

From Buruli anti-mycobacterial treatment to Ulcer wound healing - A prospective evaluation of clinical and laboratory parameters to distinguish mycobacterial disease, paradoxical reaction and secondary bacterial infection for targeted treatment (extension)

Initiative: Wissen für morgen – Kooperative Forschungsvorhaben im subsaharischen Afrika (beendet)

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In the first phase we have successfully explored the use of an innovative heat application system to treat Mycobacterium ulcerans disease in a proof-of-principle study, confirmed the efficacy of heat treatment at the primary endpoint (six months after heat treatment) in a large GCP-conform trial, and studied the treatmentinduced microbiological, histopathological and immunological evolution of BU lesions of relevance for the understanding of wound healing processes and clinical BU treatment management. We found strikingly heterogeneous treatment responses of BU lesions. Based on our standardized clinical observation and documentation and extensive laboratory work-up of the patients enrolled in the heat treatment trial we can now understand the determinants of heterogeneous treatment responses of BU lesions better. Disturbance of wound healing may be related to: (1) M.ulcerans activity, (2) M.ulcerans-host interactions (so called paradoxical reaction) and (3) secondary bacterial infection. The investigation of these often co-existing processes contributes to the clarification of unresolved principle questions such as the nature of immune protection against M.ulcerans disease after successful treatment and the resolution of the mycolactoneinduced local immunosuppression during therapy. The project now aims to improve our ability to define clinical, microbiological, cell biological and immunological features that influence progression of BU lesions to wound healing and tissue repair. To achieve this, we will analyze in-depth the large clinical and laboratory dataset of our clinical trial and prospectively study markers of wound healing and their ability to predict clinical outcome in a cohort of BU patients in Ghana. In addition, we will complete the heat treatment trial with the assessment of the relapse rate at month 24 after completion of heat treatment as well as the secondary endpoints and establish a modular reference data base for BU patients.

### Projektbeteiligte

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