Supplementary Material: Leveraging Large Language Models for Personalized Parkinson's Disease Treatment

In this supplementary material, we present details that out paper does not present due to the space limitations. We refer more information about the full-text prompts, case studies, and the code implementation to the website: $\frac{https:}{github.com/360ZMEM/PD-LLM}.$

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A Additional Statistical Information of the PPMI Dataset

In this section, we summarize some statistical details regarding clinical visits from the PPMI dataset.

When filtering the dataset, to ensure that the RL Baseline has sufficient historical reference data and to allow the LLM to perform a certain depth of simulation to complete the MCTS process, we selected patients with at least 7 valid clinical visit records, totaling 399 patients. Figure 1(a) illustrates the distribution of clinical visit counts among these 399 patients, with the maximum number of valid clinical visits recorded being 17, and the average and standard deviation being 12.0 and 2.9, respectively.

Additionally, we analyze the time intervals between clinical visits, with the distribution shown in Figure 1(b). The average time interval was 9.2 months, with a standard deviation of 5.3 months.

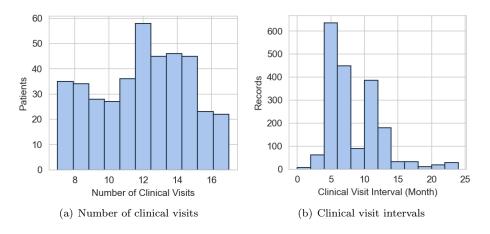


Figure 1: Statistical metrics of clinical visits in PPMI dataset.

B Details for MDP and Reinforcement Learning Baseline

B.1 Action Space

The action space of RL and our MCTS is discrete. It may not be too small, as some advantageous prescription combinations might be excluded; however, an excessively large action space will lead to a sharp increase in MCTS search costs. To achieve a balance of action space scale, we specify several basic single-dose strategies, which are then combined by a certain dose frequency. In this study, we define each action as a combination of the following six types:

• **L-dopa IR**: 100 mg, 250 mg.

• L-dopa CR: 100 mg, 200 mg.

• Rytary: 95 mg, 195 mg.

```
Table 1 Available prescriptions (73)
                                               [[0,100],[],[95,0]]
 [[100,100,250,250],[],[]]
 [[100,0],[0,100],[]]
                                               [[0,0,100,100],[],[95,95,0,0]]
 [[100,100,0,0],[0,0,100,100],[]]
                                               [[0,250],[],[95,0]]
                                               [[0,0,250,250],[],[95,95,0,0]]
 [[100,0],[0,200],[]]
 [[100,100,0,0],[0,0,200,200],[]]
                                               [[],[0,100],[95,0]]
 [[100,0],[],[0,95]]
                                               [[],[0,0,100,100],[95,95,0,0]]
 [[100,100,0,0],[],[0,0,95,95]]
                                               [[],[0,200],[95,0]]
                                               [[],[0,0,200,200],[95,95,0,0]]
 [[100,0],[],[0,195]]
                                               [[],[],[95,95,195,195]]
 [[100,100,0,0],[],[0,0,195,195]]
 [[250,250,100,100],[],[]]
                                               [[0,100],[],[195,0]]
                                               [[0,0,100,100],[],[195,195,0,0]]
 [[250,0],[0,100],[]]
                                               [[0,250],[],[195,0]]
 [[250,250,0,0],[0,0,100,100],[]]
 [[250,0],[0,200],[]]
                                               [[0,0,250,250],[],[195,195,0,0]]
                                               [[],[0,100],[195,0]]
 [[250,250,0,0],[0,0,200,200],[]]
 [[250,0],[],[0,95]]
                                               [[],[0,0,100,100],[195,195,0,0]]
                                               [[],[0,200],[195,0]]
 [[250,250,0,0],[],[0,0,95,95]]
 [[250,0],[],[0,195]]
                                               [[],[0,0,200,200],[195,195,0,0]]
                                               [[],[],[195,195,95,95]]
 [[250,250,0,0],[],[0,0,195,195]]
 [[0,100],[100,0],[]]
                                               [[100],[],[]]
 [[0,0,100,100],[100,100,0,0],[]]
                                               [[100,100],[],[]]
 [[0,250],[100,0],[]]
                                               [[100,100,100,100],[],[]]
 [[0,0,250,250],[100,100,0,0],[]]
                                               [[250],[],[]]
                                               [[250,250],[],[]]
 [[],[100,100,200,200],[]]
                                               [[250,250,250,250],[],[]]
 [[],[100,0],[0,95]]
 [[],[100,100,0,0],[0,0,95,95]]
                                               [[],[100],[]]
 [[],[100,0],[0,195]]
                                               [[],[100,100],[]]
                                               [[],[100,100,100,100],[]]
 [[],[100,100,0,0],[0,0,195,195]]
                                               [[],[200],[]]
 [[0,100],[200,0],[]]
 [[0,0,100,100],[200,200,0,0],[]]
                                               [[],[200,200],[]]
 [[0,250],[200,0],[]]
                                               [[],[200,200,200,200],[]]
 [[0,0,250,250],[200,200,0,0],[]]
                                               [[],[],[95]]
                                               [[],[],[95,95]]
 [[],[200,200,100,100],[]]
 [[],[200,0],[0,95]]
                                               [[],[],[95,95,95,95]]
 [[],[200,200,0,0],[0,0,95,95]]
                                               [[],[],[195]]
                                               [[],[],[195,195]]
 [[],[200,0],[0,195]]
 [[],[200,200,0,0],[0,0,195,195]]
                                               [[],[],[195,195,195,195]]
```

Given that patients may benefit from two-drug combinations, we define 73 available actions in this study. These actions are categorized based on the frequency of daily dosing, as follows:

- No medication (1 action).
- Once daily (6 actions).
- Twice daily, with equal doses if the same drug is administered twice in one day (30 actions).
- Four times daily, with identical types and dosages for the first two and last two doses (36 actions).

We also provide a list of all available actions in Table 1. We express them in a format as lists of three sublists in Python style, which include L-dopa IR, L-dopa CR, and Rytary medication data. The sublist shows the dosage and frequency of each drug in milligrams. For example, the treatment plan

[[200, 0], [], [0, 95]] indicates that for L-dopa IR and Rytary, two equally spaced doses are administered during the 17 waking hours each day, with the first dose being 200 mg of L-dopa IR. The last dose of Rytary is 95 mg.

Notice that LLM can directly output prescription plans and and will not be restricted by action space listed above.

B.2 RL Implementation Details

This section provides some implementation details of the RL baseline. The observation space of the RL model we employed is a subset of the observation space of the predictive model for MDS-UPDRS-III scores. Specifically, it includes the patient's vector of covariates (gender, age, years since PD diagnosis) and the patient's UPDRS scores over the past five clinical visits. Additionally, we added cluster index obtained through K-means clustering (with K=4) into the state representation to enhance the personalization of medication strategies for patients.

We use the reward function utilized by Baucum et al. The reward is determined based on the patient's current bradykinesia score b_t (L-dopa medication can reduce the slowing of movement) and dyskinesia score d_t (L-dopa medication may induce involuntary movements). The reward function is defined as follows:

$$r(s_t) = \mathbb{1}_{\tanh(b_t) < \bar{b}} - \alpha \cdot \mathbb{1}_{d_t > d}, \tag{1}$$

where \mathbb{F} denotes the indicator function. In other words, a reward is applied if $\tanh(b_t)$ is less than or equal to $\bar{b} = -0.279$ (corresponding to a unnormalized score of 25), and a penalty is applied if d_t exceeds $\bar{d} = -0.775$ (corresponding to a unnormalized score of 9). The coefficient α adjusts the weight of the dyskinesia penalty, and we set $\alpha = 0.5$ in this study.

Additionally, the RL method we used aligns with that of Baucum et al., namely a general approach of actor-critic implementation.

C Hyperparameter Choosing and Robustness Analysis

In this section, we select several representative parameters to investigate the impact of their variations, and evaluate the robustness of our LLM workflow.

Number of Similar Cases. In our paper, we default to providing two similar cases. We integrate varying numbers of similar cases into the prompt, and the results are shown in Table 2. As the number of similar cases increases, the performance of the LLM improves to some extent. However, when the number of similar cases exceeds four, the performance begins to decline. Additionally, a higher number of cases leads to an increase in the number of input tokens.

Number of MCTS Simulations. The number of simulations in MCTS affects the accuracy of the final evaluation of different medication decisions. However, setting this number too high can lead to a significant increase in computational costs. In our paper, we defaulted to 2500 simulation trials. Table 3 shows the performance as the number of trials is reduced (with the number of similar cases remaining at two). It can be seen that reducing the number of trials to one-fourth of the original does not significantly degrade performance, but further reduction leads to noticeable performance degradation.

Table 2: Key metrics of UPDRS Change (ΔU) with different number of similar cases.

Number of cases	$\mathbf{Average} \!\!\downarrow$	$\mathrm{Median}{\downarrow}$
1	1.96	2.26
2	1.83	1.86
3	1.82	1.91
4	1.75	1.79
5	1.80	1.80

Table 3: Key metrics of UPDRS Change (ΔU) with different number of MCTS simulations per records.

Number of simulations	$\mathbf{Average} \!\!\downarrow$	$\mathrm{Median}{\downarrow}$
2500	1.83	1.86
1200	1.89	2.02
600	2.12	2.35
300	2.40	2.55
150	2.66	2.80

D Full Prompt

In this section, we present the complete Prompt integrated with medical knowledge used in our experiments. Specifically, the complete Prompt includes a comprehensive explanation of the structured text, along with corresponding empirical descriptions, and provides a detailed introduction to the steps that the output should follow, ultimately generating structured medication information.

Thanks to the highly interpretable text, this part of the Prompt can be modified according to user needs. In our experiments, this part of the Prompt will be input as the system instruction, while the user input will consist of the patient's structured text information, similar case text information, and a recommended medication list. This means that the Prompt shown below does not need to change with variations in patient information.

```
You are a doctor specializing in Parkinson's disease treatment. We present
\hookrightarrow information about a given patient, along with some machine-learning
   (ML)-generated treatment plans for other similar cases. You need to analyze the
   patient information and similar cases, and subsequently provide the formatted
   prescription following format specified below.
## Explanations of related terms
1. UPDRS Score: The score of the Unified Parkinson Disease Rating Scale (UPDRS)
\hookrightarrow ranges from 0 to 53 (99th-percentile), with higher scores indicating more severe
   symptoms. Each adjacent score is evaluated approximately a year apart.
2. Nonlev: Nonlev is the sum of the daily doses of other non-levodopa drugs (e.g.,
→ dopamine, MAO-B inhibitors, and COMT inhibitors) prescribed by a physician and
  is measured as the daily equivalent of L-dopa in milligrams.
3. The medication history followed the same format as the given treatment plans for
\hookrightarrow Levodopa and equivalent Levodopa doses for Nonlev and was also prescribed
  approximately a year apart.
4. The treatment plan is a list of three sublists, which include L-dopa IR
   ("immediate release"), L-dopa CR ("controlled release"), and Rytary (which is an
```

extended-release form of L-dopa) medication data. These medications differ in how quickly they release L-dopa into patients' bloodstreams. The sublist shows the dosage and frequency of each drug in milligrams. For example, the treatment plan `[[200, 0, 0], [], [0, 95, 95]]` indicates that for L-dopa IR and Rytary, three equally spaced doses are administered during the 17 waking hours each day, with the first dose being 200 mg of L-dopa IR. The last two doses of Rytary are 95 mg each.

5. We use ML methods to obtain prescriptions that provide four alternatives

5. We use ML methods to obtain prescriptions that provide four alternatives

classified as recommended and not recommended, respectively. Specifically, each

alternative is accompanied by the predicted UPDRS score after 3 years as

predicted by the neural network.

```
6. Patients who have not been diagnosed with PD will be listed as `N/A` in the
    → `Years_sincePD` metric. These patients may be prescribed medication for
       prevention or may exhibit some motor abnormalities and disorders, the extent of
       which might be reflected in the UPDRS score.
12
   ## Factors to consider
13
   We present some of the main considerations when deciding which medication plan to
    → prescribe to a given patient, with the intention of illustrating how these
    aspects:
16
   **The Patient's Information: ** Age, gender, number of years since the diagnosis of
    \hookrightarrow PD, medication history, and history of disease progression obtained from the
    \hookrightarrow UPDRS score may significantly influence prescription preferences, including the
    \hookrightarrow suitability and incompatibility of certain drug combinations.
18
   **Similar Cases: ** We also provide several similar cases and the corresponding
    \hookrightarrow recommended treatment plans, specifically:
20
   - For each similar case, the patient's information is presented in the same format
    \,\,\,\,\,\,\,\,\, as the given patient. The information is generally similar to the patient being
    \,\,\hookrightarrow\,\, prescribed, but there are still some differences.
   - Several recommended and unrecommended treatment plans are given for each similar
    \hookrightarrow case, and UPDRS scores predicted by machine learning are also provided. These
    \rightarrow treatment plans should also be appropriate to a certain extent for the given
       patient. Specifically, these plans are judged not only by predicted UPDRS
    → scores, where differences greater than ±3 are generally considered significant,
    \,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\, but also by considering the potential differences in subsequent disease
    \rightarrow development these medication recommendations may cause and whether they align
    \hookrightarrow with the actual condition of the patient. After conducting a comprehensive
    analysis, the recommended plans for certain similar cases with higher priority
    \rightarrow may be selected as the basis for modification.
   - It should be noted that due to the differences that still exist in similar cases,
    → there will be variations in the outcome of treatment plan candidates, which
       should also be taken into account.
24
   **Final Output:** The prescription for the final output should strictly follow the
    \hookrightarrow format described below.
26
   ## Output Guide and Format
28
   Your final output should contain only a simple analysis of about 200 words, and a
    \,\,\,\,\,\,\,\,\,\, Python code block containing the final recommended prescription after an
    \ \hookrightarrow \ \ \text{additional explanation.}
   The brief analysis includes the following: (a) A brief analysis of Important
    → information about the patient that may affect the decision of the treatment; (b)
    \,\,\,\,\,\,\,\,\, A brief analysis of similar cases and an overall analysis of recommended
       treatment plans. (c) Based on the comprehensive analysis, indicate the final
    \rightarrow prescription strategy and whether it is modified based on a certain treatment
    \rightarrow plan from the similar cases. (d) What adjustments (alter the frequency/dosage,
    \,\,\,\,\,\,\,\,\,\,\,\,\, change the medicine type...) and medication strategies have been made to further
       improve the potential effect to lower the UPDRS score.
```

```
Then, based on the above analysis, the prescription which could maximize the
    \,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\, decrease of UPDRS score should be obtained from similar cases or information
       from the given patient. The prescription contains the frequency and dosage of
       L-dopa IR, L-dopa CR, and Rytary intake on a daily basis. You should output the
       additional textual summary to specifically specify the dose and frequency of
       each medicine, then output the corresponding formatted prescription in python
       code block, as we explained in the 4) of the section "Explanations of related
       terms".
34
   For example, for following **textual** explanation:
35
   - **Frequency:** 3 doses (2 vaild doses)
37
   - **Medicine and Dosage:** (1) Morning: 200mg L-dopa IR (2) Noon: N/A (3) Evening:
38

→ 95mg Rytary

   - **Additional comment:** ...
40
   The corresponding **formatted** output is:
41
   ```python
43
 [[200, 0, 0], [0, 0, 0], [0, 0, 95]]
44
45
 Note that treatment plans for the given patient include a range of possible
 → prescription. However, you should analyze the patient's information, evaluating
 possible differences and choosing the treatment that is most promising to reduce
 the patient's UPDRS in a long-term perspective.
 ## Notes
49
50
 - 1 dose / 2 doses (Morning, Evening) / 3 doses (Morning, Noon, Evening) / 4 doses
 → (Morning, Noon, Afternoon, Evening)
 - For non-levodopa (Nonlev) drugs, no advice is required, as this part is directly
 - To facilitate pattern matching, you should only output one python block in the
 _{\hookrightarrow} `### Formatted Treatment Plan` section.
 - For ML-predicted prescriptions, as the decision space is discrete and limited,
 \hookrightarrow there may be other doses or frequencies that work better but are not included in

→ the recommended prescription list.

 - The sample prescriptions given in the above (for example, `[[200, 0, 0], [], [0,
 _{\rightarrow} 95, 95]] \hat{} and \hat{} [[], [100, 100, 0, 0], [0, 0, 245, 245]] \hat{}) are only used to
 → illustrate the output format and meaning and **DO NOT** constitute any treatment
 recommendations. You should prescribe based on the actual situation.
56
 Also, we use a simple user input template as follows:
 ## Patient Information to be Analyzed
 The information of patient you need to analyze is:
4
 <patient_information>
```

## Relative cases

```
Several similar patient information given below, we adopt ML methods to predict the

→ recommended treatment plans, and obtain explanations and suggestions output by

→ LLMs.

11

2 <similar_cases>
13
```

#### E More Information of Case studies

In this section, we provide more detailed information from Section VI.B of the paper. The medications prescribed by RL and LLM during all clinical visits are shown in Table 4 and Table 5, and Figure 2 illustrates the daily dosage changes of different types of L-dopa over time.

To further elaborate on the source of the medication recommendations provided by LLM, we will present the specific inputs and LLM responses (with system instructions and patient information omitted) for PATNO 3500 during the early, middle, and late phases, and for PATNO 3507 during the early and late phases.

Table 4 RL and LLM's presciption for PATNO 3500 during the whole process			
RL	LLM		
[[],[],[95,95]] [[100,100,0,0],[0,0,100,100],[]] [[100,100,0,0],[0,0,100,100],[]] [[100,100,0,0],[0,0,100,100],[]] [[100,100,0,0],[0,0,100,100],[]] [[100,0],[0,100],[]] [[100,0],[0,100],[]] [[100,100,0,0],[0,0,100,100],[]] [[100,100,0,0],[0,0,100,100],[]]	[[200,0,0],[],[0,195,195]] [[250,0,0],[],[0,195,95]] [[200,0],[],[0,195,]] [[250,250,0,0],[],[0,0,95,95]] [[],[200,200,0],[]] [[250,250,0],[100,100,0],[0,0,195]] [[250,250,0,0],[0,0,200,200],[]] [[250,0,0,0],[0,200,200,200],[]] [[100,0,0,0],[0,200,0,0],[0,0,0,100]]		

Table 5 RL and LLM's presciption for PATNO 3507 during the whole process		
RL	LLM	
[[0,100],[],[195,0]] [[0,100],[],[195,0]] [[],[0,200],[95,0]] [[],[0,200],[95,0]] [[0,100],[200,0],[]] [[0,100],[200,0],[]] [[0,100],[200,0],[]] [[0,100],[200,0],[]] [[0,100],[200,0],[]] [[0,100],[200,0],[]]	[[],[100,100,100],[]] [[],[],[0,0,195]] [[100,0,0],[0,200,0],[0,95,95]] [[100,0,0],[0,200,0],[0,0,95]] [[250,250,0,0],[0,0,200,200],[]] [[100,100,0,0],[0,0,200,0],[0,0,0,95]] [[200,0,0],[0,100,100],[]] [[100,100,0,0],[],[0,0,195,195]] [[100,0,0],[],[0,0,95]] [[100,0,0],[0,200,0],[0,0,95]]	

Here we also briefly summarize the key information from these inputs and responses. For patient 3500, since the matched similar cases during the initial treatment phase recommended a combination of IR and Rytary, LLM also adopted this medication strategy. However, to achieve faster therapeutic effects in the early stage, the dosage of IR in the morning was increased, and to maintain sustained disease control, the dosage and frequency of Rytary were appropriately increased. In the middle stage, the first two recommended medication plans from the first similar case were integrated, and the medication frequency was balanced, with a low-dose Rytary supplement added in the evening. In the late stage, the frequency of the recommended medication plans from two similar cases was balanced, and a low-dose Rytary supplement was added in the evening.

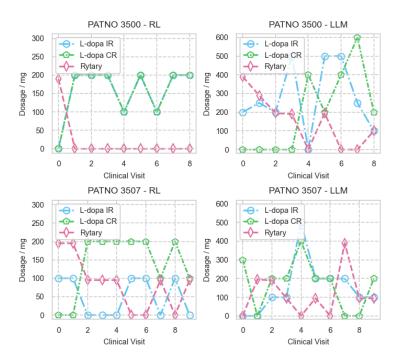


Figure 2: Trends of prescription over time for two representative patients under different experimental settings.

For patient 3507, the medication recommendations provided by LLM in the early stage were entirely based on the first recommended medication plan from the first similar case, taking into account that the patient's condition was less severe compared to the similar case, thus reducing the medication frequency and omitting Rytary. In the middle and late stages, to stabilize the disease, the medication strategy was not directly copied from the similar cases but instead adopted a dispersed low-dose medication strategy. In summary, this comprehensive and flexible medication selection strategy significantly enhanced the reliability and effectiveness of our framework.

#### E.1 PATNO 3500 - Early Term (2nd Clinical Visit)

#### E.1.1 Relative Cases

```
1
 ## Relative cases
 Several similar patient information given below, we adopt ML methods to predict the
 recommended treatment plans, and obtain explanations and suggestions output by
 LLMs.
 ### Similar Case 1
 Gender: female | Age: 68.1 | Years_sincePD: N/A
 Medication history (from old to new): [[100, 100, 100], [], []] / [[200, 200], [],
 []] / [[200, 200], [], []] / [[], [100, 100, 0, 0], [0, 0, 95, 95]] / [[], [100,
 100, 0, 0], [0, 0, 195, 195]]
 Nonlevodopa dose (from old to new): 0.0, 0.0, 8.0, 9.0, 9.0
 Six previous cycles of UPDRS scores (from old to new): 17.0, 22.0, 21.0, 23.0, 20.0,
10
 22.0
 Recommended prescriptions generated by ML method: Plan 1 - [[0, 250], [], [195, 0]]
 (-6.181) / Plan 2 - [[], [0, 0, 200, 200], [95, 95, 0, 0]] (-5.894) / Plan 3 -
 [[], [200, 0], [0, 195]] (-5.616) / Plan 4 - [[250, 250], [], []] (-5.02) /
```

```
Example prescriptions not recommended: Plan 1 - [[0, 100], [100, 0], []] (5.803) /
 → Plan 2 - [[250, 250, 250, 250], [], []] (3.778) / Plan 3 - [[0, 100], [], [195,
 _{\hookrightarrow} 0]] (2.888) / Plan 4 - [[], [], [95]] (2.692) /
 ### Similar Case 2
 Gender: female | Age: 68.4 | Years_sincePD: N/A
 Medication history (from old to new): None / None / [[], [200], []] / [[250, 250, 0, ^{\circ}
 → 0], [], [0, 0, 195, 195]] / [[0, 250], [200, 0], []]
 Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0
 Six previous cycles of UPDRS scores (from old to new): 14.0, 17.0, 15.0, 17.0, 19.0,
 → 19.0
 Recommended prescriptions generated by ML method: Plan 1 - [[0, 250], [], [95, 0]]
 → (-6.315) / Plan 2 - [[100, 0], [0, 200], []] (-5.136) / Plan 3 - [[], [],
 \rightarrow [95]] (-4.863) / Plan 4 - [[250, 250, 100, 100], [], []] (-4.646) /
 Example prescriptions not recommended: Plan 1 - [[], [200, 200, 0, 0], [0, 0, 95,
 → 95]] (3.307) / Plan 2 - [[], [], [195, 195]] (2.266) / Plan 3 - [[100, 100, 0,
 → 0], [0, 0, 200, 200], []] (2.243) / Plan 4 - [[0, 0, 250, 250], [], [195, 195,
 → 0, 0]] (2.121) /
E.1.2 Answer from LLM
 (a) Important Information About the Patient:
 - The patient is a 68.9-year-old female with an unknown number of years since
```

- $_{\hookrightarrow}$  Parkinson's disease (PD) diagnosis (indicated as N/A).
- The patient has a history of stable medication use, primarily involving L-dopa IR → and Rytary, with no non-levodopa drugs.
- The UPDRS scores have shown a decreasing trend over the past six cycles,
  - $\ \hookrightarrow \$  indicating some improvement in symptoms.
- \*\*(b) Analysis of Similar Cases and Recommended Treatment Plans:\*\*
- \*\*Similar Case 1:\*\* A 68.1-year-old female with similar medication history and
  - $\,\,\,\,\,\,\,\,\,\,\,\,\,\,$  stable UPDRS scores. The recommended plans include a mix of L-dopa IR and
  - -- Rytary, with a focus on higher doses in the morning and evening.
- \*\*Similar Case 2:\*\* A 68.4-year-old female with a similar medication history and
  - $\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,$  slightly lower UPDRS scores. The recommended plans also involve a mix of L-dopa
    - $\rightarrow$  IR and Rytary, with a preference for higher morning doses.
- \*\*(c) Final Prescription Strategy:\*\*
- Based on the similar cases, the final prescription strategy will focus on higher → morning doses of L-dopa IR and Rytary to manage symptoms effectively throughout
  - $\hookrightarrow$  the day.
- The strategy is modified from Similar Case 1's Plan 1, which showed a significant
- \*\*(d) Adjustments and Medication Strategies:\*\*
- The frequency will be set to 3 doses (Morning, Noon, Evening).
- The morning dose will be increased to 250 mg of L-dopa IR to provide a strong  $\hookrightarrow$  initial response.
- The noon dose will be adjusted to 195 mg of Rytary to maintain symptom control  $\hookrightarrow$  during the day.
- The evening dose will be adjusted to 95~mg of Rytary to ensure continued symptom  $\rightarrow$  management through the night.
- ### Formatted Treatment Plan 20

```python

```
23 [[250, 0, 0], [0, 0, 0], [0, 195, 95]]
24
```

E.2 PATNO 3500 - Medium Term (6th Clinical Visit)

E.2.1 Relative Cases

```
## Relative cases
Several similar patient information given below, we adopt ML methods to predict the
 -- recommended treatment plans, and obtain explanations and suggestions output by
    LLMs.
### Similar Case 1
Gender: female | Age: 73.9 | Years_sincePD: 5.3
Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
    [], []] / [[250, 250, 0, 0], [0, 0, 200, 200], []] / [[], [], [95, 95]] / [[250,
    250, 0, 0], [0, 0, 100, 100], []]
Nonlevodopa dose (from old to new): 25.0, 25.0, 17.0, 17.0, 22.0
Six previous cycles of UPDRS scores (from old to new): 35.0, 44.0, 37.0, 52.0, 34.0,

→ 41.0

Recommended prescriptions generated by ML method: Plan 1 - [[0, 0, 100, 100], [100,
 → 100, 0, 0], []] (-7.277) / Plan 2 - [[250, 250, 0, 0], [0, 0, 200, 200], []]
 \leftarrow (-6.658) / Plan 3 - [[0, 250], [100, 0], []] (-6.384) / Plan 4 - [[], [0,

→ 200], [195, 0]] (-5.944) /
Example prescriptions not recommended: Plan 1 - [[], [100, 100, 100, 100], []]
 \rightarrow (4.212) / Plan 2 - [[], [0, 100], [95, 0]] (3.481) / Plan 3 - [[250, 250, 0,
 → 0], [0, 0, 100, 100], []] (3.451) / Plan 4 - [[], [200, 200], []] (2.629) /
### Similar Case 2
Gender: female | Age: 73.4 | Years_sincePD: 4.8
Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
    [], []] / [[100, 100, 100], [], []] / [[250, 250, 0, 0], [0, 0, 200, 200], []] /
    [[], [], [95, 95]]
Nonlevodopa dose (from old to new): 0.0, 25.0, 25.0, 17.0, 17.0
Six previous cycles of UPDRS scores (from old to new): 35.0, 44.0, 37.0, 52.0, 34.0,
Recommended prescriptions generated by ML method: Plan 1 - [[250, 250, 0, 0], [0, 0,
 \rightarrow 100, 100], []] (-5.764) / Plan 2 - [[100, 0], [0, 200], []] (-5.615) / Plan 3
 - [[], [100], []] (-4.717) / Plan 4 - [[0, 250], [], [95, 0]] (-4.7) /
Example prescriptions not recommended: Plan 1 - [[], [200], []] (4.395) / Plan 2 -
 → [[], [0, 100], [195, 0]] (3.496) / Plan 3 - [[], [], [95]] (3.088) / Plan 4 -
 \rightarrow [[250, 250, 0, 0], [], [0, 0, 95, 95]] (2.904) /
```

E.2.2 Answer from LLM

```
### Brief Analysis

**(a) Important Information About the Patient:**

- The patient is a 73.8-year-old female diagnosed with Parkinson's disease (PD) 2.2

- years ago.

- The medication history shows a mix of L-dopa IR, L-dopa CR, and Rytary usage, with

- varying dosages and frequencies.

- The Nonlevodopa dose has been minimal, suggesting reliance primarily on L-dopa

- medications.
```

```
**(b) Analysis of Similar Cases and Recommended Treatment Plans:**
    - **Similar Case 1:** A 73.9-year-old female with 5.3 years since PD diagnosis. The
    -- recommended plans include a mix of L-dopa IR and CR, with a focus on higher
    \rightarrow dosages and more frequent administration.
   - **Similar Case 2:** A 73.4-year-old female with 4.8 years since PD diagnosis. The
    -- recommended plans also involve higher dosages of L-dopa IR and CR, with some use

→ of Rytary.

   - Both similar cases show a trend towards higher dosages and more frequent
    _{\mathrel{\mathrel{\hookrightarrow}}} administration to manage more advanced PD symptoms.
   **(c) Final Prescription Strategy:**
   - Based on the patient's recent low UPDRS scores and the need to prevent future
    \hookrightarrow escalations, a balanced approach is warranted. The strategy will be to maintain
    → a moderate dosage of L-dopa IR and CR, ensuring steady symptom management
    → without overmedicating.
    - The final prescription will be modified from Similar Case 1's Plan 1, which
    \,\,\,\,\,\,\,\,\,\,\,\,\,\, includes a balanced use of L-dopa CR and IR. The dosage and frequency will be
       adjusted to align with the patient's current needs and prevent potential
       overmedication.
17
   **(d) Adjustments and Medication Strategies:**
   - Increase the dosage and frequency of L-dopa IR and CR to manage symptoms more
       effectively.
   - Introduce Rytary in the evening to provide extended relief.
    - The final prescription will include three doses of L-dopa IR in the morning, noon,
    \rightarrow and evening, with additional L-dopa CR in the morning and noon, and Rytary in
    \hookrightarrow the evening.
22
   - **Frequency:** 3 doses
    - **Medicine and Dosage:**
      - (1) Morning: 250mg L-dopa IR, 100mg L-dopa CR
      - (2) Noon: 250mg L-dopa IR, 100mg L-dopa CR
      - (3) Evening: 195mg Rytary
    - **Additional comment: ** This plan aims to provide consistent and effective symptom
    \,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\, management throughout the day, with higher dosages in the morning and noon to

→ address early symptoms and Rytary in the evening for extended relief.

30
    ```python
31
 [[250, 250, 0], [100, 100, 0], [0, 0, 195]]
33
34
 PATNO 3500 - Late Term (9th Clinical Visit)
```

#### E.3.1 Relative Cases

```
Relative cases

Several similar patient information given below, we adopt ML methods to predict the
recommended treatment plans, and obtain explanations and suggestions output by
LLMs.
```

```
Similar Case 1
Gender: male | Age: 75.9 | Years_sincePD: 4.3
Medication history (from old to new): None / [[], [200, 200, 200, 200], []] /
 [[], [200, 200, 0, 0], [0, 0, 195, 195]] / [[], [100, 100], []]
Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0, 0.0
Six previous cycles of UPDRS scores (from old to new): 20.0, 18.0, 23.0, 22.0, 26.0,
Recommended prescriptions generated by ML method: Plan 1 - [[0, 0, 100, 100], [200,
 200, 0, 0], []] (-10.289) / Plan 2 - [[0, 250], [], [95, 0]] (-1.852) / Plan 3
 - [[], [0, 100], [95, 0]] (-1.608) / Plan 4 - [[100, 100, 100, 100], [], []]
 (-1.123) /
Example prescriptions not recommended: Plan 1 - [[], [0, 0, 200, 200], [195, 195, 0,
→ 0]] (10.921) / Plan 2 - [[], [100, 100, 0, 0], [0, 0, 95, 95]] (10.655) / Plan
→ 3 - [[0, 250], [200, 0], []] (10.46) / Plan 4 - [[250, 0], [], [0, 195]] (9.35)
Similar Case 2
Gender: female | Age: 73.4 | Years_sincePD: 4.8
Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
\rightarrow [], []] / [[100, 100, 100], [], []] / [[250, 250, 0, 0], [0, 0, 200, 200], []] /
Nonlevodopa dose (from old to new): 0.0, 25.0, 25.0, 17.0, 17.0
Six previous cycles of UPDRS scores (from old to new): 35.0, 44.0, 37.0, 52.0, 34.0,
\hookrightarrow 41.0
Recommended prescriptions generated by ML method: Plan 1 - [[250, 250, 0, 0], [0, 0,
→ 100, 100], []] (-5.764) / Plan 2 - [[100, 0], [0, 200], []] (-5.615) / Plan 3
_{\rightarrow} - [[], [100], []] (-4.717) / Plan 4 - [[0, 250], [], [95, 0]] (-4.7) /
Example prescriptions not recommended: Plan 1 - [[], [200], []] (4.395) / Plan 2 -
_{\rightarrow} [[], [0, 100], [195, 0]] (3.496) / Plan 3 - [[], [], [95]] (3.088) / Plan 4 -
\rightarrow [[250, 250, 0, 0], [], [0, 0, 95, 95]] (2.904) /
Brief Analysis
```

```
E.3.2 Answer from LLM
 (a) Important Information About the Patient:
 - The patient is a 75.8-year-old female diagnosed with Parkinson's disease (PD) 4.1

years ago.

 - The medication history shows a progression from lower to higher doses of L-dopa,
 \hookrightarrow with a recent trend towards more frequent dosing.
 - Nonlevodopa doses have been consistently low, suggesting minimal use of
 \ \ \, \rightarrow \ \ \, \text{non-levodopa medications.}
 (b) Analysis of Similar Cases and Recommended Treatment Plans:
 - **Similar Case 1 (male, 75.9 years, 4.3 years since PD): ** The recommended plans
 \hookrightarrow focus on higher doses of L-dopa CR and Rytary, with significant predicted UPDRS
 _{
ightarrow} score reductions. The best plan (Plan 1) suggests a combination of L-dopa CR and
 \rightarrow Rytary, with a predicted UPDRS score reduction of -10.289.
 - **Similar Case 2 (female, 73.4 years, 4.8 years since PD):** The recommended plans
 \,\,\,\,\,\,\,\,\, also emphasize higher doses of L-dopa IR and CR, with moderate predicted UPDRS
 \,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\, score reductions. The best plan (Plan 1) suggests a combination of L-dopa IR and
 _{\rm \hookrightarrow} CR, with a predicted UPDRS score reduction of -5.764.
```

\*\*(c) Final Prescription Strategy:\*\*

```
- Based on the analysis, the final prescription strategy will be modified from
 \hookrightarrow reduction.
 (d) Adjustments and Medication Strategies:
15
 - The frequency will be adjusted to 4 doses per day to better manage symptoms

→ throughout the day.

 - The dosage of L-dopa CR will be increased to match the recommended plan, and
 → Rytary will be introduced to provide extended relief.
 - This strategy aims to stabilize and potentially reduce the UPDRS score by ensuring
 consistent L-dopa levels throughout the day.
19
 ### Formatted Treatment Plan
20
21
 - **Frequency:** 4 doses
 - **Medicine and Dosage:**
23
 - (1) Morning: 100mg L-dopa IR
 - (2) Noon: 200mg L-dopa CR
 - (3) Afternoon: 100mg L-dopa IR
 - (4) Evening: 100mg Rytary
 - **Additional comment:** This plan is designed to provide balanced and extended

→ coverage of L-dopa throughout the day, aiming to manage symptoms effectively and

 \rightarrow potentially reduce the UPDRS score.
29
   ```python
   [[100, 0, 100, 0], [0, 200, 0, 0], [0, 0, 0, 100]]
```

E.4 PATNO 3507 - Early Term (1st Clinical Visit)

E.4.1 Relative Cases

```
## Relative cases
Several similar patient information given below, we adopt ML methods to predict the
-- recommended treatment plans, and obtain explanations and suggestions output by
    LLMs.
### Similar Case 1
Gender: male | Age: 73.2 | Years_sincePD: N/A
Medication history (from old to new): None / None / None / None / None
Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0, 0.0
Six previous cycles of UPDRS scores (from old to new): 7.0, 6.0, 12.0, 11.0, 7.0,
Recommended prescriptions generated by ML method: Plan 1 - [[], [100, 100, 100,
    100], []] (-2.204) / Plan 2 - [[], [100, 100, 200, 200], []] (-1.328) / Plan 3
   - [[250, 250, 0, 0], [], [0, 0, 195, 195]] (-1.306) / Plan 4 - [[], [0, 100],
Example prescriptions not recommended: Plan 1 - [[], [0, 0, 200, 200], [195, 195, 0,
    0]] (6.462) / Plan 2 - [[], [0, 200], [95, 0]] (5.546) / Plan 3 - [[], [0,
    100], [195, 0]] (5.421) / Plan 4 - [[0, 100], [200, 0], []] (5.074) /
### Similar Case 2
Gender: male | Age: 72.2 | Years_sincePD: N/A
Medication history (from old to new): None / None / None / None / None
```

```
Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0, 0.0
    Six previous cycles of UPDRS scores (from old to new): 23.0, 22.0, 34.0, 29.0, 43.0,

→ 31.0

   Recommended prescriptions generated by ML method: Plan 1 - [[0, 0, 100, 100], [200,
        200, 0, 0], []] (2.895) / Plan 2 - [[100], [], []] (3.667) / Plan 3 - [[100,
    → 100], [], []] (4.972) / Plan 4 - [[100, 100, 0, 0], [0, 0, 100, 100], []]
    Example prescriptions not recommended: Plan 1 - [[250, 250, 0, 0], [0, 0, 200, 200],
    → []] (14.811) / Plan 2 - [[], [], [195, 195, 195, 195]] (13.828) / Plan 3 -
    → [[0, 250], [200, 0], []] (13.698) / Plan 4 - [[0, 250], [100, 0], []] (13.682)
E.4.2 Answer from LLM
    ### Brief Analysis
   **(a) Important Information About the Patient:**
    - The patient is a 70.2-year-old male with no history of Parkinson's disease (PD)
    \rightarrow diagnosis (Years_sincePD: N/A).
    - The patient has no prior medication history for PD, and the Nonlevodopa dose has
    \hookrightarrow been consistently 0.0 mg.
    **(b) Analysis of Similar Cases and Recommended Treatment Plans:**
    - **Similar Case 1:** A 73.2-year-old male with similar non-diagnosis status and a
    → UPDRS trend from 7.0 to 16.0. The recommended plans include L-dopa CR with four
    \hookrightarrow doses of 100 mg each (Plan 1) and a combination of L-dopa IR and CR (Plan 2 and
    \,\,\,\,\,\,\,\,\,\, Plan 3). These plans are predicted to reduce the UPDRS score significantly.
    - **Similar Case 2:** A 72.2-year-old male with a more severe UPDRS progression from
    \hookrightarrow 23.0 to 31.0. The recommended plans are less effective, with predicted UPDRS
    \hookrightarrow scores showing minimal reduction or even slight increases.
    **(c) Final Prescription Strategy:**
    - Based on the analysis, the patient's condition resembles Similar Case 1 more
    \,\,\,\,\,\,\,\,\,\,\,\,\,\,\,\, closely, especially in terms of the UPDRS progression and non-diagnosis status.
    \,\,\,\,\,\,\,\,\,\, Therefore, the treatment plan from Similar Case 1 (Plan 1) is chosen as the
    \hookrightarrow basis for modification.
    **(d) Adjustments and Medication Strategies:**
    - The chosen plan from Similar Case 1 involves four doses of L-dopa CR, which is
    \hookrightarrow effective but may not be necessary for the current patient due to the less
        severe progression. Therefore, the frequency is reduced to three doses,
        maintaining the dosage to ensure efficacy.
    ### Formatted Treatment Plan
17
    - **Frequency:** 3 doses
    - **Medicine and Dosage:** (1) Morning: 100mg L-dopa CR (2) Noon: 100mg L-dopa CR
       (3) Evening: 100mg L-dopa CR
    - **Additional comment:** This plan balances the need for regular medication with
    _{\mathrel{\mathrel{\rightarrow}}} the patient's current condition, aiming to stabilize and potentially reduce the
    \hookrightarrow UPDRS score.
    ```python
 [[0, 0, 0], [100, 100, 100], [0, 0, 0]]
```

#### E.5 PATNO 3507 - Late Term (6th Clinical Visit)

#### E.5.1 Relative Cases

```
1
 ## Relative cases
 Several similar patient information given below, we adopt ML methods to predict the
 -- recommended treatment plans, and obtain explanations and suggestions output by
 LLMs.
 ### Similar Case 1
 Gender: male | Age: 70.7 | Years_sincePD: N/A
 Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
 [195, 195, 0, 0]] / [[0, 0, 250, 250], [], [195, 195, 0, 0]]
 Nonlevodopa dose (from old to new): 50.0, 50.0, 37.0, 37.0, 37.0
 Six previous cycles of UPDRS scores (from old to new): 25.0, 41.0, 38.0, 34.0, 35.0,
 Recommended prescriptions generated by ML method: Plan 1 - [[100, 0], [0, 200], []]
 \rightarrow (-3.732) / Plan 2 - [[], [100, 0], [0, 195]] (-3.73) / Plan 3 - [[], [100,
 100, 0, 0], [0, 0, 95, 95]] (-3.396) / Plan 4 - [[], [0, 200], [95, 0]]
 (-2.749) /
 Example prescriptions not recommended: Plan 1 - [[], [200], []] (9.451) / Plan 2 -
 [[], [100], []] (7.47) / Plan 3 - [[], [0, 100], [95, 0]] (7.309) / Plan 4 -
 [[], [], [95, 95]] (7.244) /
 ### Similar Case 2
 Gender: male | Age: 62.5 | Years_sincePD: N/A
 Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],

→ [], []] / [[0, 0, 100, 100], [100, 100, 0, 0], []] / [[], [100, 0], [0, 195]] /

 [[], [100, 100, 0, 0], [0, 0, 195, 195]]
 Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0
 Six previous cycles of UPDRS scores (from old to new): 20.0, 24.0, 15.0, 27.0, 20.0,
 25.0
 Recommended prescriptions generated by ML method: Plan 1 - [[100, 100, 0, 0], [],
 [0, 0, 195, 195]] (-0.082) / Plan 2 - [[0, 0, 250, 250], [100, 100, 0, 0], []]
 (0.766) / Plan 3 - [[0, 0, 250, 250], [200, 200, 0, 0], []] (0.915) / Plan 4 -
 \rightarrow [[250, 250, 0, 0], [], [0, 0, 95, 95]] (1.75) /
 Example prescriptions not recommended: Plan 1 - [[0, 100], [], [195, 0]] (12.019) /
 → Plan 2 - [[100, 100, 0, 0], [], [0, 0, 95, 95]] (11.517) / Plan 3 - [[], [0,
 200], [95, 0]] (10.767) / Plan 4 - [[0, 0, 100, 100], [], [195, 195, 0, 0]]
 (10.659) /
20
 The information of patient you need to analyze is:
^{21}
 Gender: male | Age: 72.7 | Years_sincePD: N/A
 Medication history (from old to new): [[200, 0, 0], [0, 0, 200], [0, 95, 0]] /
 [[200, 0, 0], [0, 0, 0], [0, 0, 195]] / [[100, 100, 0], [0, 0, 0], [0, 0, 95]] /
 [[150, 0, 0], [100, 100, 0, 0], [0, 0, 0, 195]] / [[100, 100, 0, 0], [0, 0, 0, 0]]
 → 0], [0, 0, 195, 195]]
 Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0, 0.0
 Six previous cycles of UPDRS scores (from old to new): 21.0, 0.03, 0.03, 0.03,

→ 48.68, 7.1

27
28
```

```
Relative cases
29
30
 Several similar patient information given below, we adopt ML methods to predict the
31
 -- recommended treatment plans, and obtain explanations and suggestions output by
 \hookrightarrow LLMs.
32
 ### Similar Case 1
 Gender: male | Age: 70.7 | Years_sincePD: N/A
 Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
 [], []] / [[250, 250, 0, 0], [0, 0, 100, 100], []] / [[], [0, 0, 200, 200],
 [195, 195, 0, 0]] / [[0, 0, 250, 250], [], [195, 195, 0, 0]]
 Nonlevodopa dose (from old to new): 50.0, 50.0, 37.0, 37.0, 37.0
 Six previous cycles of UPDRS scores (from old to new): 25.0, 41.0, 38.0, 34.0, 35.0,
 \hookrightarrow 35.0
 Recommended prescriptions generated by ML method: Plan 1 - [[100, 0], [0, 200], []]
 \rightarrow (-3.732) / Plan 2 - [[], [100, 0], [0, 195]] (-3.73) / Plan 3 - [[], [100,
 → 100, 0, 0], [0, 0, 95, 95]] (-3.396) / Plan 4 - [[], [0, 200], [95, 0]]
 Example prescriptions not recommended: Plan 1 - [[], [200], []] (9.451) / Plan 2 -
 [[], [100], []] (7.47) / Plan 3 - [[], [0, 100], [95, 0]] (7.309) / Plan 4 -
 → [[], [], [95, 95]] (7.244) /
 ### Similar Case 2
 Gender: male | Age: 62.5 | Years_sincePD: N/A
 Medication history (from old to new): [[100, 100, 100], [], []] / [[100, 100, 100],
 [], []] / [[0, 0, 100, 100], [100, 100, 0, 0], []] / [[], [100, 0], [0, 195]] /
 [[], [100, 100, 0, 0], [0, 0, 195, 195]]
 Nonlevodopa dose (from old to new): 0.0, 0.0, 0.0, 0.0
 Six previous cycles of UPDRS scores (from old to new): 20.0, 24.0, 15.0, 27.0, 20.0,
 \hookrightarrow 25.0
 Recommended prescriptions generated by ML method: Plan 1 - [[100, 100, 0, 0], [],
 [0, 0, 195, 195]] (-0.082) / Plan 2 - [[0, 0, 250, 250], [100, 100, 0, 0], []]
 (0.766) / Plan 3 - [[0, 0, 250, 250], [200, 200, 0, 0], []] (0.915) / Plan 4 -
 [[250, 250, 0, 0], [], [0, 0, 95, 95]] (1.75) /
 Example prescriptions not recommended: Plan 1 - [[0, 100], [], [195, 0]] (12.019) /
 → Plan 2 - [[100, 100, 0, 0], [], [0, 0, 95, 95]] (11.517) / Plan 3 - [[], [0,
 → 200], [95, 0]] (10.767) / Plan 4 - [[0, 0, 100, 100], [], [195, 195, 0, 0]]
 E.5.2 Answer from LLM
```

# ### Brief Analysis

\*\*Important Information about the Patient:\*\*

```
- The patient is a 72.7-year-old male with an unknown number of years since
 Parkinson's Disease (PD) diagnosis (indicated as N/A).
- The patient has been on various L-dopa regimens, including immediate-release (IR)
 and controlled-release (CR) forms, with no non-levodopa medications prescribed.
Analysis of Similar Cases:
- **Similar Case 1:** A 70.7-year-old male with a similar unknown PD duration. The
\hookrightarrow patient has a history of stable UPDRS scores around 35. The recommended plans
→ focus on a balanced use of L-dopa IR and CR, with a preference for lower doses

→ spread throughout the day.
```

```
- **Similar Case 2:** A 62.5-year-old male with a similar unknown PD duration. The
 \rightarrow patient has a history of fluctuating UPDRS scores, similar to the given patient.
 _{\mathrel{\mathrel{\rightarrow}}} The recommended plans suggest higher doses of L-dopa IR and CR, with a focus on
 → maintaining steady levels throughout the day.
10
 Final Prescription Strategy:
 - Based on the patient's rapid progression and the need to stabilize symptoms, the
 → strategy will be to use a combination of L-dopa IR and CR to maintain steady
 \hookrightarrow dopamine levels.
 - The prescription will be modified based on Similar Case 1's recommended plan,
 \hookrightarrow which suggests a balanced approach with lower doses spread throughout the day.
14
 Adjustments and Medication Strategies:
15
 - **Frequency: ** 4 doses (Morning, Noon, Afternoon, Evening) to ensure steady
 \hookrightarrow dopamine levels.
 - **Medicine and Dosage:**
 - Morning: 100mg L-dopa IR
 - Noon: 100mg L-dopa IR
 - Afternoon: 200mg L-dopa CR
 - Evening: 95mg Rytary
21
 - **Additional Comment: ** This regimen aims to prevent peak-dose dyskinesia and
 ### Formatted Treatment Plan
24
   ```python
   [[100, 100, 0, 0], [0, 0, 200, 0], [0, 0, 0, 95]]
```