 2012;380:1387-95

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- (9) Lloyd-Jones *et al.* Heart disease and stroke statistics 2010 update: a report from the American Heart Association. *Circulation*. 2010;121:e46-215
- (10) Zannad F. *et al.*, Heart failure burden and therapy, *Europace* 2009, 11; v1-v9.
- (11) Loefer LR, Rosamond WD, Chang PP, Folsom AR, Chambless LE. Heart failure incidence and survival (from the Atherosclerosis Risk in Communities study). *Am J Cardiol*. 2008;101(7):1016
- (12) Doughty R.N., The survival of patients with heart failure with preserved or reduced left ventricular ejection fraction: an individual patient data meta-analysis, *European Heart Journal* (2012);33(14):1750-1757
- (13) Clinicaltrials.gov, NCT01035255 last accessed online 19 March 2014
- (14) Novartis press release PARADIGM-HF trial of Novartis LCZ696 for chronic heart failure stopped early based on strength of interim results issued on March 31, 2014
- (15) Mosterd A, Hoes, A, Clinical epidemiology of heart failure, *Heart* 2007;93:1137
- (16) Cook C, Cole G, Asaria P, Jabbour R, Francis DP. The annual global economic burden of heart failure. *Int J Cardiol*. 2014.;171(3):368-76
- (17) Neumann T, Biermann J, Erbel R, Neumann A, Wasem J, Ertl G, et al., Heart failure: the commonest reason for hospital admission in Germany: medical and economic perspectives. *Dtsch Arztebl Int*. 2009;106:269-75.

(18) Stewart S, Jenkins A, Buchan S, McGuire A, Capewell S, McMurray JJ. The current cost of heart failure to the National Health Service in the UK. Eur J Heart Fail. 2002 Jun;4(3):36-71.

Novartis Media Relations

Central media line : +41 61 324 2200

Eric Althoff

Novartis Global Media Relations

+41 61 324 7999 (direct)

+41 79 593 4202 (mobile)

eric.althoff@novartis.com

Dermot Doherty

Novartis Global Media Relations

+41 61 696 8653 (direct)

+41 79 536 9755 (mobile)

dermot.doherty@novartis.com

e-mail: media.relations@novartis.com

For Novartis multimedia content, please visit www.thenewsmarket.com/Novartis

For questions about the site or required registration, please contact: journalisthelp@thenewsmarket.com.

Novartis Investor Relations

Central phone:

Samir Shah

Pierre-Michel Bringer

Thomas Hungerbuehler

Isabella Zinck

+41 61 324 7944

+41 61 324 7944

+41 61 324 1065

+41 61 324 8425

+41 61 324 7188

North America:

Stephen Rubino

Susan Donofrio

+1 862 778 8301

+1 862 778 9257

e-mail: investor.relations@novartis.com

e-mail: investor.relations@novartis.com

SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned, thereunto duly authorized.

Novartis AG

Date: August 30, 2014

By: /s/ MALCOLM B. CHEETHAM

Name: Malcolm B. Cheetham
Title: Head Group Financial
Reporting and Accounting