A Model-Checking Framework for Neuro-Degenerative Deficit Screening and Personalized Training

Elisabetta De Maria and Christopher Leturc Université Côte d'Azur, CNRS, I3S, France {elisabetta.de-maria, christopher.leturc}@univ-cotedazur.fr

- Keywords: Model-Checking, Probabilistic Model, Neuro-Cognitive Diseases, Medical Application, Serious Game, Cognitive Training.
- Abstract: Serious games are established as an effective tool to screen cognitive deficits and assess diagnosis in patients affected by neuro-degenerative diseases such as Alzheimer or Parkinson. They are also known for their cognitive training benefits. According to the latest DSM-5 classification, we can discriminate mild Neuro-Cognitive Disorders (mild NCDs) and Major Neuro-Cognitive Disorders (Major NCDs). In this article, we consider three classes of patients: healthy, mild NCD, and Major NCD. For each class, we use Discrete Time Markov Chains to model the behaviour shown while playing serious games. Model checking techniques allow us to spot the difference between the expected and the observed behaviour. As a main contribution, we provide a new theoretical framework allowing us to evaluate how the confidence level of practitioners on the patient's Alzheimer degree evolves after each game session, i.e., *help to diagnose*, and to set up an experimental protocol in which the levels of the proposed subsequent game sessions automatically depend on the patient behaviour observed in the previous sessions, i.e., *help to train*.

1 INTRODUCTION

Neuro-degenerative pathologies such the as Alzheimer or Parkinson disease often lead to the decline of cognitive functions and, more generally, to cognitive impairments. To provide timely and individualized actions, the presence of neurocognitive disorders should be detected as soon as possible and constantly monitored by clinicians. Presently, an accurate diagnosis typically involves a comprehensive series of neuropsychological tests, frequently accompanied by biomarker tests. Conducting these tests can be demanding and time-consuming, both for the practitioner and the patient. There is an increasing interest in discovering behavioral markers that are objective, quick to conduct, and can supplement traditional clinical assessments, aiding in the early detection of alterations in cognitive performances. Serious games are very promising in this context (Philippe et al., 2014). They are digital or physical games designed with a primary purpose beyond entertainment, such as education, training, health, or social change, while maintaining engaging gameplay elements (Alvarez et al., 2007). Several feasibility studies have demonstrated the value of serious games in assessing cognitive impairment

(Tong et al., 2016; Vanessa et al., 2017; Kato and Klerk, 2017). Additionally, research highlights their potential in Alzheimer's disease therapy as cognitive training tools (Anguera et al., 2013; Kathryn et al., 2014). Furthermore, studies indicate that elderly people exhibit a preference for games compared to traditional cognitive exercises (Melenhorst et al., 2006).

Based on the most recent DSM-5 (American Psychiatric Association, 2013) classification, cognitive impairments involve both a decline in cognitive functions and behavioral issues that can disrupt everyday activities. Depending on the severity of these deficits and on their impact, this classification discriminates mild Neuro-Cognitive Disorder (mild NCD) and Major Neuro-Cognitive Disorder (Major NCD). Patients with mild NCD and major NCD require supervision from medical practitioners and psychologists.

In this work we advocate the use of serious games to help practitioners in screening cognitive deficits and proposing training activities suited to patients. For some games, several difficulty levels can be proposed. For each Alzheimer degree (healthy, mild NCD, Major NCD), the activity of patients while playing is modeled using discrete Markov chains, in the style of (De Maria et al., 2019; L'Yvonnet et al.,

De Maria, E. and Leturc, C.

DOI: 10.5220/0013095300003911

Paper published under CC license (CC BY-NC-ND 4.0)

In Proceedings of the 18th International Joint Conference on Biomedical Engineering Systems and Technologies (BIOSTEC 2025) - Volume 2: HEALTHINF, pages 351-360

ISBN: 978-989-758-731-3; ISSN: 2184-4305

A Model-Checking Framework for Neuro-Degenerative Deficit Screening and Personalized Training.

2021). As a matter of fact, when patients play serious games, some scenarios frequently happen while others are rare. We quantify these variations in the patient behaviour by associating probabilities with the important actions of games. Certain actions will definitely be carried out by the patient while other actions depend on the Alzheimer degree of the patient. For these actions, practitioners provide us with *a priori* weights or probabilities are initially integrated into our models and refined based on clinical study results.

For the correct implementation of our methodology, practitioners initially need to give us: (i) for each game and for each Alzheimer degree, the a priori weights associated with key activities of the game (if available, weights coming from clinical experiments will replace a priori weights); (ii) the hypothesis on the Alzheimer degree, i.e., impairment degree, of each patient, based on a preliminary set of pen and paper neuropsychological tests. To ensure the validity of important properties, e.g., all executions reach a final state, our models are validated thanks to the use of probabilistic model checking (Hansson and Jonsson, 1994), which allows to automatically check if some dynamic properties are respected. By computing probabilities associated with execution traces, model checking also helps in spotting the differences between the patient expected behaviour and the observed one.

As a main contribution, we propose the introduction of a meta-automaton whose nodes are Markov chains representing the expected behaviour of a class of patients for a given game. Each state of the meta-automaton represents a game session. This meta-automaton allows one: (i) to infer how the confidence level of practitioners on the patient's Alzheimer degree evolves after each game session, and (ii) to set up an experimental protocol in which the difficulty level of the proposed subsequent game sessions depends on the patient behaviour observed in the previous game sessions (improvements/regressions/constant behaviour). Some suited final conditions allow us to determine when the game sessions should end. At the end of the protocol, we may suggest to practitioners to reconsider the patient's Alzheimer degree provided at the beginning (e.g., a patient known as "Major NCD" could be considered to be re-classed as "mild MCD" or vice-versa). Instabilities in the patient performances (e.g., oscillations) can also be automatically detected thanks to model checking techniques. Model checking (Clarke et al., 1999b) is thus applied at two levels: to validate the Markov chains describing the activity of patients playing serious games, and to detect cru-



Figure 1: Screenshot of the Match Items game.

cial properties of traces of the meta-automaton.

The paper is organized as follows. In Section 2 we introduce, as a case study, a simple serious game developed at Claude Pompidou Institute, Nice, France. In Section 3 we give some preliminaries on the formal tools we adopt: probabilistic automata and temporal logics. Section 4 is devoted to the formal framework we propose to help with diagnosis and training of Alzheimer patients. In Section 5 we propose a formal validation of our approach and in Section 6 we outline important future developments.

2 CASE STUDY

As a simple case study, we introduce the Match Items game (Tran et al., 2015), which has been developed in 2014 at Claude Pompidou Institute, Nice, France and has already been implied in several clinical protocols. In particular, one of the clinical experiments validated this game as a suitable tool to discriminate between mild NCD patients and healthy ones. The Match Items game targets selective and sustained visual attention functions. In this game patients interact with a touch-pad. Their task consists in matching a random picture shown at the center of the touch-pad with its corresponding element from a list of pictures located at the bottom of the screen (see Figure 1). The game lasts at most five minutes.

When the patient selects the correct picture, a happy smiley appears, and a new picture is displayed. In case of an incorrect choice, a sad smiley is displayed, prompting the patient to try again. If there is over 10 seconds of inactivity on the touch-pad, the game reminds the patient to select a picture. Exiting the game zone results in the game being stopped. In the rest of the paper, key actions of this game will be referred as follows: $\alpha :=$ "the patient chooses the right picture", $\beta :=$ "the patient chooses the wrong picture", $\gamma :=$ "the patient is inactive", $\theta :=$ "the patient quits the game zone".

3 PRELIMINARIES

In this section, we present the Probabilistic Finite Deterministic Automata (PDFA) and temporal logics. In the following subsections, we delve into the details of these two concepts, laying the groundwork for our subsequent exploration of model-checking and its application to serious game-based neuro-degenerative screening.

3.1 Probabilistic Deterministic Finite Automata

Let us consider patients engaged in playing serious games. Given our focus on studying the patients activity and the unpredictability of patients' actions, we adopt Probabilistic Deterministic Finite Automata (PDFA) (Rabin, 1963). Given a serious game, we conceive an automaton for each class of patients. This automaton serves a dual purpose by representing the development of the serious game and specifying the activity the user is supposed to display if she belongs to the class being tested by the automaton.

Each serious game is represented as a PDFA, where the states represent the different game configurations, e.g., the user has to choose a picture, or the end of the game. The input symbols of the alphabet represent the actions the user can make, denoted by $\Sigma_g = \{\alpha, \beta, \gamma, \theta\}$. We note $\Sigma_g^* \subseteq \Sigma_g^{\mathbb{N}^*}$ the set of words we can write with Σ_g . A word is just a concatenation of symbols on Σ_g , $w = \alpha\beta\gamma\beta\gamma\beta\alpha\beta\theta$ is a word that uses the symbols of Σ_g , thus $w \in \Sigma_g^*$.

Definition 1. *A* Deterministic Finite Automaton (*DFA*) *is a 5-tuple A* = ($Q, \Sigma, \delta_d, q_0, F$) *where:*

- *Q* is a finite set of states.
- Σ is a finite alphabet of input symbols.
- $\delta_d : Q \times \Sigma \to Q$ is the transition function, where $\delta_d(q,a)$ represents the next state when being in state q and reading input symbol a.
- $q_0 \in Q$ is the initial state.
- $F \subseteq Q$ is the set of accepting (final) states.

We define δ^* the transition function w.r.t. δ which operates on a set of words:

$$\delta_d^* : Q \times \Sigma^* \to Q \tag{1}$$

$$\delta_d^*(q,()) = q \tag{2}$$

$$\delta_d^*(q, xa) = \delta_d(\delta_d^*(q, x), a)$$
 for $x \in \Sigma^*, \ a \in \Sigma$ (3)

A language L_d recognized by a DFA is defined as:

$$L_d = \{ w \in \Sigma^* \mid \delta_d^*(q_0, w) \in F \}$$

We say that $A = (Q, \Sigma, \delta_d, P, q_0, F)$ is a Probabilistic Deterministic Finite Automaton (*PDFA*) if and

only if, $A = (Q, \Sigma, \delta_d, q_0, F)$ is a DFA and $P : Q \times \Sigma \times Q \rightarrow [0, 1]$ is a probabilistic function such that :

(1)
$$\forall (q, a, q') \in Q \times \Sigma \times Q, \text{ if } \delta_d(q, a) \neq q'$$

then $P(q, a, q') = 0$
(2) $\forall q \in Q, \sum_{(a,q') \in \Sigma \times Q} P(q, a, q') = 1$

When the future states of a PDFA depend only on the present state and are independent of the sequence of events that preceded it, the Markov property holds (Norris, 1998). In other words, given the present, the past has no additional information to offer about the future. A PDFA with the Markov property is called a *Markov chain*.

The use of Markov chains provides a powerful tool for modeling the probabilistic behavior of patients while playing serious games. In the sequel, we present probabilistic model-checking, a formal technique that can be used to automatically analyze the probabilistic behavior of patients and verify the validity of specific properties while they are playing.

3.2 Temporal Logics

Temporal logic formulae describe the dynamical evolution of a given system. The Computation Tree Logic CTL* (Clarke et al., 1999a) allows one to describe properties of computation trees. Its formulas are obtained by (repeatedly) applying Boolean connectives, path quantifiers, and state quantifiers to atomic formulas. The path quantifier A (resp., E) can be used to state that all paths (resp., some path) starting from a given state have some property. The state quantifiers are the following ones. The next time operator X can be used to impose that a property holds at the next state of a path. The operator F (sometimes in the future) requires that a property holds at some state on the path. The operator G (always in the future) specifies that a property is true at every state on the path. The until binary operator U holds if there is a state on the path where the second of its argument properties holds, and, at every preceding state on the path, the first of its two argument properties holds. The Branching Time Logic CTL (Clarke et al., 1986) is a fragment of CTL* that allows quantification over the paths starting from a given state. Unlike CTL*, it constrains every state quantifier to be immediately preceded by a path quantifier. The Linear Time Logic LTL (Sistla and Clarke, 1985) is another known fragment of CTL* where one may only describe events along a single computation path. Its formulas are of the form $A\phi$, where ϕ does not contain path quantifiers, but it allows the nesting of state quantifiers.

CTL and LTL have a non-empty intersection. As an example, the property A $((x=1) \cup (y=3))$ belongs both to CTL and LTL. It holds in a state if, for each path starting from the state, *x* equals 1 until the moment when *y* equals 3. There exists several tools to automatically check whether a model verifies a given CTL or LTL formula, e.g., NuSMV (Cimatti et al., 1999) and SPIN (Holzmann, 2004).

The dynamics of probabilistic models can be specified using Probabilistic Computation Tree Logic (PCTL) (Hansson and Jonsson, 1994), which extends CTL by replacing the classical CTL path quantifiers A and E with probabilities. Thus, instead of saying that some property holds for all paths or for some paths, we say that a property holds for a certain fraction of the paths. The most important operator in PCTL is P, which allows to reason about the probability of event occurrences. The property **P** bound [prop] is true in a state s of a model if the probability that the property *prop* is satisfied by the paths from state s satisfies the bound bound. As an example, the PCTL property P = 0.4 [X (y = 2)]holds in a state if the probability that y = 2 is true in the next state equals 0.4. To compute the likelihood that some behavior of a model happens, the P operator can take the form P=?. As an example, the property P = ? [G (y = 1)] assesses the probability that y always equals 1. Several model-checkers allow to automatically check whether a given probabilistic model satisfies a given PCTL formula, or to automatically compute the probability for a given formula to be satisfied. State-of-the-art probabilistic model checkers are PRISM (Kwiatkowska et al., 2011), UP-PAAL (Behrmann et al., 2004), STORM (Dehnert et al., 2017), and PAT (Sun et al., 2009).

4 THE FRAMEWORK: A DOXASTIC META-AUTOMATON

In this section, we present the framework while instantiating it to our medical application. First, we define three PDFA to model the behaviour of three classes of patients while playing the game. Second, we define what we call a "meta-automaton", whose aim is to model the protocol. Such a meta-automaton suggests to practitioners to which class the patient is supposed to belong and helps the patient to training. After a game session, based on patient performances, the meta-automaton suggests a class the patient is supposed to belong and the next game session for the patient. For the sake of compactness, in

4.1 Three PDFA for Three Classes of Patients

In the case study concerning the Match Items game (see Section 2), we consider the following finite alphabet $\Sigma_g = \{\alpha, \beta, \gamma, \theta\}$, which represents the different possible actions defined in Section 2. As an example, a word as $w_1 = \alpha\beta\beta\alpha \in \Sigma_g^*$ signifies that the user first does action α , then β , then β , and finally α .

In our medical application, we consider three deteministic automata $Q = \{A_{\mathfrak{h}}, A_{\mathfrak{m}}, A_{\mathfrak{M}}\}$ since there are three classes of Alzheimer patients. Each automaton represents the activity of a class of patients while playing the serious game. In order to validate or reject one hypothesis about a state of a patient, we consider that $A_{\mathfrak{h}}$ represents the test for $\mathfrak{h}, A_{\mathfrak{m}}$ for \mathfrak{m} , and $A_{\mathfrak{M}}$ for \mathfrak{M} . In these automata we consider one initial state q_0 in which the user has to launch the game, and two final states: f_1 when the game is over, and f_2 when the user left the game before it was over. Let $F = \{f_1, f_2\}$ be the final states. We define the following three PDFAs: for all $x \in \{\mathfrak{M}, \mathfrak{m}, \mathfrak{h}\}, A_x = (Q_x \cup F, \Sigma_g, \delta_x, P_x, q_x, F)$, where L_x is the language recognized by A_x .

For the Match Items game, clinicians already provided us with (a priori) empirical probabilities on the different actions to be performed depending on the different classes. To obtain these probabilities, 10 clinicians-including medical doctors, nurses, and psychologists who are familiar with patients' performance while playing the game-each filled out a questionnaire. The questionnaire included questions such as: "For a patient in a given class, what are the chances of selecting the correct image at each step?" and "For a patient in a given class, what are the chances of not interacting with the game for at least 10 seconds?" The responses were given as numbers from 0 to 10. Table 1 represents the average probability given by 10 clinicians. We assume that, for each of the three automata, the probabilistic function follows this table, e.g., $P_{\mathfrak{h}}$ is such that for all $\begin{array}{l} (q,q') \in (\mathcal{Q}_{\mathfrak{h}} \setminus F)^2, P_{\mathfrak{h}}(q, \alpha, q') = 0.8, P_{\mathfrak{h}}(q, \beta, q') = \\ 0.1, P_{\mathfrak{h}}(q, \gamma, q') = 0.05, P_{\mathfrak{h}}(q, \theta, q') = 0.05. \end{array}$

For the sake of clarity, Figure 2 illustrates the automaton $A_{\mathfrak{h}}$ in a simplified manner by depicting each transition between states for each action. The initial state is denoted as q_0 , and there are two final states: f_1 signifies the normal end of the game, while f_2 indicates the user left the game. Furthermore, let us notice that $A_{\mathfrak{h}}$ has the Markov chain property as $A_{\mathfrak{m}}$ and $A_{\mathfrak{M}}$.

Table 1: Average probability given by 10 clinicians for each class of patients.

Action	h	m	M
α	0.84	0.5	0.17
β	0.11	0.30	0.58
γ	0.0499	0.1999	0.24
θ	0.0001	0.0001	0.01



Figure 2: Automaton $A_{\mathfrak{h}}$ describing the expected behaviour of healthy people while playing the Match Items serious game.

4.2 An Experimental Protocol as A Meta-Automaton

The experimental protocol aims to monitor and assess the patient's condition. This protocol involves organizing various tests within a meta-automaton. After each test, a *belief function* provides a confident score about the class the patient belongs to. Thanks to this score, the meta-automaton informs the decision-making process for the next test to apply.

Definition 2. A Doxastic Deterministic Finite Meta-Automaton (*DDFMA*) is a 7-tuple $\mathcal{A} = (Q, \Sigma_O, \Sigma_A, \delta, \{\mathcal{B}_q\}_{q \in Q}, q_0, F)$ where¹:

- *Q* is a finite set of PDFA.
- $\Sigma_Q = \bigcup_{q \in Q} \Sigma_q$ is a finite alphabet of input symbols recognized by all automata in Q.
- Σ_A is a finite alphabet of input symbols.
- $\delta: Q \times \Sigma_A \to Q$ is the transition function.

¹To distinguish between PDFA and DDFMA, we denote the latter with a calligraphic letter \mathcal{A}

• $\forall q \in Q, \mathcal{B}_q : L_q \times \Sigma_A \times Q \rightarrow [0,1]$ is a belief function that represents from an automaton q, given an accepted word $w \in L_q$, the practitioner belief for the patient to be in the class associated with the next automaton q'. Furthermore \mathcal{B}_q is such that:

(1)
$$\forall q \in Q, \forall (a,q') \in \Sigma_A \times Q, \text{ if } \delta(q,a) \neq q'$$

then $\forall w \in L_q, \mathcal{B}_q(w,a,q') = 0$
(2) $\forall q \in Q, \forall w \in L_q, \sum_{(a,q') \in \Sigma_A \times Q} \mathcal{B}_q(w,a,q') = 1$

• $q_0 \in Q$ is the initial state, i.e., the first automaton.

• *F* is the set of final states.

We define δ^* the transition function w.r.t. δ which operates on a set of words defined on the alphabet $\Sigma = \Sigma_Q \cup \Sigma_A$, and for all $q \in Q$ and $x \in \Sigma^*$:

$$\begin{split} \delta^* : & Q \times \Sigma^* \to Q \\ \delta^*(q, ()) &= q \\ \delta^*(q, xa) &= \delta(\delta^*(q, x), a) \quad \text{if } a \in \Sigma_A \\ \delta^*(q, xa) &= \delta^*(q, x) \quad \text{if } a \in \Sigma_Q \end{split}$$

A language $\mathcal{L}_{\mathcal{A}}$ recognized by a DDFMA is defined as:

$$\mathcal{L}_{\mathcal{A}} = \{ w \in \Sigma^* \mid \delta^*(q_0, w) \in F \}$$

Let us notice that the constraints (1) and (2) for the function \mathcal{B} translate that (1) if a transition does not exist in the meta-automaton, then the practitioner belief for the patient to be in the class associated with this transition cannot be different from 0, and (2) the sum of all beliefs associated with other transitions is equal to 1.

For our medical application, we consider the three classes $C = \{\mathfrak{h}, \mathfrak{m}, \mathfrak{M}\}$ and the DDFMA $\mathcal{A} = (Q, \Sigma_C, \Sigma_A, \delta_{exp}, P_{exp}, h, F)$, where

- Q = {A_b, A_m, A_m} is the set of states, which is composed by the three automata;
- $\Sigma_{\mathcal{C}} = \Sigma_{\mathfrak{h}} \cup \Sigma_{\mathfrak{m}} \cup \Sigma_{\mathfrak{M}}$ is the set of all symbols recognized by the automaton;
- $\Sigma_A = C$: a symbol corresponds to a class that will be tested in the next state of the automaton
- δ is given in Figure 3;
- for all $A_q \in Q$, for a word $w \in L_q$ recognized by A_q , for all $O \in \Sigma_{exp}$, the belief function \mathcal{B}_{exp} is

Table 2: Factors of \mathcal{B}_q , for all $q \in Q$.

q	$A_{\mathfrak{h}}$	$A_{\mathfrak{m}}$	$A_{\mathfrak{M}}$
$\Delta_q^{\mathfrak{h}}$	2.016	0	0
$a_q^{\mathfrak{h}}$	0.5	1	-0.1
$b_q^{\mathfrak{h}}$	-3.6	1.2	0.1
$c_q^{\mathfrak{h}}$	1	0.3	0.1
$d_q^{\mathfrak{h}}$	0.7	2.2	-2.4
$v_q^{\mathfrak{h}}$	0	0	1
$z_q^{\mathfrak{h}}$	0	-1	1.1
$\Delta_q^{\mathfrak{M}}$	6.256	0	3.769
$a_q^{\mathfrak{M}}$	2.4	-1	-6.6
$b_q^{\mathfrak{M}}$	2.1	1	0.4
$c_q^{\mathfrak{M}}$	-1	1.4	0.01
$d_q^{\mathfrak{M}}$	0.24	0.8	1.6
$v_q^{\mathfrak{M}}$	1	1	1
1 977		62	04

such that for all $q \in Q$, $w \in L_q$, $x \in \Sigma_A$:



 $\mathcal{B}_q(w,\mathfrak{m},A_\mathfrak{m}) = 1 - \mathcal{B}_q(w,\mathfrak{h},A_\mathfrak{h}) - \mathcal{B}_q(w,\mathfrak{M},A_\mathfrak{M})$

- *h* ∈ *Q* is the initial state, that corresponds to the initial hypothesis on the class in which the patient belongs to.
- *F* = *Q*: we consider each state to be an accepting state.

We defined a function $\Delta : L_q \to \mathbb{R}$ representing a *confidence score* for the patient not to belong to the class tested by the automata \mathcal{A}_q . The domain of this function is defined on [0,m], with $m \in \mathbb{R}$ which is the max score (here m = 10). 0 indicates that the patient did not do any mistake, while *m* indicates that the patient did 100% mistakes or left the game, i.e., she did the θ action. We consider a factor for each action: $k_\beta = 1$ and $k_\alpha = 1$ since we aim to count the number of mistakes. However, since we have also θ and γ as possible, to count these actions we associate a factor

to each one. We consider a factor $k_{\gamma} = 0.2$ indicating that five γ are equivalent to do one mistake. Finally, $k_{\theta} = 1 \times 10^9$: if the patient leaves the game, we consider it as 100% mistakes. Thus, Δ computes the number of mistakes (i.e., β actions) and weights the number of waiting actions (i.e., γ actions). To do so, we introduce a function for each $x \in \Sigma_C$, $|.|_x : \Sigma_C^* \to \mathbb{N}$ that counts the number of *x* in a word $w \in \Sigma_C^*$ and we note $|w|_x$ this number. The formal definition is the following one, for all words $w \in \Sigma_C^*$:

$$\Delta(w) = \begin{cases} m & \text{if } \theta \in w \\ \frac{m \times (k_{\beta} \times |w|_{\beta} + k_{\gamma} \times |w|_{\gamma} + k_{\theta} \times |w|_{\theta})}{k_{\alpha} \times |w|_{\alpha} + k_{\beta} \times |w|_{\beta} + k_{\gamma} \times |w|_{\gamma} + k_{\theta} \times |w|_{\theta}} & \text{otherwise} \end{cases}$$

In Table 2, we give the factors defined for each \mathcal{B}_q . These factors considered in each \mathcal{B}_q have been computed based on Table 1 so that the output class from a test corresponds to the class given by this table. Figure 4, 5 and 6 represent the different belief functions for each automaton $A_{\mathfrak{h}}$, $A_{\mathfrak{m}}$ and $A_{\mathfrak{M}}$, respectively. Thus, to be considered in h, the patient has to produce at least 80% of good answers (the value given in Table 1 is 0.84). So, the delta is computed as 1 - 0.84 = 0.16. This value corresponds to the abscissa of the first intersection between the green and the black curves in Figure 5. The green curve represents the variation of the belief $\mathcal{B}_{A_{\mathfrak{m}}}(w,\mathfrak{h},A_{\mathfrak{h}})$, i.e., when we believe the patient could be in h. The black curve represents the variation of the belief $\mathcal{B}_{A_{\mathfrak{m}}}(w,\mathfrak{m},A_{\mathfrak{m}})$, i.e., when we believe the patient could be in m. This intersection between the green curve and the black one is approximately for a $\Delta(w) = 0.16$, it represents the moment where the practitioner considers the patient in m and should do the test again. After 75% mistakes, we believe that the patient could be in \mathfrak{M} and this is represented by the intersection between the black curve and the purple curve. The purple curve represents the variation $\mathcal{B}_{A_{\mathfrak{m}}}(w,\mathfrak{M},A_{\mathfrak{M}})$. Figure 4 represents the change of belief in function of the result w of the patient when the test $A_{\mathfrak{h}}$ has been done. The green curve represents $\mathcal{B}_{A_{\mathfrak{h}}}(w,\mathfrak{h},A_{\mathfrak{h}})$, the black curve $\mathcal{B}_{A_{\mathfrak{h}}}(w,\mathfrak{m},A_{\mathfrak{m}})$, and the purple curve $\mathcal{B}_{A_{\mathfrak{h}}}(w,\mathfrak{M},A_{\mathfrak{M}})$. After more than 20% mistakes, the belief to be in h decreases a lot to reach the intersection with the belief to be in m. Let us notice that, contrary to the green curve in Figure 5, the belief to be in \mathfrak{h} is slightly shifted to the right, since we consider the test $A_{\mathfrak{h}}$ to be harder, and therefore tolerate the patient to make a few more errors. After 80% mistakes, we start to believe the patient could belong to the class \mathfrak{M} and we would have to do the \mathfrak{M} transition. Figure 6 represents the change of belief in function of the result w of the patient when the test $A_{\mathfrak{M}}$ has been done. We do not tolerate more than 13% mistakes to be in class h. This is depicted by



Figure 3: Automaton of the experimental protocol.

the intersection point between the green curve and the black curve. Let us notice that, since we applied the easiest test $A_{\mathfrak{M}}$, we cannot fully believe the patient belongs to the class \mathfrak{h} even if the patient does no mistake. This is represented by the fact that, for $\Delta(w) = 0$, $\mathcal{B}_{A_{\mathfrak{M}}}(w, \mathfrak{h}, A_{\mathfrak{h}}) = 0.8$.

Figure 3 represents the experimental protocol as a DDFMA. The transitions represent the next test to apply. To know the next transition, i.e., the next test to apply, we compute a score from the current test with the Δ function. Then, this score is considered as an input for the belief function \mathcal{B} . This function represents the beliefs of a clinician or a set of clinicians about the results obtained from the test.

The first test to apply, i.e., the initial state, depends on the initial hypothesis h we consider. Then, we start by the corresponding test q to validate or reject h and get a score; then this score is considered as an input for the function \mathcal{B}_q that returns the next test to apply.

Let us admit the initial hypothesis is $h = \mathfrak{h}$, i.e., the patient is assumed to be in the class \mathfrak{h} . We then apply the test $A_{\mathfrak{h}}$: after the test is over, we have evaluated if the patient is in \mathfrak{h} or if we have to reject the hypothesis. To compute the score we consider the Δ function previously defined, which computes the number of actions that are not the right action (i.e., α).



Figure 4: Evolution of $\mathcal{B}_{A_{\mathfrak{h}}}$ in function of $\Delta(w)$.



Figure 5: Evolution of $\mathcal{B}_{A_{\mathfrak{m}}}$ in function of $\Delta(w)$.



Figure 6: Evolution of $\mathcal{B}_{A_{\mathfrak{M}}}$ in function of $\Delta(w)$.

If we could not reject the hypothesis, the evaluation is \mathfrak{h} , which means we confirm the hypothesis as acceptable w.r.t. the test. If the result is \mathfrak{m} , it is highly possible that the patient is in \mathfrak{m} . The change of class will be suggested to practitioners

5 FORMAL VALIDATION

In this section, we apply the DDFMA framework to the serious game introduced in Section 2, employing a PDFA-based representation to describe the expected behavior of each class of patients while playing the game. Firstly, we provide examples of PCTL properties to test the aptitude of the models to dispaly some interesting behaviours. Secondly, we apply LTL to the execution traces of the meta-automaton in order to define stopping conditions for the protocol. Thirdly, we give concrete decision-making capabilities of the meta-automaton and show some properties holding in our medical application, including examples of acceptable and unacceptable traces.

5.1 Using PCTL to Assess Model Probabilistic Behavior

The behaviour of the game for each class of patients is modelled with a PDFA that has the Markov Chain Property. In order to apply model-checking to our application, we define *PDFA models* :

Definition 3. Let Atm be a set of symbols called atomic propositions. We call $M = (A, \mathcal{V})$ a PDFA

model iff A is PDFA and \mathcal{V} : Atm $\rightarrow 2^Q$ is a valuation function where Q is the set of states of A. We call $\mathcal{M} = (\mathcal{A}, \mathcal{V})$ a DDFMA model iff \mathcal{A} is a DDFMA and \mathcal{V} : Atm $\rightarrow 2^Q$ is a valuation function where Q is the set of states of \mathcal{A} .

We define the following PDFA models: $M_{\mathfrak{h}} = (A_{\mathfrak{h}}, \mathcal{V}_{\mathfrak{h}}), M_{\mathfrak{m}} = (A_{\mathfrak{m}}, \mathcal{V}_{\mathfrak{m}})$ and $M_{\mathfrak{M}} = (A_{\mathfrak{M}}, \mathcal{V}_{\mathfrak{M}})$, for each automaton $A_{\mathfrak{h}}, A_{\mathfrak{m}}$ and $A_{\mathfrak{M}}$. We consider each action to be represented as an atomic proposition, denoted by $Atm = \{ \alpha, \beta, \gamma, \theta \}$, verified in the accessible state with the corresponding transition. Given a PDFA model $M \in \{M_{\mathfrak{h}}, M_{\mathfrak{m}}, M_{\mathfrak{M}}\}$, if we access a state *s* with the action β , then the atom β is verified in *s*, denoted by $M, s \models \beta$. Each final state f_1 (resp. f_2) has a corresponding atom a_1 (resp. a_2), i.e., $M, f_1 \models a_1$ and $M, f_2 \models a_2$. Given a model M_c , with $c \in \{\mathfrak{h}, \mathfrak{m}, \mathfrak{M}\}$, and a state *s* in M_c , we check if the state *s* in M_c satisfies the following PCTL useful property examples.

Reachability of a Final State. Is the probability of reaching a final state equal to 1? The PCTL formula to test is P = 1 [F (a1 or a2)].

Reachability of a State Without Violating Some Constraints. Is the probability to reach a certain state *d* without violating some constraints *c* greater than 0? The PCTL formula to test is P > 0 [c U d].

Reachability of a State Without Passing From Another. Which is the probability to reach a state d without passing from a state b? The PCTL formula to test is P = ?[(not b) U d].

5.2 LTL Analysis of Patient Experiences: Protocol Stop Conditions

In this section we consider the DDFMA \mathcal{A} defined in Section 4.2 and we note its corresponding DDFMA model $\mathcal{M} = (\mathcal{A}, \mathcal{V})$. The valuation function \mathcal{V} is such that each transition $t \in C$ (where $C = \{\mathfrak{h}, \mathfrak{m}, \mathfrak{M}\}$) is associated with an atomic proposition verified in a state $q \in Q$. For instance, if we do a transition \mathfrak{m} to reach a state q_m , then the corresponding atomic proposition \mathfrak{m} is verified in q_m , i.e., $\mathcal{M}, q_m \models \mathfrak{m}$.

5.2.1 Oscillating Behaviour

If we detect a trace $\tau = (q_0, q_1, \dots, q_n) \in Q^{n+1}$ showing that the patient alternated between different classes, then we want to stop the protocol. Hereafter we provide some examples of traces we aim to detect by providing a regular expression: $(\mathfrak{m} \cdot * \mathfrak{M})^4$ (resp. $(\mathfrak{h} \cdot * \mathfrak{M})^4$, $(\mathfrak{h} \cdot * \mathfrak{m})^4$), i.e., the patient oscillates between \mathfrak{m} and \mathfrak{M} (resp. \mathfrak{h} and \mathfrak{M} , \mathfrak{h} and \mathfrak{m}) four consecutive times. Here "." denotes any character of C and * is the quantification "zero or more occurrences".

Let q_0 be the first state of a trace τ . q_0 verifies the stop condition if it satisfies the following LTL formula: \mathfrak{m} and F (\mathfrak{M} and F (\mathfrak{m} and F (\mathfrak{M} and F \mathfrak{m} and F (\mathfrak{M} and F (\mathfrak{m} and F (\mathfrak{M})))))).

5.2.2 The Permitted Number of Tests Has Been Exceeded

We consider that a patient will not do more than 10 tests. Given an initial state q_0 for a trace τ , q_0 verifies this stop condition if it satisfies the LTL formula: X^{10} true and not X^{11} false, where X^n is a shortcut for X...X with *n* occurrences of *X*.

5.2.3 Reaching A Steady-State Condition

If a patient stays in the same class for at least 3 tests, we stop the test. For instance, given a trace $\tau = (q_0, \dots, q_n)$, a state q_{n-2} verifies this stop condition for the healthy class if it verifies the LTL formula: \mathfrak{h} and X (\mathfrak{h} and X \mathfrak{h}).

5.3 Acceptable and Unacceptable Traces

Some traces are unacceptable. For instance, if the patient gives wrong answers but he is considered as \mathfrak{h} , then we do not want her to stay in the class \mathfrak{h} . In the same way, if the patient has a very positive outcome but is considered as \mathfrak{m} , then we would like to check in a more accurate way if she could be classed as \mathfrak{h} . But we allow a mitigated outcome to make the patient stay in the same state. In the following, given a DDFMA model $\mathcal{M} = (\mathcal{A}, \mathcal{V})$, we formally show that our belief functions are compatible with these properties.

5.3.1 An Extremely Poor Outcome In A_β Cannot Classify a Patient as h

A trace in the DDFMA looks like $\tau = h\beta\gamma * \beta\gamma * \betah$. This trace recognized by a DDFMA means that we initialize the DDFMA in the state $A_{\mathfrak{h}}$ to make the patient perform this test. In this test, the word recognized by the automaton $A_{\mathfrak{h}}$ is $\beta\gamma * \beta\gamma * \beta$. This trace signifies that the patient only does wrong answers β , and may wait between actions, i.e., $\gamma *$.

We formally show that, according to the DDFMA model defined in Section 4.2, this is impossible. Let consider a word $w = \beta \gamma \gamma \beta \beta \beta \gamma \beta^5 \gamma \beta$ recognized by the

test $A_{\mathfrak{h}}$, i.e., this is the execution of actions of the patient. Here obviously $\Delta(w) = m = 10$. Thus according to Figure 4, the belief $\mathcal{B}_{A_{\mathfrak{h}}}(w,\mathfrak{M},A_{\mathfrak{M}}) = 0.72$ and $\mathcal{B}_{A_{\mathfrak{h}}}(w,\mathfrak{m},A_{\mathfrak{m}}) = 0.28$. Since the belief of belonging to \mathfrak{M} is stronger than belonging to \mathfrak{m} , the automaton should propose a transition towards $A_{\mathfrak{M}}$ and so τ cannot be verified in this configuration for our DDFMA.

5.3.2 A Very Good Outcomes In A_m Can Classify a Patient as h

Let consider that a patient does 100% of α in the automaton $A_{\mathfrak{M}}$. A word recognized by $A_{\mathfrak{M}}$ could be $w = \alpha^{10}$. Thus, $\Delta(w) = 0$ and $\mathcal{B}_{A_{\mathfrak{M}}}(w, \mathfrak{h}, A_{\mathfrak{h}}) = 0.75$ if we look at Figure 6. In such situation an acceptable trace would be $\mathfrak{M}w\mathfrak{h}$.

5.3.3 A Medium Outcomes In A_m Can Classify A Patient As m

A medium outcome in $A_{\mathfrak{m}}$ corresponds to all words w recognized by $A_{\mathfrak{m}}$ such that $\Delta(w) \in [1.364, 7.45]$. For instance a word $w = (\alpha\beta)^5$ have a $\Delta(w) = 5$ and so $\mathcal{B}_{A_{\mathfrak{m}}}(w, \mathfrak{m}, A_{\mathfrak{m}}) = 0.8765$. In such situation an acceptable trace by the DDFMA would be $\mathfrak{m}w\mathfrak{m}$.

6 CONCLUSION AND FUTURE WORK

Artificial intelligence is increasingly being embraced for medical applications, revolutionizing healthcare with its innovative capabilities. In this work we proposed a framework to model the behaviour of Alzheimer patients while playing serious games over several game sessions. The utility of the proposed methodology is twofold: (i) complement the diagnosis of medical doctors thanks to our formal analysis of the patient performances while playing serious games; (ii) help practitioners by dynamically suggesting the next step, i.e., game or difficulty level, to propose to the patient after each game session. One of the main strengths of the proposed methodology is to be very general, and thus suited to be exploited for other kinds of medical protocols, e.g., diagnosis and training of children affected by attention disorders.

The approach we propose in this article was devised after numerous discussions with the clinicians of Claude Pompidou Institute, Nice, France. Before implementing the methodology, we need a theoretical model to present to practitioners. As a next step, we intend to implement all the models in the Probablistic Model Checker PRISM (Kwiatkowska et al., 2011) and to develop an automated tool at the clinicians' disposal. For the sake of simplicity, we provided only one game and one difficulty level. Actually, the protocol can include the possibility to alternate different games, and different difficulty levels for each game. We dispose of several games targeting different cognitive functions which may be affected in Alzheimer patients, such as memory or inhibitory control. Finally, the formal approach presented in this work opens an avenue in automating medical protocols and allows to dynamically keep practitioners aware about the possible evolution on the patient disease severity level. The meta-automaton dynamically evaluates the confidence levels of practitioners regarding a patient's diagnosis by analyzing the consistency and progression of their in-game performance across sessions. This adaptive mechanism not only enhances diagnostic accuracy over time but also personalizes the therapeutic aspect of the games. By automatically adjusting the complexity and nature of subsequent game sessions based on prior performance, our protocol ensures that each patient receives tailored cognitive training. This personalized approach maximizes the therapeutic benefits, making serious games a powerful dual-purpose tool for both diagnosing and training patients with neurodegenerative diseases.

ACKNOWLEDGEMENTS

We thank Valeria Manera from Institut Claude Pompidou, Nice, France, for her valuable insights and fruitful discussions on Alzheimer's patients.

REFERENCES

- Alvarez, J., Jessel, J.-P., and Méthel, G. (2007). Pbl and serious game. In Moore, G. and Hernandez, A., editors, *7th ALE International Workshop*, Toulouse.
- American Psychiatric Association (2013). *Diagnostic and Statistical Manual of Mental Disorders*. Number 21. American Psychiatric Association, 5th edition.
- Anguera, J. A., Boccanfuso, J., Rintoul, J. L., Al-Hashimi, O., Faraji, F., Janowich, J., Kong, E., Larraburo, Y., Rolle, C., Johnston, E., and Gazzaley, A. (2013). Video game training enhances cognitive control in older adults. *Nature*, 501(7465):97–101.
- Behrmann, G., David, A., and Larsen, K. G. (2004). A Tutorial on UPPAAL. In Formal Methods for the Design of Real-Time Systems: 4th International School on Formal Methods for the Design of Computer, Communication, and Software Systems, SFM-RT 2004, number 3185 in LNCS, pages 200–236. Springer.
- Cimatti, A., Clarke, E. M., Giunchiglia, F., and Roveri, M. (1999). NUSMV: A new symbolic model verifier. In

Computer Aided Verification, 11th International Conference, CAV '99, Trento, Italy, July 6-10, 1999, Proceedings, pages 495–499.

- Clarke, E., Grumberg, O., and Peled, D. (1999a). *Model Checking*. MIT Press.
- Clarke, E. M., Emerson, E. A., and Sistla, A. P. (1986). Automatic verification of finite-state concurrent systems using temporal logic specifications. ACM Trans. Program. Lang. Syst., 8(2):244–263.
- Clarke, E. M., Grumberg, O., and Peled, D. A. (1999b). *Model Checking*. MIT Press.
- De Maria, E., L'Yvonnet, T., Moisan, S., and Rigault, J. (2019). Probabilistic activity recognition for serious games with applications in medicine. In Hasan, O. and Mallet, F., editors, Formal Techniques for Safety-Critical Systems - 7th International Workshop, FTSCS 2019, Shenzhen, China, November 9, 2019, Revised Selected Papers, volume 1165 of Communications in Computer and Information Science, pages 106–124. Springer.
- Dehnert, C., Junges, S., Katoen, J.-P., and Volk, M. (2017). A storm is coming: A modern probabilistic model checker. In *Computer Aided Verification*, pages 592– 600. Springer.
- Hansson, H. and Jonsson, B. (1994). A logic for reasoning about time and reliability. *Formal aspects of computing*, 6(5):512–535. Springer.
- Holzmann, G. J. (2004). The SPIN Model Checker primer and reference manual. Addison-Wesley.
- Kathryn, J. M., Ewbank, D. C., Wang, H., and Trojanowski, J. Q. (2014). The impact of exercise, cognitive activities, and socialization on cognitive function: Results from the national long-term care survey. *American Journal of Alzheimer's Disease & Other Dementias*, 29.
- Kato, P. and Klerk, S. (2017). Serious games for assessment: Welcome to the jungle. *Journal of Applied Testing Technology*, 18.
- Kwiatkowska, M., Norman, G., and Parker, D. (2011). PRISM 4.0: Verification of Probabilistic Real-time Systems. In Proc. 23rd Int. Conf. on Computer Aided Verification (CAV'11), volume 6806, pages 585–591. Springer.
- L'Yvonnet, T., Maria, E. D., Moisan, S., and Rigault, J. (2021). Probabilistic model checking for human activity recognition in medical serious games. *Sci. Comput. Program.*, 206:102629.
- Melenhorst, A.-S., Rogers, W. A., and Bouwhuis, D. G. (2006). Older adults' motivated choice for technological innovation: Evidence for benefit-driven selectivity. *Psychology and Aging*, 21.
- Norris, J. R. (1998). Markov chains. Number 2.
- Philippe, R., König, A., Amieva, H., Andrieu, S., Bremond, F., Bullock, R., Ceccaldi, M., Dubois, B., Gauthier, S., Kenigsberg, P.-A., Nave, S., Orgogozo, J. M., Piano, J., Benoit, M., Touchon, J., Vellas, B., Yesavage, J., and Manera, V. (2014). Recommendations for the use of serious games in people with alzheimer's disease, related disorders and frailty. *Frontiers in Aging Neuroscience*, 6:54.

- Rabin, M. O. (1963). Probabilistic automata. *Information* and control, 6(3):230–245.
- Sistla, A. P. and Clarke, E. M. (1985). The complexity of propositional linear temporal logics. *J. ACM*, 32(3):733–749.
- Sun, J., Liu, Y., Dong, J. S., and Pang, J. (2009). PAT: Towards Flexible Verification under Fairness. In 21th International Conference on Computer Aided Verification, volume 5643, pages 709–714. Springer.
- Tong, T., Chignell, M., Tierney, M. C., and Lee, J. (2016). A serious game for clinical assessment of cognitive status: Validation study. *JMIR Serious Games*, 4(1):e7.
- Tran, M. K. P., Brémond, F., and Robert, P. (2015). Assistance for Older Adults in Serious Game Using an Interactive System. In 4th Int. Conf. on Games and Learning Alliance (GALA), pages 286–291. Springer.
- Vanessa, V., Wyss, P., Rampa, L., Mitache, A. V., Müri, R. M., Mosimann, U. P., and Nef, T. (2017). Evaluation of a novel serious game based assessment tool for patients with alzheimer's disease. *PLOS ONE*, 12.