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Effect of high-dose methotrexate chemotherapy on intestinal flora and mucosal barrier in children with acute lymphoblastic leukemia

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Background: High-dose methotrexate (HDMTX) chemotherapy is generally accepted as an effective method for the treatment and prevention of extramedullary leukemia in children. However, it is unknown whether they are the damaged effects of HDMTX chemotherapy on intestinal mucosal barrier, which may in turn influence the progress or prognosis of leukemia. The levels of diamine oxidase (DAO) and endotoxin were measured to be able to evaluate the damaged intestinal mucosal barrier. The aim of this study was to study the effects of high-dose methotrexate (HDMTX) treatment on intestinal mucosal barrier in children with acute lymphocytic leukemia (ALL).

Objective: High-dose methotrexate (HDMTX) chemotherapy is generally accepted as an effective method for the treatment and prevention of extramedullary leukemia in children. However, it is unknown whether HDMTX chemotherapy kills intestinal bacteria and affects intestinal mucosal barrier on a large scale, thus causing dysbacteriosis and mucosal disfunction, which may in turn influence the progress or prognosis of leukemia. The aim of this study was to examine changes in intestinal flora and intestinal mucosal barrier in children with acute lymphoblastic leukemia (ALL) treated with HDMTX chemotherapy.

Methods: Bacterial DNA in stool from 36 healthy children and 36 ALL children were extracted and tested at A260 with a spectrophotometer before and after HDMTX chemotherapy. The primers of *Bifidobacteria*, *Lactobacillus and Escherichia coli* were designed according to the 16SrRNA/DNA bacterial sequences. Bacteria were qualitatively and quantitatively confirmed by routine polymerase chain reaction (PCR) and fluorescent quantitative PCR, respectively. The blood samples were collected from 30 normal children and from 30 children with ALL after 1 h, 24 h, 44 h and 68 h of intravenous drip HDMTX. The levels of diamine oxidase (DAO) and endotoxin were measured by the spectrophotometer in the isolated plasma.

Results: Our data showed that the total amount of bacterial DNA in the stools of children with ALL was decreased by 29.6% compared with healthy children. The total amount of flora in the stools of children with ALL on the third and seventh days after chemotherapy were 1496.5+577.1 and 1966.6+598.3 ng/mL, respectively, which was notably less than before chemotherapy (2436.3+768.6 ng/mL) and healthy children (3479.3±870.5 ng/mL). The amount of *Bifidobacteria*, *Lactobacillus and E. coli* in the intestinal tract in the ALL group after chemotherapy had an apparent change, which decreased most clearly on the third day, and partially recovered on the seventh day after chemotherapy. The measured levels of plasma endotoxin and DAO in ALL group were significantly higher than in normal control group, the difference was statistically significant (P<0.01). The level of plasma endotoxin and DAO after 24 h and 44 h were both higher than 1 h and 68 h IV drip HDMTX, and reach the peak after 24 h, the difference was statistically significant (P<0.01).

Conclusions: The amount of bacterial DNA, *Bifidobacteria, Lactobacillus and E. coli* in the stools was in decrease significantly compared with the control group. The levels of plasma endotoxin and DAO in ALL group were apparently higher than in normal control group. Leukemia itself and HDMTX chemotherapy can all cause intestinal dysbacteriosis and intestinal mucosal barrier disfunction in children with ALL.

Keywords: Acute lymphoblastic leukemia, high-dose methotrexate, intestinal flora, dysbacteriosis, mucosal barrier, and disfunction.

Prosthodontic rehabilitation of a total glossectomy patient with magnetic detachable mandibular tongue prosthesis: A clinical report

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Total glossectomy can result in significant functional impairments in mastication, swallowing and speech. In addition to these functional problems, severe psychological problems may follow complete loss of the tongue. Placement of a mandibular tongue prosthesis obturates this large defect, increases the patient's ability to produce intelligible sounds and assists with return to a normal diet. This prosthetic rehabilitation can also improve both appearance and psychosocial adjustment. This clinical report describes the treatment of a total glossectomy patient using magnetically attached two-piece tongue prosthesis. Unconventionally, magnets were used to mechanically attach the two parts.

Keywords: Prosthodontic rehabilitation, Tongue prosthesis, Mandibular tongue prosthesis, Glossectomy, Total glossectomy.

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