KURZPROTOKOLL
CAMN107YDE19

Öffentlicher Titel	Phase Ib Studie zur kombinierten Therapie mit Nilotinib und Ruxolitinib bei CML oder PH+ALL Patienten
Wissenschaftl. Titel	A Phase Ib single-arm, open-label, multicenter study to assess the safety and tolerability of combined treatment with nilotinib 300mg BID and ruxolitinib increasing dose in CML patients in CP and in AP/BC or relapsed/refractory Ph+ ALL
Kurztitel	CAMN107YDE19
Studienart	multizentrisch, prospektiv, offen/unverblindet, einarmig, Pharma-Studie
Studienphase	Phase I
Erkrankung	Blut: Myeloische Neoplasien/Dysplasien: Chronische myeloische Leukämie (CML) Blut: Akute lymphatische Leukämie (ALL): Rezidiviert/refraktär
Ziele	- Occurrence of dose limiting toxicities (DLTs) [Time Frame: Baseline, up to day 28 (equals first cycle)] [Designated as safety issue: Yes] Occurrence of DLTs during cycle 1 - Safety and tolerability profile of nilotinib and ruxolitinib administered in combination [Time Frame: Baseline, up to month 12] [Designated as safety issue: Yes] Maximum Tolerated Dose (MTD) and/or Recommended Phase II Dose (RPIID) of ruxolitinib in combination with nilotinib. (timeframe, baseline up to month 12) - Trough levels of nilotinib and ruxolitinib administered in combination [Time Frame: Baseline, up to month 12] [Designated as safety issue: Yes] Trough levels will be determined by measuring the minimum plasma concentration (Cmin). - Clinical activity of nilotinib and ruxolitinib administered in combination [Time Frame: Baseline and at 3, 6, and 12 months] [Designated as safety issue: No] Chronic myeloid leukemia in chronic phase: assessment of molecular response: MMR (<=0.1% BCR-ABL) and MR4 (<=0.001% BCR-ABL) at 3, 6, 12 months; Advanced disease: assessment of cytogenetic response will be based on evaluating percentage of Ph+ metaphases
Einschlusskriterien	- Patients of the first stratum must have chronic myeloid leukemia receiving nilotinib first-line therapy or receiving second-line or subsequent-line treatment with nilotinib. - Patients of the second stratum must have CML in AP/BC or relapsed/refractory Ph+ ALL, or be Ph+ ALL patients with MRD with or without prior nilotinib pretreatment; - Patients must have adequate end organ function, as defined by: - Creatinine < 2.0 x upper limit of normal (ULN) - Total bilirubin < 1.5 x ULN (< 3.0 x ULN if related to disease or polymorphism, such as Mb. Gilbert) - ALT and AST < 2.5 x ULN (< 5.0 x ULN if related to disease) - Serum lipase <= 1.5 x ULN - Alkaline phosphatase <= 2.5 x ULN (< 5.0 x ULN if related to disease) - Patients must have the following electrolyte values within normal limits or corrected to within normal limits with supplements prior to the first dose of study medication: Potassium, Magnesium, Phosphate, Total calcium (corrected for serum albumin) - Female patients of childbearing potential (WOCBP) must have a negative serum pregnancy test within 7 days before initiation of study drug. All WOCBP must use highly effective contraceptive methods throughout and during 3 months after study; - Patient has an Eastern Cooperative Oncology Group (ECOG) performance status of 1 for patients in CP, <= 2 for patients in AP/BC or with relapsed/refractory Ph+ ALL or with Ph+ ALL with MRD; - Patient has the following laboratory values within 7 days of starting study drug: - For CML and Ph+ ALL patients: platelet count > 75 x 109/L and ANC > 1.0 x 109/L
Ausschlusskriterien	- Patient must not have evidence of active malignancy other than the existing CML or ALL

	- Patient must not receive drugs that interfere with coagulation or inhibits platelet function, with the exception of aspirin <= 150 mg per day or low molecular weight heparin. - Patient must not have history of platelet dysfunction, bleeding diathesis, and/or coagulopathy in the 6 months prior to screening; - Patient must not require treatment with any strong CYP3A4 inducer or inhibitor - Patient must not have history of hypersensitivity to any of the study drugs or to drugs of similar chemical classes and their excipients; - Patients must not take other investigational drugs within 28 days prior to screening; - Patient must not be pregnant or lactating at screening and/or baseline; - Patient must not have impaired cardiac functions - Other protocol-defined inclusion/exclusion criteria may apply
Alter	18 Jahre und älter
Molekularer Marker	BCR-ABL1
Fallzahl	80
Sponsor	Novartis Pharma
Förderer	Novartis Pharma
Registrierung in anderen Studienregistern	EudraCT 2014-000831-18 ClinicalTrials.gov NCT02253277