KURZPROTOKOLL
Neobil AIO-HEP-0120

Öffentlicher Titel	Phase II Studie zu Bintrafusp Alfa bei resezierbarem Gallengangskarzinom
Wissenschaftl. Titel	Neoadjuvant Bintrafusp Alfa in Patients With Resectable Biliary Tract Cancer (NEOBIL)
Kurztitel	Neobil AIO-HEP-0120
Studienart	multizentrisch, prospektiv, Therapiestudie, offen/unverblindet, einarmig, Investigator Initiated Trial (IIT)
Studienphase	Phase II
Erkrankung	Verdauung: Gallengangs-/Gallenblasenkrebs (maligne biliäre Tumoren): neoadjuvant

Einschlusskriterien

- Written informed consent granted prior to initiation of any study-specific screening procedures
- Biliary tract cancer, confirmed by histopathology, cytopathology is not sufficient
- Resectable disease limited to the liver assessed by an interdisciplinary tumor board involving a hepatobiliary surgeon
- Subject is willing and able to comply with the protocol for the duration of the study including undergoing treatment and scheduled visits and examinations including follow up
- Age >= 18 years
- Performance status ECOG 0-1
- Normal organ and bone marrow function defined as: o Hematopoetic: absolute neutrophil count >=1,500/mm3, platelet count >= 100,000/mm3, o Hemoglobin >=9 g/dL o Normal international normalized ratio (INR), PT <= 1.5 x ULN and activated partial thromboplastin time (aPTT) <= 1.5 x ULN o Hepatic: AST <=5 x ULN, ALT <= 5 x ULN, and bilirubin <= 3.0 x ULN o Renal: Creatinine level <=1.5 x ULN or estimated creatinine clearance >= 30 mL/min according to the Cockcroft-Gault formula (or local institutional standard method)
- Special medical conditions and comorbidities: o Maximum Child Pugh stage A in patients with cirrhosis o HIV: stable on ART for at least 4 weeks, no documented evidence of multi-drug resistance, viral load of < 400 copies/mL and CD4+ T-cells <= 350 cells/µL o HBV infection: participant on a stable dose of antiviral therapy, HBV viral load below the limit of quantification.
- Women of childbearing potential must have a negative serum or highly sensitive urine pregnancy test performed within 7 days prior to the first dose of IMP
- Women of childbearing potential (WOCBP) must use HIGHLY EFFECTIVE method(s) of contraception to avoid pregnancy for the duration of study treatment and further 2 months after the last dose of IMP

Ausschlusskriterien

- Metastatic disease
- Prior surgery, systemic therapy, radiation therapy, chemoradiation, transarterial chemoembolisation (TACE), Radiofrequency ablation (RFA) or selective intraarterial Radiotherapy (SIRT) for treatment of CCA. NOTE: Laparoscopy for diagnostic procedures is allowed
- Drug or alcohol addiction, medical or psychological condition that may interfere with the patient´s participation in the study
- Participation in another clinical trial with any investigational study drug (whatever the use, curative, prophylactic or diagnostic intent) within 30 days prior to enrollment
- Pregnancy or breast feeding women
- Regulatory and ethical criteria: Patient who has been incarcerated or involuntarily institutionalized by court order or by the authorities [§ 40 Abs. 1 S. 3 Nr. 4 AMG]. Patients who are unable to consent because they do not understand the nature, significance and implications of the clinical trial and therefore cannot form a rational intention in the light of the facts [§ 40 Abs. 1 S. 3 Nr. 3a AMG]

- IMMUNOSUPRESSANTS: "Current use of immunosuppressive medication, EXCEPT for the following: a. intranasal, inhaled, topical steroids, or local steroid injection (e.g., intra-articular injection); b. Systemic corticosteroids at physiologic doses <= 10 mg/day of prednisone or equivalent; c. Steroids as premedication for hypersensitivity reactions (e.g., CT scan premedication)."
- AUTOIMMUNE DISEASE: "Active autoimmune disease that might deteriorate when receiving an immuno-stimulatory agent. Patients with diabetes type I, vitiligo, psoriasis, or hypo- or hyperthyroid diseases not requiring immunosuppressive treatment are eligible."
- PREVIOUS MALIGNANT DISEASE: within the last 3 years except for a. superficial/non-invasive bladder cancer, or basal or squamous cell carcinoma in situ treated with curative intent; b. endoscopically resected GI cancers limited to the mucosal layer without recurrence in > 1 year
- INFECTIONS: "Active infection requiring systemic therapy. "
- VACCINATION: has received or will receive a live vaccine within 30 days prior to the first administration of study intervention. Seasonal flu vaccines that do not contain a live virus are permitted
- HYPERSENSITIIVTY TO BINTRAFUSP ALFA: "Known severe hypersensitivity [Grade >= 3 NCI CTCAE 5.0]) to investigational product or any component in its formulations, any history of anaphylaxis, or recent, within 5 months, history of uncontrollable asthma
- CARDIOVASCULAR DISEASE: "Clinically significant (i.e., active) cardiovascular disease: cerebral vascular accident/stroke (< 6 months prior to enrollment), myocardial infarction (< 6 months prior to enrollment), unstable angina, congestive heart failure (>= New York Heart Association Classification Class II), or serious cardiac arrhythmia requiring medication."
- BLEEDING: "history of bleeding diathesis or recent major bleeding events (i.e. Grade >= 2 bleeding events in the month prior treatment)
- Other severe acute or chronic medical conditions: "including drug-induced interstitial lung disease (ILD) or participant has had a history of drug-induced pneumonitis that has required oral or IV steroids", and/or other diseases, which in the opinion of the Investigator might impair the participant's tolerance for the study or ability to consistently participate in study procedures

Alter	18 Jahre und älter
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