Contributions of Hypoxia-Awareness Training to the Familiarization of Personal Symptoms for Occupational Safety in the Flight Environment

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Abstract: Hypoxia remains a flight-safety issue in terms of aviation medicine. Hypoxia-awareness training has been used to help aircrew members recognize personal hypoxia symptoms. There is still no study, as yet, to establish the association of within-subject data between inflight hypoxia events and the altitude chamber. The main purpose of our study was to use paired subjects’ data on inflight hypoxia symptoms compared with those experienced during training. A questionnaire was developed to obtain information on military aircrew members in 2018. Among 341 subjects, 46 (13.49%) suffered from inflight hypoxia. The majority of the subjects detected ongoing inflight hypoxia on the basis of their previous experience with personal hypoxia symptoms or sensations in previous chamber flights. Of the top five hypoxia symptoms, the data revealed that hot flashes, poor concentration, and impaired cognitive function appeared both during the inflight events and during the hypoxia-awareness training. The occurrence rate of hypoxia symptoms was found to be not significantly different between the in-flight events and the past chamber flights through an analysis of within-subject data. Because the individual memory had faded away over time, fresher hypoxia awareness training is still mandatory and valuable to recall personal hypoxia experience for military aircrew members.

Keywords: hypoxia; aviation medicine; hypoxia-awareness training; altitude chamber

1. Introduction

Based on Dolton’s law, atmospheric pressure at sea level is 760 mmHg, but it decreases with increasing altitude. Moreover, oxygen partial pressure decreases and leads to fewer oxygen molecules being inhaled and less air exchanged by human lungs. In this situation, there is insufficient oxygen being delivered to vital organs, thus impairing those functions,
especially those related to the brain and nervous system. In flight, physiological altitude is generally defined as 10,000 feet, at which pilots who are occupationally exposed to such an environment have no significant degraded performance [1]. In the early days of aviation, however, the pioneers flew hot air balloons, and two French scientists, Croce-Spinelli and Sivel, in their attempt to reach 25,000 feet, were the first hypoxia victims due to a high-altitude environment [2].

Since then, supplemental oxygen and cabin-pressurization systems have been specially developed to prevent in-flight hypoxia. However, fatal hypoxia incidents occasionally occur, which warns all pilots that the hazard of hypoxia to flight safety is always present [3]. On 25 October 1999, a Learjet 35, a small fixed-wing aircraft, was on schedule during a flight from Orlando, FL, USA, to Dallas, TX, USA. However, this plane tragically crashed into the ground, killing both pilots and all the passengers. After the investigation, the significant contributor to this serious accident was deemed hypoxia due to the loss of cabin pressurization [4]. In another case, Helios Airways Flight 522 was flying from Larnaka, Cyprus, to Prague, Czech Republic, on 14 August 2005. Cabin pressurization failure rendered all the aircrew members and passengers unconscious. Thus, there was no one conscious to fly the aircraft, and it ran out of fuel and catastrophically crashed into the terrain below [5]. From the above two cases, we can see that the aircrew members were unaware that they were being exposed to a hypoxic in-flight environment and increasingly lost their ability to take the proper emergency actions.

In World War II, an altitude chamber was constructed to enhance hypoxic awareness and allow military pilots to remember their personal symptoms during hypoxia-awareness training [6]. Island and Fraley (1993) analyzed in-flight hypoxia events from 1976 to 1990 in the United States Air Force (USAF) [7]. If the aircrew members had no experience with hypobaric-awareness training, then there was an extremely high possibility of them losing consciousness during exposure to a hypoxic environment. According to other studies, pilots could recognize ongoing in-flight hypoxia events based on their familiarity with their own personal symptoms and experience during altitude-chamber flights [6,8]. In addition, several studies have reported a consistency in hypoxia symptoms between different sessions of chamber training. The main symptoms during hypoxia-awareness training were impaired mentation, visual changes, dizziness/lightheadedness, paresthesia, tingling, and slurred speech [9–12]. Currently, refresher training courses are mandatory in many countries (e.g., every three years in Saudi Arabia, and every five years in New Zealand and the United States) [13].

To the best of our knowledge, few studies have explored the characteristics of in-flight hypoxia events and investigated their relationship to individual-level symptoms of hypoxia-awareness training [8,14]. Therefore, the aim of this survey was to describe the details of the in-flight hypoxia events and examine the within-subject consistency of the in-flight hypoxia symptoms with those of hypoxia-awareness training. These findings will be useful for enhancing the benefits of hypoxia-awareness training, and in terms of amendments to future programs.

2. Materials and Methods

2.1. Study Design

In Taiwan, the Aviation Physiology Research Laboratory (APRL) organizes and executes an aviation physiology training program for military aircrew members. Prior to flight training, aircrew members must complete an initial altitude chamber flight, and then undertake a refresher training course every four years.

A chamber provides a simulated high-altitude environment that emphasizes the effects of changes in atmospheric pressure on the human body (e.g., sinus and ear blockage, tooth pain, and gastrointestinal discomfort). In addition, the other purpose of altitude-chamber training is to recognize the individual-specific hypoxia symptoms of aircrew members at simulated training altitudes.
We conducted this retrospective and noninvasive/experimental survey to collect information related to the inflight hypoxia experience and hypoxia symptoms from previous chamber flights. This research was assessed as low-risk work and approved by the Institutional Review Board of Kaohsiung Armed Forces General Hospital in Kaohsiung City, Taiwan (No. KAFGH 107-02).

2.2. Study Subjects

Study subjects were recruited from all the triservice military aircrews that had experience with altitude-chamber flights. We encouraged and invited them to join this research from 1 January to 31 December 2018, when they received the refresher training course at the APRL. Prior to the prechamber flight briefing, the aviation physiologist/instructor obtained informed consent from all the trainees and delivered the questionnaires to collect data on the inflight hypoxia conditions they experienced and their recall memory of the hypoxia symptoms they experienced during previous chamber flights. In total, 341 subjects voluntarily participated in this study. Among them, 46 (13.49%) had experienced an inflight hypoxia problem at some point during their flight career, and were finally enrolled in the data analysis.

2.3. Data Collection

We designed an anonymous and structured questionnaire to collect information about the inflight hypoxia events and previous hypoxia experience during the chamber flights. This questionnaire was composed of several parts: Part I, demographic items about the subjects’ personal characteristics (age, gender, station, airframe, flight years, and flight hours); Part II, experience with inflight hypoxia (yes or no) and the characteristics of inflight hypoxia episodes, including the cabin/flight altitude at which these episodes occurred, potential causes of in-flight hypoxia events, and how to detect such events in the future and immediately take corrective measures obtained by the close type options; and Part III, memory of frequently described hypoxia symptoms recognized during the inflight hypoxia incidents and past chamber flights, such as poor concentration, hot flashes, impaired cognitive function, dizziness/lightheadedness, visual disturbances, numbness, air hunger, paresthesia, fatigue, anxiety, tingling, and nausea [15–17].

2.4. Statistical Analyses

The frequency and proportion of the descriptive statistics were applied to characterize the distribution of all the questionnaire’s data in this survey. Using the McNemar test, hypoxia symptoms that occurred during the inflight hypoxia incidents were then compared with those during the past chamber flights for the eligible participants. All data from this survey were entered and handled using the SPSS 24 program (IBM, Armonk, NY, USA). The significance level was determined by a two-tailed \( p \) value < 0.05.

3. Results

We received 46 questionnaire responses about the inflight hypoxia cases during the study period. The distribution of age subgroups, <30, 30–39, and \( \geq 40 \) years, were 17.4%, 43.5%, and 39.1%, respectively; only one case involved a female pilot. Of the participants, over 90% were pilots. Fighter pilots (60.9%) most commonly encounter hypoxic conditions, followed by cargo/transporter pilots (26.1%). Moreover, 30.4%, 54.3%, and 15.3% of the participants had <10, 10–19, and \( \geq 20 \) years of flight experience, respectively. The flight hours of nearly half of all participants totaled more than 2000 (Table 1).
Table 1. Characteristics of study participants.

| Variables               | n (%) |
|-------------------------|-------|
| Age (years)             |       |
| <30                     | 8 (17.4) |
| 30–39                   | 20 (43.5) |
| ≥40                     | 18 (39.1) |
| Gender                  |       |
| Male                    | 45 (97.8) |
| Female                  | 1 (2.2) |
| Station                 |       |
| Pilot                   | 42 (91.3) |
| Nonpilot                | 4 (8.7) |
| Airframe                |       |
| Fighter                 | 28 (60.9) |
| Jet trainer             | 6 (13.0) |
| Cargo/transporter       | 12 (26.1) |
| Flight years            |       |
| <10                     | 14 (30.4) |
| 10–19                   | 25 (54.3) |
| ≥20                     | 7 (15.3) |
| Flight hours            |       |
| <1000                   | 10 (21.8) |
| 1000–1999              | 14 (30.4) |
| ≥2000                   | 22 (47.8) |

Table 2 summarizes the data obtained in relation to the inflight hypoxia events. The two most common causes of the hypoxia events occurred at altitudes of 15,000–19,999 (37.0%) and 25,000–29,999 (19.6%) feet. Approximately 40% of the inflight hypoxia events were due to abnormal cabin pressurization: depressurization (17.4%) or failure to pressurize (21.7%). There were 13 participants (28.2%) who stated that mask leakage/unsealing and hose disconnection may have triggered the hypoxia events they experienced. There were 10 respondents that reported the reasons for such events were physiological. Five respondents (11.0%) did not explain the causes for such events. The majority of the respondents reported similar hypoxia symptoms and feelings relative to those they experienced during their previous chamber flights. One-fifth of the respondents recognized hypoxia occurrence when conducting oxygen-system checks. Over 80% described that there was mild (58.7%) and significant degradation (21.7%) of flight performance due to the hypoxia impact. They immediately followed emergency oxygen usage (69.2%), descent (69.2%), and even mission-abortion (34.8%) procedures to prevent loss of consciousness.

We calculated the prevalence of the inflight and chamber-flight hypoxia symptoms (Table 3). The majority of aircrew members experienced several hypoxia symptoms. In flight, the aircrew members frequently detected the top five hypoxia symptoms—hot flashes (43.5%), poor concentration (34.8%), impaired cognitive function (32.6%), paresthesia (28.3%), and fatigue (26.1%). In chamber flights, hot flashes, poor concentration, and impaired cognitive function repeatedly appeared during hypoxia-awareness training. Of these symptoms, the most common symptoms that occurred both in flight and during the chamber flights were poor concentration, hot flashes, impaired cognitive function, paresthesia, and visual disturbance, in descending order. We also found that there was no significant difference in the frequency of the in-flight hypoxia symptoms when compared to those during the chamber flights (all p values > 0.05).
Table 2. Characteristics of inflight hypoxia events.

| Variables                        | n (%)  |      |      |
|----------------------------------|--------|------|------|
| Altitude (feet)                  |        |      |      |
| <15,000                          | 8 (17.4)|      |      |
| 15,000–19,999                    | 17 (37.0)|      |      |
| 20,000–24,999                    | 7 (15.2)|      |      |
| 25,000–29,999                    | 9 (19.6)|      |      |
| ≥30,000                          | 5 (10.8)|      |      |
| Causes                           |        |      |      |
| Depressurization                 | 8 (17.4)|      |      |
| Failure to pressurize            | 10 (21.7)|      |      |
| Mask leakage/unsealing           | 3 (6.5)|      |      |
| Hose disconnection               | 10 (21.7)|      |      |
| Physiology                       | 10 (21.7)|      |      |
| No response                      | 5 (11.0)|      |      |
| Detection                        |        |      |      |
| Personal hypoxia symptoms        | 16 (34.8)|      |      |
| Sensations like those experienced in previous chamber flights | 14 (30.4)|      |      |
| Abnormal sensation               | 9 (19.6)|      |      |
| Oxygen system check              | 10 (21.7)|      |      |
| Impact of flight performance     |        |      |      |
| None                             | 9 (19.6)|      |      |
| Mild degradation                 | 27 (58.7)|      |      |
| Significant degradation          | 10 (21.7)|      |      |
| Procedures                       |        |      |      |
| Oxygen usage                     | 32 (69.2)|      |      |
| Descent                          | 32 (69.2)|      |      |
| Abortion                         | 16 (34.8)|      |      |

Table 3. Comparison of inflight and chamber-flight hypoxia symptoms.

| Symptoms                      | In Flight n (%) | Chamber Flight n (%) | Both n (%) | p Value |
|-------------------------------|-----------------|----------------------|------------|--------|
| Hot flashes                   | 20 (43.5)       | 22 (47.8)            | 13 (28.3)  | 0.804  |
| Poor concentration            | 16 (34.8)       | 23 (50.0)            | 13 (28.3)  | 0.092  |
| Impaired cognitive function   | 15 (32.6)       | 17 (37.0)            | 8 (17.4)   | 0.804  |
| Paresthesia                   | 13 (28.3)       | 14 (30.4)            | 8 (17.4)   | 1.000  |
| Fatigue                       | 12 (26.1)       | 11 (23.9)            | 4 (8.7)    | 1.000  |
| Visual disturbance            | 11 (23.9)       | 20 (43.5)            | 6 (13.0)   | 0.064  |
| Air hunger                    | 10 (21.7)       | 14 (30.4)            | 5 (10.9)   | 0.424  |
| Dizziness/lightheadedness     | 6 (13.0)        | 18 (39.1)            | 4 (8.7)    | 0.250  |
| Tingling                      | 6 (13.0)        | 7 (15.2)             | 3 (6.5)    | 1.000  |
| Numbness                      | 5 (10.9)        | 10 (21.7)            | 2 (4.3)    | 0.227  |
| Anxiety                       | 4 (8.7)         | 10 (21.7)            | 3 (6.5)    | 0.070  |
| Nausea                        | 0 (0.0)         | 3 (6.5)              | 0 (0.0)    | -      |

4. Discussion

From this survey, we discovered 46 inflight hypoxia events (13.49%) from the military aircrew members who attended refresher altitude-chamber training in 2018. Hypoxia is still a hazard that threatens flight safety, and the main cause of hypoxia events is a pressurization-system malfunction. Most events were detected by the participants because of their experience with personal hypoxia symptoms or those that they experienced during previous chamber flights. There was a high similarity in terms of symptoms between the inflight hypoxia events and those during the hypoxia-awareness training.

The previous study described that there was a smaller proportion of the inflight hypoxia incidents in the population of student pilots [14]. Simply stated, this study showed that fewer hypoxia cases occurred in aircrew members who were young, had fewer flight
years, or had fewer flight hours. The explanation for this may be that when compared with
experienced pilots, the junior pilots usually fly during monitored training courses and at
relatively low altitudes due to the type of aircraft. These findings noted that the majority of
events occurred for the fighter pilots, consistent with the data analyzed by Deussing et al.
(2011) [18]. Because there were relatively small samples in this study, we conservatively
explained the tendency related to the flight experience. Future extended work will be
conducted to investigate the relationship between the inflight hypoxia characteristics and
the demographic factors by collecting more cases.

The current data have shown that hypoxia events often occur at altitudes from 15,000
to 19,999 feet, similar to previous data [8,14]. One study used cabin altitude to present the
environment in which the pilots actually stayed at that time of inflight hypoxia attacks [6,8].
Because almost all of our respondents did not focus on cabin altitude during such episodes,
we were not able to provide an outcome related to altitude in this study. During the period
from 1970 to 1980, 298 inflight hypoxia events were recorded by the USAF. Cabin or cockpit
decompression (19.46%) was found to be the leading cause of inflight hypoxia, followed
by regulator malfunction (15.77%), oxygen-hose disconnection (9.40%), and poor mask fit
(4.03%). Our survey also demonstrated that nearly 40% of incidents were attributed to
pressurization-system problems, and 20% of them were induced by oxygen-hose discon-
nection. Interestingly, some participants reported that physiological issues hindered them
and were related to hypoxia, such as fatigue, nutrition, and smoking. These self-imposed
stressors may increase pilots’ physiological altitude and make them more susceptible to
hypoxic environments in flight [19]. These issues are worth assessing in future studies
to determine how they affect the hypoxia tolerance of aircrew members. Oxygen-hose
disconnection was another important cause of inflight hypoxia conditions. In this study,
the results revealed that the majority of cases occurred for fighter pilots. These pilots
usually perform acrobatic combat maneuvers and are exposed to a high G environment.
The potential explanation for this majority is that the oxygen hose can be disconnected
after a high G load if it was not assembled properly before the flight.

In the USAF, an analysis showed that there were 656 inflight hypoxia incidents that
involved 606 altitude-chamber-trained and 50 untrained aircrew members. Only 3.8% of
those trained experienced a loss of consciousness; however, 94% of the untrained members
suffered from a loss of consciousness [20]. A chamber provides a simulated high-altitude
environment at ground level. There are several purposes for hypobaric training: (1) to real-
ize the physiological effects of changes in atmospheric pressure on the body, (2) to recognize
personal hypoxia symptoms and behavior affected by hypoxia, and (3) to learn the proper
corrective techniques using onboard oxygen systems. The abovementioned USAF survey
also indicated that nearly one-fourth of aircrew members detected inflight hypoxia on the
basis of their experience during chamber flights [7]. Several studies have also shown that
aircrew members use their personal hypoxia symptoms from hypoxia awareness training
to recognize inflight hypoxia events [8,14,18]. Corresponding to the previous findings,
two-thirds of the respondents in the current study also detected hypoxia attacks through
their memory or experience during past altitude-chamber training. However, some air-
crew members found ongoing inflight hypoxia when they inspected the oxygen system
during the flight. The reason for this may be that they were always educated to follow the
rules when inspecting the oxygen system in a disciplined manner so as to avoid hypoxia
incidents before and during the flight. This check technique is always mentioned during
the chamber flight training. Therefore, the aircrew members are regularly accustomed
to performing inflight oxygen-system checks and then possibly detecting and correcting
hypoxia occurrence in the early flight stage, even without any symptoms. Few studies
have revealed data on the behavioral changes affected by these inflight hypoxia conditions.
We extended this work to collect information on flight-performance degradation among
the participants. More than 20% of the participants described the significant impact of a
hypoxia attack on their performance. When exposed to an environment with a low oxygen
concentration, humans have a short period of time, called the time of useful consciousness
(TUC), to make life-saving decisions. All of the study participants still had the ability to carry out reactive procedures during the TUC. As aforementioned, the military aircrew members could discover and handle inflight hypoxia events based on personal symptoms and sensations from their experience during chamber flights. Therefore, our findings also enhance the importance of hypoxia-awareness training inside altitude chambers.

From the literature review, it can be seen that the results demonstrated that there were variable orders and frequencies of hypoxia symptoms during the chamber flights. Nonetheless, the dominant hypoxia symptoms included mental impairment, coordination issues/slowing response, visual changes, lightheadedness/dizziness, and hot flashes [9,11]. Consistent with former findings, the top five symptoms of our survey by their order of frequency were poor concentration, hot flashes, visual disturbance, impaired cognition, and dizziness/lightheadedness inside the chamber. In terms of the inflight hypoxia events, the most common symptoms reported were paresthesia, lightheadedness, dizziness, decreased mental ability, and hot/cold flashes [8,18]. Our findings present the highly overlapping symptoms of the inflight hypoxia episodes with those of other studies. Although the majority of the main hypoxia symptoms were similar to those of the chamber flights, there was still a mild difference between them, which can be related to the characteristics of multitasking flight duty for the pilots, in contrast to the fact that they merely focus on experienced hypoxia symptoms inside the chamber.

A significant number of military aviators have stated that they recognized their own symptoms and that they were becoming hypoxic and took timely corrective steps because they recalled their altitude-chamber training [8,14]. In terms of the United States Navy, Deussing et al. (2011) described that the overall symptom spectrum of inflight hypoxia was consistent with that of the hypoxia-awareness training [18]. A comparison of the occurrence of “every single” hypoxia symptom between the inflight and chamber-flight conditions has still not been addressed. Of the 16 symptoms in this study, only five illustrated a significant difference in reporting the scenarios by a comparison of group analyses, but included three main symptoms: tingling, difficulty concentrating, and air hunger. In the current study, the method of individual analysis was substituted to manage the within-subject data of every single hypoxia symptom from the two different exposure events. Surprisingly, dominant hypoxia symptoms reappeared in the different patterns, and all the frequencies were not significantly different between the inflight hypoxia and that during the chamber-flight training. Previous studies have identified that the memory of hypoxia symptoms from chamber flights gradually fades away over time [12]; however, the proportion of primary symptoms was not different between the repetitive exposures, especially for senior aircrew members. In other words, hypoxia-symptom memories can be recalled and enhanced by repetitive training. Similarly, nearly 80% of the participants that recalled hypoxia memories from previous trainings had more flight years and experiences with hypoxia-awareness training, which can likely be explained by the consistency of the symptoms between the inflight hypoxia events and the chamber-flight training. Hypoxia-awareness training is properly repeated in fixed time intervals of three to six years, suitable to refresh the memory of hypoxia symptoms, correct use of emergency equipment, and recovery procedures [9,13]. Similar to the former reports, our findings emphasize the necessity of periodic altitude-chamber indoctrination for all aircrew members.

There are still inherent limitations in the current study. First, the underestimation of inflight hypoxia events may exist, because the pilots are more sensitive to this safety issue. Second, we designed a structured questionnaire to obtain the details of hypoxia attacks described by the participants. Because memory can diminish after several years, reporting errors cannot be ruled out from this study. In the future, monitoring devices integrated with the flight or hypoxia-awareness training would be developed to record objective physiological data and reduce the nondifferential errors. Third, some symptoms of physiological events, such as hyperventilation, are very similar to those of a hypoxic condition. Aircrew members probably misdiagnose this event as being a hypoxia case. However, the subjects declared the inflight hypoxia symptoms corresponded to their
experiences inside the altitude chamber. They complied with the disciplines and recovered from the event by using oxygen. Thus, misclassification of other events was unlikely to exist in this study. Next, we sampled the participants during the physiological refresher training over one year. Thus, these results are limited and should not be generalized to the whole community of military aircrew members. Finally, a reduced-oxygen breathing device (ROBD) was used to execute hypoxia-awareness training. The hypoxia symptoms induced by this ROBD are in line with those in the altitude chambers [21,22]. Because there is no training capability in our country, we cannot compare the inflight hypoxia data with the ROBD information.

5. Conclusions

In summary, we have illustrated that the malfunction of the pressurization system was the main cause of inflight hypoxia events among military aircrew members. Meanwhile, they managed the ongoing inflight hypoxia that relied on the experience of the chamber training. The dominant hypoxia symptoms inside the chamber were poor concentration, hot flashes, visual disturbance, impaired cognition, and dizziness/lightheadedness, in that order. Our analysis emphasized the within-subject reappearance and consistency of considerable hypoxia symptoms during the inflight hypoxia events and those during the chamber flights. Therefore, hypoxia-awareness training plays an important role in acquainting aircrew members with personal hypoxia symptoms and training them how to respond when facing hypoxic situations, contributing to enhanced flight safety.

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References

1. Pickard, J.S. The Atmosphere and Respiration. In Fundamentals of Aerospace Medicine, 3rd ed.; DeHart, R.L., Davis, J.R., Eds.; Lippincott Williams and Wilkins: Baltimore, MD, USA, 2002; pp. 19–38.
2. Dille, J.R. In the beginning. In Fundamentals of Aerospace Medicine, 3rd ed.; DeHart, R.L., Davis, J.R., Eds.; Lippincott Williams and Wilkins: Baltimore, MD, USA, 2002; pp. 1–18.
3. Ghosh, P.C.; Pant, P. In-flight hypoxia—still a worry bane. Ind. J. Aerosp. Med. 2010, 54, 7–12.
4. National Transportation Safety Board. Report on the Crash of Sunjet Aviation, Learjet Model 35, N47BA on 25 October 1999. (Report Number: AAB-00-01). 11 December 2012. Available online: https://www.ntsb.gov/investigations/AccidentReports/Reports/AAB0001.pdf (accessed on 21 January 2021).
5. Air Accident Investigation & Aviation Safety Board. Report on Accident of the a/c 5B-DBY of Helios Airways, Flight HCY522 on 14 August 2005, in the area of Grammatiko, Attikis, 33 km Northwest of Athens International Airport. November 2006. Available online: https://reports.aviation-safety.net/2005/20050814-0_B733_5B-DBY.pdf (accessed on 21 January 2021).
6. Files, D.S.; Webb, J.T.; Pilmanis, A. Depressurization in military aircraft: Rates, rapidity, and health effects for 1055 incidents. *Aviat. Space Environ. Med.* **2005**, *76*, 523–529. [PubMed]

7. Island, R.T.; Fraley, E.V. Analysis of USAF hypoxia incidents January 1976 through March 1990. In Proceedings of the 7th International Symposium on Aviation Psychology, Dayton, OH, USA, 22–29 April 1993; pp. 664–668.

8. Cable, G.G. In-flight hypoxia incidents in military aircraft: Causes and implications for training. *Aviat. Space Environ. Med.* **2003**, *74*, 169–172. [PubMed]

9. Johnston, B.J.; Iremonger, G.S.; Hunt, S.; Beattie, E. Hypoxia Training: Symptom Replication in Experienced Military Aircrew. *Aviat. Space Environ. Med.* **2012**, *83*, 962–967. [CrossRef] [PubMed]

10. Smith, A.M. Hypoxia Symptoms in Military Aircrew: Long-Term Recall vs. Acute Experience in Training. *Aviat. Space Environ. Med.* **2003**, *74*, 169–172. [PubMed]

11. Tu, M.-Y.; Chiang, K.-T.; Cheng, C.-C.; Li, F.-L.; Wen, Y.-H.; Lin, S.-H.; Lai, C.-Y. Comparison of hypobaric hypoxia symptoms between a recalled exposure and a current exposure. *PLoS ONE* **2020**, *15*, e0239194. [CrossRef] [PubMed]

12. Woodrow, A.D.; Webb, J.T.; Wier, G.S. Recollection of Hypoxia Symptoms Between Training Events. *Aviat. Space Environ. Med.* **2011**, *82*, 1143–1147. [CrossRef] [PubMed]

13. Neuhaus, C.; Hinkelbein, J. Cognitive responses to hypobaric hypoxia: Implications for aviation training. *Psychol. Res. Behav. Manag.* **2014**, *7*, 297–302. [CrossRef] [PubMed]

14. Rayman, R.B.; McNaughton, G.B. Hypoxia: USAF experience 1970-1980. *Aviat. Space Environ. Med.* **1983**, *54*, 357–359. [PubMed]

15. Artino, A.R.; Folga, R.V.; Swan, B.D. Mask-on hypoxia training for tactical jet aviators: Evaluation of an alternate instructional paradigm. *Aviat. Space Environ. Med.* **2006**, *77*, 857–863. [PubMed]

16. Patrño, L.; Zorro, S.; Silva, J.; Castelo-Branco, M.; Ribeiro, J. Flight physiology training experiences and perspectives: Survey of 117 pilots. *Aviat. Space Environ. Med.* **2013**, *84*, 620–624. [CrossRef] [PubMed]

17. Sampson, J.B.; Kobrick, J.L. The environmental symptoms questionnaire: Revisions and new field data. *Aviat. Space Environ. Med.* **1980**, *51*, 872–877. [PubMed]

18. Deussing, E.C.; Artino, A.R., Jr.; Folga, R.V. In-flight hypoxia events in tactical jet aviation: Characteristics compared to normobaric training. *Aviat. Space Environ. Med.* **2011**, *82*, 775–781. [CrossRef] [PubMed]

19. Yoneda, I.; Watanabe, Y. Comparisons of altitude tolerance and hypoxia symptoms between nonsmokers and habitual smokers. *Aviat. Space Environ. Med.* **1997**, *68*, 807–811. [PubMed]

20. Dully, F.E. Altitude chamber training: Is it worth the risk? *Hum. Factors Aviat. Med.* **1992**, *39*, 1–8.

21. Singh, B.; Cable, G.G.; Hampson, G.V.; Pascoe, G.D.; Corbett, M.; Smith, A. Hypoxia Awareness Training for Aircrew: A Comparison of Two Techniques. *Aviat. Space Environ. Med.* **2010**, *81*, 857–863. [CrossRef] [PubMed]

22. Vacciano, C.A.; Vagedes, K.; Gonzalez, D. Comparison of the physiological, cognitive, and subjective effects of sea level and altitude-induced hypoxia [abstract]. *Aviat. Space Environ. Med.* **2004**, *75*, B56.