1530. De-implementation Strategy to Reduce the Inappropriate Use of Urinary and Intravenous CATheters: the RICAT Study

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Background. Catheter-associated urinary tract infection (UTI) and catheter-associated bloodstream infection (BSI) are common healthcare-associated infections (HAI). Therefore, catheters should only be used if indicated. However, based on the literature up to 65% of the urinary catheters and 56% of the peripheral intravenous catheters have an inappropriate indication. So, an efficient way to reduce HAIs is to avoid unnecessary use of catheters. Our quality improvement project aims to reduce unnecessary use of catheters.

Methods. In a multicenter, interrupted time series study, several interventions to avoid inappropriate use of catheters were carried out in internal medicine and nonsurgical subspecialty wards in seven hospitals in the Netherlands. The indications for catheter use were based on (inter)national guidelines. The primary endpoint is the percentage of inappropriate indications on the day of data collection. Secondary endpoints are catheter-associated infections, length of hospital stay and mortality. Data were collected once per 2 weeks during baseline (7 months) and post-intervention (7 months). Preliminary analyses compared incidence rates before and after the intervention.

Results. Data were obtained from 5,691 observed patients. The rate of inappropriate use of urinary catheters decreased from 32.1 to 23.7% (incidence rate ratio 0.74, 95% CI 0.58–0.94, *P* = 0.013), and inappropriate use of peripheral intravenous catheters decreased from 22.0 to 15.2% (incidence rate ratio 0.69, 95% CI 0.60–0.80, *P* < 0.001). The overall urinary and intravenous catheter use was stable, resp. 12.2% (*n* = 324) to 12.5% (*n* = 380) and 62.8% (*n* = 1,665) to 62.1% (*n* = 1,887). Most inappropriate indications were due to prolonged catheter use. The indications which expire frequently are "Accurate measurements of urinary output in critically ill patients' for urinary and "IV fluids and antibiotic therapy' for intravenous catheters. Subsequent analyses will take into account the interrupted time series design, and evaluate catheter-associated UTI and BSI rates.

Urinary catheters





Peripheral intravenous catheters

Conclusion. Our de-implementation strategy reduces unnecessary use of urinary and intravenous catheters in non-ICUs. It is important to increase awareness for inappropriate use of catheters.

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1531. A CMV Vaccine Based on Non-Replicating Lymphocytic Choriomeningitis Virus Vectors Expressing gB and pp65 Is Safe and Immunogenic in Healthy Volunteers, Allowing for Development of a Phase II Clinical Trial in Living Donor Kidney Transplant Recipients

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Background. Cytomegalovirus (CMV) is a major pathogen in pregnancy and immunocompromised patients. Antiviral prophylaxis is limited by toxicities, recurrent infection, and antiviral resistance. A safe and protective CMV vaccine is highly desirable.

Methods. HB-101 is a CMV vaccine consisting of two nonreplicating lymphocytic choriomeningitis virus vectors, one expressing the human CMV antigen pp65 and the other a truncated, more antigenic isoform of the CMV fusion protein gB. The safety and immunogenicity of HB-101 were evaluated in a randomized, placebo-controlled, double-blind phase I dose-escalating trial (NCT02798692). Three dosing cohorts (1: 2.6×10^6 ; $2: 2.6 \times 10^7$ and $3: 2.6 \times 10^8$ FFU) of 18 subjects each were enrolled. On Day 0, Month 1, and Month 3, HB-101 or placebo was administered to 14 and 4 subjects, respectively. Immunogenicity studies included cellular responses against pp65, and humoral and cellular responses against gB and the LCMV vector.

Results. Injection site pain was the most frequently reported solicited adverse event (SAE). It affected 57.1% of HB-101 recipients in both cohorts 1 and 2 and 92.9% in cohort 3. Among the general SAE malaise, fatigue and generalized myalgia were most frequently reported. All SAE were generally mild to moderate and lasted <8 days. No serious adverse events and no abnormal lab tests were noted during the active phase of the study. HB-101-induced gB-specific IgG antibody responses at all doses, in a dose-dependent manner. All three dose levels also induced antibodies that neutralized HCMV infection in cultured human fibroblasts (MRC-5 cells), and resulted in a robust, boosterable and durable T-cell response by IFNY ELISPOT for CMV gB and pp65. Polychromatic flow cytometry indicated induction of a high proportion of polyfunctional CMV-specific CD8 and CD4 T-cells. CD8 T-cells expressing IFNy, IL2 and TNFa without CD107a were among the most prominent populations induced against CMV pp65.

Conclusion. HB-101 is a novel CMV vaccine with a good safety profile in healthy volunteers, eliciting strong humoral and cellular immune responses. We are starting a Phase 2 trial in kidney transplant candidates at higher risk for CMV infection. We plan to give multiple vaccinations prior to living donor kidney transplant, and will follow post-transplant for safety, immunogenicity, and efficacy.

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1532. Increased Risk of Bacterial, Fungal and Other Viral Infections During CMV Infection: Decreased Cytokine Production in Response to Toll-Like Receptor Ligands

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Background. In the solid-organ transplant (SOT) setting, CMV is an immunomodulatory virus that indirectly increases the risk for bacterial, fungal and viral infections. However, the pathogenesis of this phenomenon is poorly understood. The aim of our study was to determine whether inflammatory responses to different Toll-like receptor ligands are blunted during CMV infection in SOT patients.

Methods. CMV D+/R- SOT patients had blood drawn at the end of CMV prophylaxis and then weekly after onset of CMV viremia. PBMCs were extracted and incubated for 24 hours in the presence of bacterial (LPS), fungal (Zymosan [ZYM]), and viral (Resiquimod [R848]) ligands. Proinflammatory (IL1 β), Th1 (IFN γ), Th2 (IL4), immunoregulatory (IL10), and chemotactic (MCP1) cytokines were measured in the supernatant by multiplex ELISA.

Results. Thirty-eight SOT patients were followed for at least 9 months. Patients who developed subsequent CMV infection had lower cytokines in response to bacterial, fungal and viral ligands (LPS, ZYM, and R848) at the end of prophylaxis compared with those with no CMV infection. These results were independent of immunosuppression and peripheral blood cell counts. Specifically, these trends were significantly different with respect to IFN γ , IL1 β , and IL10 production in response to

LPS (P = 0.003, 0.003, and 0.039, respectively), R848 (P < 0.001, 0.039, and < 0.001, respectively) and ZYM (P = 0.039, 0.003, and 0.003, respectively), as well as for MCP1 in response to R848 or ZYM (P = 0.039 for both). In the cohort with CMV infection, cytokine responses to TLR ligands were even lower during the acute CMV infection when compared with the end of prophylaxis, although this was significant only for IL10 production after R848 stimulation (P = 0.034). There was no influence of CMV viral load or duration of viremia on cytokine levels.

Conclusion. Response to non-CMV antigens during CMV infection was blunted supporting the clinical observation in transplant recipients that CMV infection increases susceptibility to bacterial, fungal, and other viral infections. However, inherent differences in patients that are neither directly related to CMV nor to their net level of immunosuppression also contribute to this increased susceptibility, as cytokine levels at the end of prophylaxis were lower among patients with compared with those without subsequent CMV infection.

Disclosures. All authors: No reported disclosures.

This abstract has been withdrawn at the author's request.

1534. Prevalence and Outcome of Neutropenic Enterocolitis Among Pediatric Acute Myeloid Leukemia Patients: A Developing Country Experience Reham Khedr, MD, PhD, Mohamed Abdelwaged, MSc², Marwa Fathy, PhD³ Wael Zekri, MD, PhD4; ²National Cancer Institute-Cairo University / CCHE57357, Nasr City, Egypt, ²Cairo University, Cairo, Egypt, ³National Cancer Institute- Cairo university, Cairo, Egypt and ⁴Cairo University / CCHE57357, Cairo, Egypt

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Background. Neutropenic enterocolitis (NEC) is a life-threatening disease with substantial morbidity and mortality, seen primarily in patients with hematologic malignancies. The frequency of NEC has increased with the widespread use of chemotherapeutic agents such as the taxanes, which cause severe gastrointestinal mucositis.

Methods. This was a retrospective study at the National Cancer Institute, Cairo University. The computerized records were screened for ultrasound or computerized tomographic scan requests for abdominal pain for all acute myeloid pediatrics inpatients (2012-2016). Retrospective case note analysis was used to collect clinical data for patients with features of Typhlitis. D 30 morbrtality was reported.

Results. The incidence of NEC among our inpatients was 24% (49/203). Fortythree children had radiologically confirmed typhlitis, and six had clinical features alone. Most (93%) patients were profoundly neutropenic (ANC <100). All of the patients were subjected to conservative management. All of them needed ICU admission. Eighteen children had a variable period of bowel rest, including 12 patients who were supported with total parenteral nutrition. Three patients had laparotomy that revealed extensive colonic bowel necrosis (1), perforated bowel loop (1), and a perforated appendix (1). Two out of three cases of Laparotomy were diagnosed with Mucormycosis. 30-Days mortality was 44.8% (22/49).Relapsing typhlitis in subsequent courses was observed in 6/27 (22%) patients. Fulminant Gram-negative sepsis without surgical intervention was the leading cause of death in this cohort.

Conclusion. The diagnosis of typhlitis was based on clinical features, supported by radiologic evidence in almost half of the study group. Surgical intervention should be reserved for specific complications or where another surgical pathologic condition cannot reasonably be ruled out. Though rare, fungal infection should be suspected specially in cases with worsening signs of typhlitis despite broad antimicrobial coverage. Disclosures. All authors: No reported disclosures.

1535. Utility of CT Abdomen in Evaluation of Neutropenic Fever in Patients with Hematological Malignancies

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Background. Infections is a serious complication of severe neutropenia and is associated with significant morbidity and mortality. Pan CT scan or CT abdomen is frequently ordered to identify infection source in neutropenic fever. However, utility of CT abdomen in this clinical scenario has not been systematically analyzed.

Methods. We retrospectively reviewed all adults hospitalized at our institution with neutropenic fever from January 2006 to December 2016 and had CT abdomen for source identification. Demographic, clinical, imaging, and outcome data were abstracted and analyzed using descriptive statistics.

Results. Overall, 156 patients (61.5% males) met the study criteria. The most common underlying hematologic malignancies were leukemia in 83 (53.2%) and malignant lymphoma 46 (29.5%). Others included multiple myeloma, myelodysplasia, and benign hematological malignancies. The most common presenting symptoms, besides fever, at the time of CT abdomen were chills (33.5%), abdominal pain (23.9%), nausea (23.2%), diarrhea (20.6%), cough (19.5%), shortness of breath (12.3%), and skin rash (18.4%). Initial CT abdomen was positive in 45 (28.8%). Repeat CT abdomen was obtained in 22 (14.3%) for persistent fevers and had positive findings for infection source in 85.7%. Sources of infection identified on CT abdomen were involving gastrointestinal tract (46.7%), hepatobiliary system (24.4%), urinary tract (21.1%) and peritoneum (7.8%). In terms of microbiology, a causative organism was identified in blood in 53 (34.9%), urine in 15 (9.9%), stool in 15 (9.9%), and respiratory secretions in 8 (5.3%). Causative pathogens included Gram-positive bacteria in 30 (62.5%), Gramnegative bacteria in 23 (47.9%) and Anaerobes in 5 (10.4%) cases. CT abdomen finding resulted in antimicrobial changes in 75 (59.5%) of patients and procedural intervention in 14 patients (9.3%).

Conclusion. While routine use of CT abdomen for evaluation of neutropenic fevers is low yield, CT findings can help identify a source of infection, necessitating change in antimicrobial therapy or procedural intervention, in patients with abdominal symptoms or persistent fever despite broad-spectrum antimicrobial therapy.

Disclosures. All authors: No reported disclosures.

1536. Donor-Derived Mycobacterium tuberculosis Infection After Solid-Organ Transplantation: A Comprehensive Review

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Background. Donor derived Mycobacterium tuberculosis (DDTB) has occasionally been reported after solid-organ transplantation (SOT).

Methods. To characterize DDTB, MEDLINE OVID, and EMBASE were reviewed from inception to December 31, 2016 using key words donor-derived infection, tuberculosis and solid-organ transplant.

Results. A total of 36 cases of proven (17), probable (8) and possible (11) DDTB were identified among 16 lung, 13 kidney, six liver, and one heart recipient. Most patients were male (21/35, 60%); median age was 48 (range 23-68) years. Median time to DDTB was 2.7 (0.2-29) months after SOT. Donor residence in TB-endemic area (13/28, 46.4%) was common. Fever was the most frequent symptom (20/36, 56.5%). DDTB was classified as pulmonary (36%), extra-pulmonary (28%) or disseminated (36%), with common involvement of the allograft (31/36, 86%). Diagnosis was made by smear or culture (30/36, 83.3%). Three (10/24, 41.7%) or four-drug therapy (13/24, 54.2%) was given for median of 10.5 (range, 6-24) months. Graft loss, all-cause and TB-attributable mortality occurred in 18% (4/22), 25% (9/36) and 44% (4/9), respectively.