The Effect of Multi-Component Interventions on the Incidence Rate, Severity, and Duration of Post Open Heart Surgery Delirium among Hospitalized Patients

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Abstract

**Background:** Delirium is one of the prevalent complications of post open heart surgery. The present research aimed to assess the effect of multi-component interventions on the incidence rate, severity, and duration of post open heart surgery delirium among hospitalized patients.

**Methods:** In this quasi-experimental study, 96 patients under open heart surgery were selected using convenience sampling and were divided into a control and an intervention group. The interventions included patients’ preoperative education, nurses’ education, and intra-ward environmental interventions. The demographic information and Mini-Mental State Examination (MMSE) questionnaires were completed a day before surgery. The patients in both groups were also surveyed after extubation until the fourth day post operation using Delirium Observation Screening (DOS) scale considering the incidence, severity, and duration of delirium. The data were analyzed using the SPSS statistical software, version 20.

**Results:** The incidence rate of delirium was 14.6% and 6.2% in the control and intervention groups, respectively (p>0.05). Besides, the mean severity of delirium was 0.53 in the control group and 0.40 in the intervention group (p>0.05). Finally, the mean duration of delirium was 4.5 and 3.25 hours in the two groups, respectively (p>0.05).

**Conclusions:** Since prevention of delirium can play a considerable role in patients’ recovery after heart surgery, it is necessary to carry out some measures to prevent such complications. Even though the interventions performed in this study did not cause significant changes in this regard, the results suggested that prevention interventions should be performed with stronger and more integrated planning for achieving better outcomes.

Background

In spite of emphasis on improvement and prevention of risk factors, cardiovascular diseases are still supposed to be the main disability and mortality factor in developing countries. Nonetheless, progress in new therapeutic methods, such as thrombolytic therapy, and interventional methods, such as Percutaneous Coronary Intervention (PCI), have led to improvement in management and treatment of cardiovascular diseases. Yet, surgical interventions are still propounded as the elective treatment for some patients, and the number of patients under open heart surgery is increasing daily in industrial countries.

Heart surgery may be accompanied by several complications, among which delirium is prevalent with the incidence rate of 30–50%. “Post-surgery delirium” was used by Blachly PH and Starr A. for the first time. According to the American Psychological Association, delirium has been defined as a mental transient syndrome with sudden onset and cognitive impairment, decreasing the level of consciousness, attention disorders, increase or decrease in psychomotor activity, and sleep and wake cycle disorder are among its characteristics. Delirium and dementia should be differentiated from each other because delirium is
specified with sudden onset, fluctuation, attention disturbance, and disorganized thoughts and speech, while dementia is accompanied by gradual onset, no fluctuations, no inattention, and no speech disturbances. Additionally, delirium is recurrent, while dementia is not.\textsuperscript{7}

Post-surgery delirium often appears in the first three days and mostly on the first day after surgery.\textsuperscript{8} The prevalence of post-surgery delirium has been reported to be between 15.0\% and 60.0\%, but this rate is higher after heart surgery.\textsuperscript{9} Delirium is highly prevalent (up to about 80.0\%) at Intensive Care Unit (ICU) and may result in increase in the number of days being under mechanical ventilation, increase in the length of ICU stay and hospital stay, unintentional pulling of tubes and catheters, and mortality.\textsuperscript{10} Delirium not only leads to increase in the length of hospital stay, morbidity, and mortality, but also results in some long-term effects, including functional and cognitive impairment and further hospitalization. Therefore, it imposes great economic burden on the health system as well as extra expenses up to 60,000 dollars on each patient per year.\textsuperscript{11} Hence, some studies mostly using prevention approaches have been carried out to control post-surgery delirium. Accordingly, music therapy and haloperidol-prophylaxis have been introduced as effective methods in prevention of delirium.\textsuperscript{12}

Up to now, no precise pathophysiology and etiology has been mentioned for delirium, but it has been stated to result from some predisposing and underlying factors.\textsuperscript{13} Some of the predisposing factors of delirium include age above 65 years, history of psychotic disorders like schizophrenia, sensory impairment (especially sight), brain injury, history of stroke, sleep deprivation, history of alcohol abuse, prolonged heart and lung bypass, more than one-liter blood transfusion during surgery, hypothermia during surgery and anesthesia, cardiogenic shock, hypoxemia, dehydration, and drugs consumption (psychoactive, analgesic, and hypnotic drugs).\textsuperscript{14} Environmental risk factors are other risk factors the effects of which on the severity of delirium have been shown in a study by Mccusker J. et al. in 2013. Some of these factors include patient's isolation, too few or too many sensory stimulations, inappropriate environmental light and sound, and absence of family at patient's bedside.\textsuperscript{15}

Nowadays, researchers have mostly focused on prevention of delirium and the necessity to carry out studies regarding recognition of its risk factors.\textsuperscript{16} Some researchers admitted that multi-component interventions could reduce the incidence of delirium among heart surgery patients.\textsuperscript{17,18} However, others have reported that prevention pharmacological interventions did not have significant effects on the incidence of delirium.\textsuperscript{19} Based on the previous studies, the main performed interventions were merely related to the post heart surgery period. Also, the incidence rate of delirium was the main outcome evaluated in the previous studies, but the effects of such interventions on the severity and duration of delirium have remained unanswered. Therefore, the present study aims to survey the effect of multi-component interventions on the incidence rate, severity, and duration of post open heart surgery delirium among hospitalized patients.

\textbf{Methods}
Participants and Sampling: In this quasi-experimental study, the research population included all the patients who underwent open heart surgery at Kosar Hospital (Fars Heart Foundation), Shiraz, Iran in February to November 2019. The participants were selected through convenience sampling and in case they fulfilled the inclusion criteria of the study and signed written informed consents, they were recruited into the study. Considering the significance level of 5.0% and power of 80.0%, a 96-subject sample size was determined for the study (48 subjects in each group). The demographic information questionnaire was completed by all the participants and then, they were divided into a control and an intervention group. Since two similar research environments were not available simultaneously and the interventions caused environmental changes as well as changes in the process of patients’ caretaking, first 48 individuals from the control group and then, the 48 ones from the intervention group underwent investigation. The inclusion criteria of the study were aging below 65 years, not suffering from and not having the history of debilitating mental illness and neurological diseases, obtaining minimum Mini-Mental State Examination (MMSE) score of 25.0, not being addicted to drugs and alcohol, and having the ability to communicate. On the other hand, the exclusion criteria of the study were mechanical ventilation for more than 12 hours, heart and lung bypass for more than 90 minutes, and repeated heart surgery for any reason, such as bleeding and tamponade.

Measurement instruments: The applied instruments in this study were as follows

1- Demographic information form, which was prepared based on review of the literature and experts’ opinions. This form included sex, age, place of residence, education level, occupation, history of underlying diseases, and type of surgery.

2- MMSE, which contained 11 questions with a total score between 0 and 30. This scale evaluated cognitive defects and changes over time and could be completed within only 5–10 minutes. In order to investigate the psychometric properties of this instrument in Iranian society, Cronbach's alpha coefficient method (0.87) and split-half technique (0.71) were used. In addition, the validity of MMSE was assessed by concurrent criterion method, revealing a significant inverse relationship between the score of this test and the score produced from the Global Deterioration Scale (GDS) (p = 0.0001). The sensitivity and specificity of the questionnaire were also determined as 0.90 and 0.84, respectively.

3- Delirium Observation Screening (DOS) scale, which consisted of 13 items. The maximum total score of this scale was 13, with scores equal to or more than 3 representing the incidence of delirium. DOS scale was used in this study because it is allocated to nurses, can be completed in less than 5 minutes, and provides the possibility to determine the severity of delirium. The ability to assess the severity of delirium is yet another strong point of this instrument. In this study, DOS scale was scored three times a day by nurses via observation and the final score was computed as the mean of the three scores. The validity of DOS scale was surveyed by Delirium Rating Scale-Revised-98 (DRS-R-98) and was estimated to be 0.67. Additionally, its reliability was assessed by Inter Class Correlation (ICC) and was reported to be 0.99. In order to assess the psychometric properties of DOS scale in Iranian society, this instrument was firstly translated into Persian by an expert translator. Thereafter, in order to determine its content validity, 11
experts were required to express their opinions regarding the relevancy, clarity, and simplicity of the items using Waltz and Bausell validity index. Face validity of the instrument was also assessed by 6 occupied nurses at ICU of heart surgery ward and nursing experts. Moreover, the reliability of DOS scale was evaluated using inter-rater reliability method. In doing so, the instrument was completed for 13 heart surgery patients by the researcher and an educated nursing expert. Then, the correlation between the scores produced by the two observers was calculated, and the results indicated acceptable reliability (r = 0.83).

**Intervention:** The multi-component intervention consisted of three parts, namely training the nursing personnel, training the patients, and environmental interventions. Training the personnel of heart surgery ICU was carried by the researcher in form of a two-hour session. Considering the personnel's rotational shifts and the necessity of their presence, this class was repeated for three times. In this way, training the personnel regarding delirium and its importance, methods of prevention and treatment of delirium, correct way of communicating with such patients, and orienting patients about time, place, and person was carried out at least twice in each shift. The educational content was presented through lecture and discussion. Moreover, the presented materials were installed as a poster at the nursing station, so as to increase the personnel's exposure to the subjects.

Pre-surgery training of the patients was carried out by the researcher at least 24 hours before the open heart surgery. This was done via individual face-to-face conversations along with some pictures from ICU for a period of at least 15 minutes. During this conversation, in addition to education, the patients’ questions and ambiguities were answered. The preoperative educational content consisted of type and process of the disease, surgery method, time of transfer to the ICU after surgery, environment and equipment of ICU, possible limitations such as inability to talk or drink water due to the existence of Endo Tracheal Tube (ETT) and complete rest on the bed, approximate length of ICU stay, limitations related to visiting one's family at the ICU, reassuring the patients regarding pain monitoring and control by the personnel, independence from analgesic drugs, and all the therapeutic and nursing measures that were going to be carried out during the days after the surgery.

In addition to training the personnel and patients, environmental interventions were also conducted in this study. These interventions consisted of light adjustment, using calendar and clock, and providing the patients with the possibility to visit their families at the ICU for 10 minutes once a day and making phone calls to their family members at least once at each shift. Using communication aids, such as glasses, hearing aids, and dentures, for the patients who were using such equipment before the surgery, rapidly sending the patients out of bed, and preventing dehydration by assessing tongue mucus and skin turgor, as one of the risk factors of postoperative delirium, were among the other interventions carried out for the patients.

**Design:** Due to the fact that the physical environment of ICU can influence the study results, it was necessary to carry out the investigation on both control and intervention groups at one ward in order to make a similar research environment. Therefore, both control and intervention groups went under study at
a single ward, but at two different times. Moreover, since the interventions could cause environmental changes as well as changes in the process of patients’ caretaking, first the control group and then the intervention group underwent investigation.

The patients in the control group received the routine care at the post open heart surgery wards of Shiraz Kosar Hospital and were assessed concerning the incidence, severity, and duration of delirium once at the end of each shift until the end of the fourth day after surgery. It should be mentioned that the incidence and severity of delirium was assessed using DOS scale by some fixed in-charges at the ward because at least one of them was present at each shift. Additionally, since they did not play any direct roles in the patients’ caretaking and doing postoperative interventions, they had no knowledge about the control and intervention groups. Before completing the DOS scale, the necessary information about using this instrument was given to the nurses in a one-hour session. Thereafter, their ability in using the instrument was evaluated by putting them at a similar position; i.e., patient’s bedside. After making sure about their knowledge and ability in using the DOS scale, they assessed the incidence and severity of delirium. In the control group, the incidence, severity, and duration of delirium were evaluated by the researcher and trained nurses using the DOS scale after the patients’ coming into consciousness and extubation thrice a day at the end of each shift (8 A.M., 2 P.M., and 8 P.M.) until the end of the fourth day after the surgery.

After completion of sampling in the control group, the other group of participants who met the inclusion criteria and signed written informed consents were entered into the intervention group. Then, the systematic interventions, including training the nursing personnel, training the patients, and environmental interventions, were performed. In this group also, the incidence, severity, and duration of delirium were evaluated as in the control group.

After all, the collected data were entered into the SPSS statistical software, version 20 and were analyzed using independent t-test, chi-square test, one-way ANOVA, and Pearson's correlation coefficient.

**Results**

This study was carried out on 96 patients (48 in the control group and 48 in the intervention group) who underwent open heart surgery. The results of chi-square test indicated that the patients in the control and intervention groups were similar with respect to sex, place of residence, education level, occupation, history of suffering from underlying diseases, and type of surgery (Table 1). Additionally, the results of independent t-test showed no significant difference between the two groups regarding age, MMSE score, length of mechanical ventilation, and length of heart and lung bypass (Table 2).
| Group                              | Control          | Intervention       | P-value |
|-----------------------------------|------------------|--------------------|---------|
|                                   | Frequency        | Frequency percentage | Frequency | Frequency percentage |         |
| **Sex**                           |                  |                    |         |                     |         |
| Female                            | 17.0             | 35.4               | 11.0    | 22.9                | 0.26    |
| Male                              | 31.0             | 64.6               | 37.0    | 77.1                |         |
| **Place of residence**            |                  |                    |         |                     |         |
| The capital (Shiraz)              | 12.0             | 25.0               | 9.0     | 18.8                | 0.34    |
| Cities                            | 24.0             | 50.0               | 31.0    | 64.6                |         |
| Villages                          | 12.0             | 25.0               | 8.0     | 16.7                |         |
| **Level of education**            |                  |                    |         |                     |         |
| Illiterate                        | 9.0              | 18.8               | 8.0     | 16.7                | 0.96    |
| Primary school                    | 11.0             | 22.9               | 12.0    | 25.0                |         |
| High school                       | 5.0              | 10.4               | 5.0     | 10.4                |         |
| Diploma                           | 17.0             | 35.4               | 19.0    | 39.6                |         |
| Academic                          | 6.0              | 12.5               | 4.0     | 8.3                 |         |
| **Occupation**                    |                  |                    |         |                     |         |
| Self-employed                     | 15.0             | 31.3               | 16.0    | 33.3                | 0.64    |
| Employed                          | 6.0              | 12.5               | 3.0     | 6.3                 |         |
| Unemployed                        | 1.0              | 2.1                | 2.0     | 4.2                 |         |
| Retired                           | 14.0             | 29.2               | 18.0    | 37.5                |         |
| Homemaker                         | 11.0             | 22.9               | 8.0     | 16.7                |         |
| Student                           | 1.0              | 2.1                | 0.0     | 0.0                 |         |
| **History of underlying diseases**|                  |                    |         |                     |         |
| Diabetes                          | 2.0              | 4.2                | 3.0     | 6.3                 | 0.57    |
| Hyperlipidemia                    | 5.0              | 10.4               | 4.0     | 8.3                 |         |
| Hypertension                      | 6.0              | 12.5               | 10.0    | 20.8                |         |
| More than one Disease             | 27.0             | 56.3               | 22.0    | 45.8                |         |
| None                              | 8.0              | 16.7               | 9.0     | 18.8                |         |
| **Type of surgery**               |                  |                    |         |                     |         |
| Coronary artery bypass graft      | 38.0             | 79.2               | 40.0    | 83.3                | 1.0     |
| Repair/replace of the valves or other types of surgeries | 10.0 | 2.8 | 8.0 | 16.7 |
Table 2
Comparison of the means of quantitative variables in the control and intervention groups

| Group                                      | Variable                        | Mean  | Standard deviation | P-value |
|--------------------------------------------|---------------------------------|-------|--------------------|---------|
|                                            | Age                             | 53.5  | 9.97               | 0.37    |
|                                            |                                 | 55.19 | 8.52               |         |
|                                            | Mini-mental state examination   | 29.6  | 1.04               | 0.76    |
|                                            |                                 | 26.67 | 0.99               |         |
|                                            | Length of mechanical ventilation| 10.1  | 1.75               | 0.5     |
|                                            |                                 | 9.88  | 1.59               |         |
|                                            | Length of heart and lung bypass | 69.79 | 15.15              | 0.41    |
|                                            |                                 | 67.40 | 13.48              |         |

The results of chi-square test also revealed no significant difference between the control and intervention groups concerning the incidence of delirium (Table 3). Besides, the mean severity of delirium was 0.53 ± 1.43 in the control group and 0.40 ± 1.20 in the intervention group, and the results of independent t-test indicated that the difference was not statistically significant (p > 0.05). Finally, the mean duration of delirium was 4.5 ± 13.86 hours in the control group and 3.25 ± 13.27 hours in the intervention group, and the results of independent t-test demonstrated that the difference was not statistically significant (p > 0.05).

Table 3
Frequency distribution of incidence of delirium in the control and intervention groups

| Group                                      | Control group | Intervention group | P-value |
|--------------------------------------------|---------------|--------------------|---------|
|                                            | Frequency     | Frequency          | Frequency| Frequency%  |                      | 0.31 |
|                                            | percentage    | percentage         | percentage|            |                      |      |
|                                            | Incidence of delirium | Yes | 7.0              | 14.6    | 3.0    | 6.2    |         |      |
|                                            |                | No                | 41.0    | 85.4    | 45.0   | 93.8   |         |      |

Discussion

The present study aimed to investigate the effect of multi-component interventions on the incidence rate, severity, and duration of post open heart surgery delirium among 96 hospitalized patients. The study results showed that multi-component interventions, including training the patients and the nursing
personnel and environmental interventions, had no effects on the incidence rate, severity, and duration of delirium.

According to the results, although the incidence of delirium in the intervention group (3 cases) was lower than that in the control group (7 cases), the difference was not statistically significant. Similarly, Bakker et al. (2014) conducted a study to assess the effect of The CareWell in Hospital program on functional status of hospitalized elderly patients and found that the preventive interventions had no effects on the incidence rate of delirium. In contrast, Mansoori et al. (2018) performed a research on hospitalized patients at Coronary Care Unit (CCU) indicated that modification of sensory stimulations led to a significant reduction in the incidence of delirium in the intervention group compared to the control group. The difference between the results of the present study and the one conducted by Mansoori et al. might be due to the fact that Mansoori et al’s study was done in CCU wards, while ours was performed in ICU after heart surgery. In addition, in the study by Mansoori et al, the patients in the control and intervention groups were at two different wards and, consequently, different personnel and physical environments could influence the results. In the present study, on the other hand, both groups were at a single ward, but underwent investigation at two different times.

Zolfaghari et al. (2011) also investigated the effectiveness of multifactorial interventions in prevention of the incidence of delirium and length of hospital stay among heart surgery patients. The results revealed that these interventions led to a decrease in the incidence of delirium, but did not have any effects on the length of hospital stay. The difference between the results of the research by Zolfaghari et al. and the current one might be attributed to the fact that a different instrument (CAM-ICU) was used in that study to assess delirium. Besides, unlike the present study, the patients exposed to the risk of delirium (including those with prolonged mechanical ventilation and heart and lung bypass) were not excluded from that study.

The findings of our study showed that the mean severity of delirium was 0.53 in the control group and 0.40 in the intervention group, but the difference was not statistically significant. These results were similar to those of the study by Watne et al. (2014), which aimed to investigate the effect of multi-component interventions on prevention of delirium in hospitalized older patients. The results of that study indicated no significant difference between the control and intervention groups regarding the severity of delirium.

The findings of the current study revealed that duration of delirium was 4.5 hours in the control group, which reduced to 3.25 hours in the intervention group. However, the difference between the two groups was not statistically significant. Bjorkelund et al (2010) and Deschodt al (2012) carried out a study to reduce delirium after hip fracture and showed no significant difference between the intervention and control groups regarding the duration of delirium after the interventions.

Overall, difference among the results of various researches in this field could be related to differences in the applied instruments for assessment of delirium, different physical environments, differences in the
nature of the participants’ diseases, not considering the effective factors in the incidence of delirium as the inclusion and exclusion criteria, and differences in therapeutic methods used for management of delirium. In the present study, multi-component interventions did not result in a significant reduction in the incidence, severity, and duration of delirium after heart surgery. However, if the number of study patients and the number of cases with delirium in the control group was more, different results might have been obtained.

One of the limitations of this study was the rotational shift of resident anesthesiologists at the ICU, because various physicians’ therapeutic methods for management and treatment of delirium can influence the results. Moreover, delirium in this study was assessed three times during 24 hours. Therefore, the symptoms of delirium or suffering from fluctuation in the severity of delirium at other times was not considered in this study, which could have affected the results. Hence, further studies with larger sample sizes and elimination of the aforementioned limitations are recommended to be conducted on the issue.

**Conclusion**

Considering the fact that reduction in the incidence, severity, and duration of delirium can play a considerable role in patients’ recovery after heart surgery, it is necessary to take measures to prevent delirium. However, the carried out interventions in the present study did not lead to significant changes in this regard. Therefore, stronger and more integrated preventive interventions should be performed to improve such patients’ recovery process.

**Abbreviations**

Mini-Mental State Examination: MMSE; Delirium Observation Screening: DOS; Percutaneous Coronary Intervention: PCI; Intensive Care Unit: ICU; Coronary Care Unit: CCU; Global Deterioration Scale: GDS; Delirium Rating Scale-Revised-98: DRS-R-98; Inter Class Correlation: ICC; Endo Tracheal Tube: ETT

**Declarations**

**Ethics approval and consent to participate**

All participants gave written informed consent to participate in the study. The present study was conducted in accordance with the principles of the revised Declaration of Helsinki, a statement of ethical principles which directs physicians and other participants in medical research involving human subjects. The participants were assured of their anonymity and confidentiality of their information. Moreover, the study was approved by the local Ethics Committee of Shiraz University of Medical Sciences, Shiraz, Iran (Ethical code: IR.SUMS.REC.7048).

**Consent for publication**
Not applicable.

**Availability of data and materials**

The data will not be shared since the participants are guaranteed full anonymity.

**Competing interests**

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**Authors’ Contributions**

BT has made a substantial contribution to the acquisition of data, analysis of data. MR and AM have made a substantial contribution to the concept, design, data analysis, interpretation of data, revised the article critically and approved the version to be published. NH has made a substantial contribution to the acquisition of data. All authors have read and approved the final version of the manuscript.

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**References**

1. Sanchis-Gomar F, Perez-Quilis C, Leischik R, Lucia A. Epidemiology of coronary heart disease and acute coronary syndrome. Annals of translational medicine. 2016 Jul;4(13).

2. Urden LD, Stacy KM, Thelan LA, Lough ME. Thelan’s critical care nursing: diagnosis and management. Mosby Inc; 2006.

3. Reichert HA, Rath TE. Cardiac surgery in developing countries. The journal of extra-corporeal technology. 2017;49(2):98.

4. Brown IV CH, Laflam A, Max L, Lymar D, Neufeld KJ, Tian J, Shah AS, Whitman GJ, Hogue CW. The impact of delirium after cardiac surgery on postoperative resource utilization. The Annals of thoracic surgery. 2016 ;101(5):1663.

5. Blachy P, Starr A. Post-cardiotomy delirum. American Journal of Psychiatry. 1964;121(4):371-5.

6. Martins S, Fernandes L. Delirium in elderly people: a review. Frontiers in neurology. 2012; 19;3:101
7. Lippmann S, Perugula ML. Delirium or dementia?. Innovations in clinical neuroscience. 2016;13(9-10):56.

8. Whitlock EL, Vannucci A, Avidan MS. Postoperative delirium. Minerva anestesiologica. 2011;77(4):448.

9. Gottesman RF, Grega MA, Bailey MM, Pham LD, Zeger SL, Baumgartner WA, Selnes OA, McKhann GM. Delirium after coronary artery bypass graft surgery and late mortality. Annals of neurology. 2010;67(3):338-44.

10. van den Boogaard M, Schoonhoven L, van Achterberg T, van der Hoeven JG, Pickkers P. Haloperidol prophylaxis in critically ill patients with a high risk for delirium. Crit Care. 2013;17(1):R9.

11. Cerejeira J, Mukaetova-Ladinska EB. A clinical update on delirium: from early recognition to effective management. Nursing research and practice. 2011.

12. Arumugam S, El-Menyar A, Al-Hassani A, Strandvik G, Asim M, Mekkodithal A, Mudali I, Al-Thani H. Delirium in the intensive care unit. Journal of emergencies, trauma, and shock. 2017;10(1):37.

13. Koster S, Hensens AG, Schuurmans MJ, van der Palen J. Risk factors of delirium after cardiac surgery: a systematic review. European Journal of Cardiovascular Nursing. 2011;10(4):197-204.

14. Horacek R, Prasko J, Mainerova B, Lstalova K, Grosmanova T. Delirium in surgery intensive care unit. Act Nerv Super Rediviva. 2011;53:121-32.

15. McCusker J, Cole MG, Voyer P, Vu M, Ciampi A, Monette J, Champoux N, Belzile E, Dyachenko A. Environmental factors predict the severity of delirium symptoms in long-term care residents with and without delirium. Journal of the American Geriatrics Society. 2013;61(4):502-11.

16. Kostas TR, Zimmerman KM, Rudolph JL. Improving delirium care: prevention, monitoring, and assessment. The Neurohospitalist. 2013;3(4):194-202.

17. Zolfaghari M, Arbabi M, Pedram Razi S, Biat K, Bavi A. Effectiveness of a Multifactor Educational Intervention on Delirium Incidence and Length of Stay in Patients with Cardiac Surgery. Hayat. 2012;18(1):67-78.

18. Guo Y, Sun L, Li L, Jia P, Zhang J, Jiang H, et al. Impact of multicomponent, nonpharmacologic interventions on perioperative cortisol and melatonin levels and postoperative delirium in elderly oral cancer patients. Archives of gerontology and geriatrics. 2016;62:112-7.

19. Ghaeli P, Shahhatami F, Zade MM, Mohammadi M, Arbabi M. Preventive intervention to prevent delirium in patients hospitalized in intensive care unit. Iranian journal of psychiatry. 2018 ;13(2):142.

20. Folstein MF, Folstein SE, McHugh PR. “Mini-mental state”: a practical method for grading the cognitive state of patients for the clinician. Journal of psychiatric research. 1975;12(3):189-98.

21. Foroughan M, Jafariei Z, Shirinbayan P, Ghaemmagham Farahani Z, Rahgozar M. Standardization of Mini-Mental State Examination in the senile in Tehran(2006). Updates in the Cognitive Sciences. 2008; 10(2):29-37

22. Scheffer AC, van Munster BC, Schuurmans MJ, de Rooij SE. Assessing severity of delirium by the Delirium Observation Screening Scale. International journal of geriatric psychiatry. 2011;26(3):284-
23. Adamis D, Sharma N, Whelan PJ, Macdonald AJ. Delirium scales: a review of current evidence. Aging & mental health. 2010;14(5):543-55.

24. Bakker FC, Persoon A, Bredie SJH, et al. The CareWell in Hospital program to improve the quality of care for frail elderly inpatients: results of a before-after study with focus on surgical patients. Am J Surg. 2014; 208(5):735–746. doi:10.1016/j.amjsurg.2014.04.009

25. Mansoori, M., et al. Reduction of the Incidence of Delirium in CCU: A Controlled Clinical Trial. Journal of Client-Centered Nursing Care, 4(2), 2018: 104-111. https://doi.org/10.32598/jccnc.4.2.104

26. Watne, L. O. et al. (2014). The effect of a pre- and postoperative orthogeriatric service on cognitive function in patients with hip fracture: randomized controlled trial (Oslo Orthogeriatric Trial). BMC Medicine, 12, 63. doi: https://doi.org/10.1186/1741-7015-12-63

27. Bjorkelund, K. B., Hommel, A., Thorngren, K. G., Gustafson, L., Larsson, S. and Lundberg, D. (2010). Reducing delirium in elderly patients with hip fracture: a multi-factorial intervention study. Acta Anaesthesiologica Scandinavica, 54, 678–688. doi: 10.1111/j.1399-6576.2010.02232.x

28. Deschodt, M. et al. (2012). Preventing delirium in older adults with recent hip fracture through multidisciplinary geriatric consultation. Journal of the American Geriatrics Society, 60, 733–739. doi: https://doi.org/10.1111/j.1532-5415.2012.03899.x