CLINICAL TRIAL FACT SHEET

Clinicaltrials.gov Identifier: NCT03275285

A phase III, randomized, open-label, multicenter study assessing the clinical benefit of isatuximab combined with carfilzomib (Kyprolis®) and dexamethasone versus carfilzomib with dexamethasone in patients with relapsed and/or refractory multiple myeloma previously treated with 1 to 3 prior lines of therapy

(IKEMA study)

Trial Description:

Approximately 300 patients who have already been treated for myeloma with from one to three prior therapy regimens will be randomly assigned by a computer to one of two groups called study "arms." Patients will know to which arm they have been assigned. Patients in the experimental arm will receive the combination of anti-CD38 monoclonal antibody isatuximab (formerly known as SAR650984) plus carfilzomib (brand name Kyprolis®) and dexamethasone (IKd), and patients in the other arm will receive carfilzomib plus dexamethasone (Kd).

Trial Objectives:

The primary objective of this trial is to determine the possible benefit of isatuximab in combination with carfilzomib and dexamethasone in prolonging the length of response time (progression-free survival, or PFS) as compared to carfilzomib and dexamethasone. Other objectives include evaluating the following:

- Overall response rate (ORR).
- Rate of very good partial response (VGPR) or better (IMWG criteria) with minimal residual disease (MRD) negativity in both arms.
- Rate of complete response (CR) in both study arms (IMWG criteria).
- Rate of overall survival (OS) in both arms.
- Safety of therapy in both study arms.
- Duration of response in both study arms.
- How isatuximab in combination with carfilzomib is metabolized by the body.
- Quality of life, disease and treatment-related symptoms, impact on the healthcare system, and health status of all trial participants.

Trial Design:

Experimental Arm

- Patients will receive 4-week cycles of isatuximab intravenously (IV, or into a vein) on days 1, 8, 15, and 22 of the 1st cycle, then on day 1 and 15 of subsequent cycles along with carfilzomib.
- Patients will receive carfilzomib intravenously on days 1, 2, 8, 9, 15, and 16 of each 4-week cycle.
- Patients will receive dexamethasone intravenously or by mouth twice a week during each 4-week cycle.

Comparator Arm

- Patients will receive intravenous carfilzomib on days 1, 2, 8, 9, 15, and 16 of each 28-day cycle.
- Patients will receive intravenous or oral dexamethasone twice weekly.
- Patients will receive oral or IV dexamethasone on days 1, 8, 15, and 22 of each 4-week cycle.

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INTERNATIONAL MYELOMA FOUNDATION

International Myeloma Foundation
12650 Riverside Drive, Suite 206, North Hollywood, CA 91607 USA
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(continued)

Duration of Treatment:

- The screening period for this study is up to 3 weeks long.
- Patients will continue study treatment until disease progression, unacceptable adverse reaction (side effect), patient's wish, or other reason for discontinuation.
- Patients who discontinue the study treatment due to disease progression will be followed every 3 months for further anti-myeloma therapies, progression-free survival (remission) after further therapy, and for survival.
- Patients who discontinue the study treatment before disease progression is documented will be followed-up every 4 weeks until confirmation of disease progression, then every 3 months for further anti-myeloma therapies, progression-free survival (remission duration) after further therapy, and survival.
- After progression-free survival (remission duration) analysis, patients will be followed yearly for 3 years.

Inclusion Criteria:

- Myeloma previously treated with 1–3 lines of therapy.
- Measurable M-protein (≥ 0.5 g/dL) and/or urine M-protein (≥ 200 mg/24 hours).

Exclusion Criteria:

- Previous treatment with carfilzomib and failure to achieve at least one minor response (≥ 25% drop in monoclonal protein level) during previous therapies and/or last previous therapy completed within last 14 days.
- Disease measurable only by free light chain level.
- Poor overall health status.
- Patients with a history of the following heart conditions: myocardial infarction, severe/unstable angina pectoris, coronary/peripheral artery bypass graft, New York Heart Association class III or IV congestive heart failure, ≥ grade 3 arrhythmias, stroke or transient ischemic attack within the last 6 months, and/or left ventricular ejection fraction lower than 40%.
- Previous cancer unless disease-free for > 5 years, or cancer that didn't spread to other tissue and was curatively treated.
- Known acquired immunodeficiency syndrome-related illness or illness requiring antiretroviral treatment, or hepatitis A, B, or C active infection.
- Females of childbearing potential, or male patients with female partners who are of childbearing potential, who do not agree to use a highly effective method of birth control.

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(continued)

Locations Enrolling Patients and Contact Information:

For site information, send an email including one or more of the below site numbers to **Contact-Us@sanofi.com**

United States, California
Investigational Site Number 8400002
San Francisco, California, United States, 94117

United States, South Carolina
Investigational Site Number 8400003
Spartanburg, South Carolina, United States, 29303-3040

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