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2 **REVIEW**

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SEVERE TRAUMATIC BRAIN INJURY MANAGEMENT AND CLINICAL OUTCOME USING THE LUND CONCEPT

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- Abstract—This review covers the main principles of the 13 O3 Lund concept for treatment of severe traumatic brain injury. This is followed by a description of results of clinical studies in which this therapy or a modified version of the therapy has been used. Unlike other guidelines, which are based on meta-analytical approaches, important components of the Lund concept are based on physiological mechanisms for regulation of brain volume and brain perfusion and to reduce transcapillary plasma leakage and the need for plasma volume expanders. There have been eight non-randomized and two randomized outcome studies with the Lund concept or modified versions of the concept. The non-randomized studies indicated that the Lund concept is beneficial for outcome. The two randomized studies were small but showed better outcome in the groups of patients treated according to the modified principles of the Lund concept than in the groups given a more conventional treatment

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Key words: severe traumatic brain injury, Lund concept, management, outcome.

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 E-mail address: Lars-Owe.Koskinen@neuro.umu.se (L.-O.D. Koskinen).
- Q4 Abbreviations: ARDS, acute respiratory distress syndrome; CPP, cerebral perfusion pressure; Hb, hemoglobin; ICP, intracranial pressure; LC, Lund concept; MAP, mean arterial pressure; PEEP, positive endexpiratory pressure; sTBI, severe traumatic brain injury; SAH, subarachnoid hemorrhage; TER, transcapillary exchange rate.

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INTRODUCTION

Originally, the Lund concept (LC) for treatment of severe Q5 41 traumatic brain injury (sTBI) was a theoretical approach 42 mainly based on the physiological and 43 pathophysiological principles of brain volume and brain 44 perfusion regulation (Asgeirsson et al., 1994; Grände 45 et al., 1997; Grände, 2006). The concept aimed at coun-46 teracting an increase in intracranial pressure (ICP) or to 47 reduce an already raised ICP after sTBI, while improving 48 compromised perfusion in and around the contusion 49 areas at the same time. It can be described as an ICP-50 and perfusion-quided approach. The main components 51 of the LC have found support in experimental and clinical 52 studies, as described later in this review. 53

So far, no TBI guidelines have been tested in a large randomized clinical trial and from that point of view there is limited high-level clinical evidence for all TBI guidelines presented today (Muzevic and Splavski, 2013). A specific therapy therefore must be based on other types of input such as smaller clinical outcome studies including metaanalysis, experimental studies and basal physiological principles.

Even though different guidelines differ in essential62aspects, the Brain Trauma Foundation's guidelines have63moved closer to the LC during the past 10 years, e.g.64concerning cerebral perfusion pressure (CPP) and the65use of vasopressors (Bullock et al., 1996, 2000; Brain66Trauma Foundation, 2007). In contrast to Brain Trauma67

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Foundation guidelines-in which the ICP-reducing ther-68 apy should start when ICP is above 20 mmHg (Brain 69 Trauma Foundation, 2007)-the LC recommends that 70 the therapy should start as early as possible after arrival 71 at the hospital, in an attempt to counteract the develop-72 ment of brain edema and to ensure that there is early opti-73 mization of the perfusion. To our knowledge, no clear side 74 75 effects have appeared with the LC, which means that it can be given early and to all patients independent of 76 severity of the injury and independent of the degree of 77 autoregulation. The LC has not changed since its intro-78 duction, except that dihydroergotamin is no longer used. 79 Dihydroergotamin, which reduces ICP via cerebral 80 81 venous constriction, was used in the initial version of the concept in patients with uncontrolled increase in ICP 82 (Asgeirsson et al., 1994). It was withdrawn because of 83 possible side effects related to peripheral vasoconstriction 84 in high doses. For details of the LC guidelines, see; 85

Asgeirsson et al. (1994), Grände (2006, 2011) and 86 Olivecrona et al. (2007, 2009a,b). A simplified schematic Q687 algorithm of the LC used in the clinical setting is shown in 88 Fig. 1. 89

MEASUREMENT OF ICP AND CPP

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Like in other guidelines, monitoring of ICP is an essential part of the LC, and the monitoring should be started as soon as possible after the arrival at the hospital. The method of ICP monitoring can either be by external ventricular drainage or by an intraparenchymal device. It is also crucial to monitor the arterial pressure and the mean arterial pressure (MAP). 91

The reference points for MAP and ICP must be identical when calculating CPP. For example, a head elevation of 15 degrees with the zero-reference point for the ICP at the external meatus and the zero-reference 101

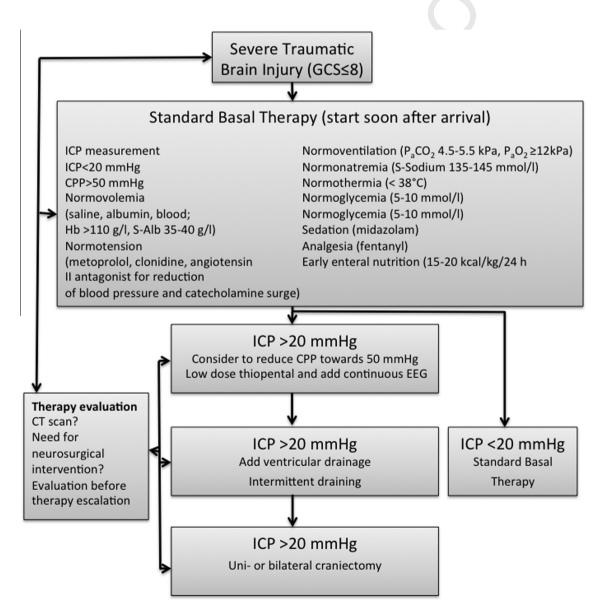


Fig. 1. A simplified schematic algorithm of the LC used in the clinical setting. For doses of the pharmacological substances used and other parameters, see Grände (2006).

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